



PMI RESEARCH & DEVELOPMENT

Study ZRHR-REXC-03-EU Clinical Study Report Appendix 15.2 Tables

Study Title:	A randomized, controlled, open-label, 3-arm parallel group, single-center study to demonstrate reductions in exposure to selected smoke constituents in smoking, healthy subjects switching to the Tobacco Heating System 2.2 (THS 2.2) or smoking abstinence, compared to continuing to use conventional cigarettes, for 5 days in confinement
Study Number:	ZRHR-REXC-03-EU
Product Name:	Tobacco Heating System 2.2 (THS 2.2)
Study Initiated (first subject screened):	29 June 2013
Study Completed (last subject last visit):	26 September 2013
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Version:	2.0
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This study was conducted in accordance with Good Clinical Practice.

Confidentiality Statement

This document is confidential. Disclosure of any of its contents to third parties is not permitted except by the prior written consent of Philip Morris Products S.A.



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15.2 SUMMARY TABLES



15.2.1 Disposition and Background Data Summary Tables

**Table 15.2.1.1 Summary of Subject Disposition - All Screened Subjects**

Status	THS 2.2 (N=80)		CC (N=41)		SA (N=39)		Overall Screened	
Total screened - n							329	
Screen failures - n (%)							160	(48.6%)
Screening failures, without product test - n (%)							160	(48.6%)
Screening failures, with product test - n (%)							0	
Enrolled - n (%)							169	(51.4%)
Enrolled, not randomized - n (%)							9	(2.7%)
Randomized - n (%)	80	(100 %)	41	(100 %)	39	(100 %)	160	(48.6%)
Total randomized - n	80		41		39		160	
Completed - n (%)	79	(98.8%)	41	(100 %)	39	(100 %)	159	(99.4%)
Discontinued - n (%)	1	(1.3%)	0		0		1	(0.6%)
Discontinued, with randomized product use	1	(1.3%)					1	(0.6%)

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled subjects are those who are eligible on Day -2; Discontinued refers to randomized subjects who discontinued the study before the planned discharge at Day 6; Completed refers to randomized subjects who did not discontinue the study before the planned discharge at Day 6.

Note: Percentages are based on the number of subjects indicated in the column header (N), apart from the Overall column where percentages of completed and discontinued refer to the total of subjects randomized.

Appendix 15.3.1.7

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_sdisp.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.1.2 Summary of Reasons for Discontinuations - Randomized Population**

	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Overall Randomized (N=160)
Total no. of discontinuations - n (%)	1 (1.3%)	0	0	1 (0.6%)
Reason for discontinuation				
Adverse events - n (%)	0			0
Protocol violation - n (%)	0			0
Withdrawal by subject - n (%)	1 (1.3%)			1 (0.6%)
Lost to follow-up - n (%)	0			0
Other - n (%)	0			0
Total no. of incomplete safety follow-up - n (%)	1 (1.3%)			1 (0.6%)
Reason for discontinuation				
Adverse events - n (%)	0			0
Protocol violation - n (%)	0			0
Withdrawal by subject - n (%)	1 (1.3%)			1 (0.6%)
Lost to follow-up - n (%)	0			0
Other - n (%)	0			0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.1.7

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.1.3 Summary of Protocol Deviations - Safety Population**

Significance/ Classification of Deviation	THS 2.2 (N=80)		CC (N=41)		SA (N=39)		Exposed Not Randomized (N=9)		Overall Safety (N=169)	
	n	(%)	n	(%)	n	(%)	n	(%)	n	(%)
Major	1	(1.3%)	0		0		0		1	(0.6%)
Protocol violation	1	(1.3%)							1	(0.6%)
Minor	32	(40.0%)	18	(43.9%)	18	(46.2%)	8	(88.9%)	76	(45.0%)
Time schedule deviation	26	(32.5%)	14	(34.1%)	15	(38.5%)	1	(11.1%)	56	(33.1%)
Procedure violation	16	(20.0%)	9	(22.0%)	4	(10.3%)	8	(88.9%)	37	(21.9%)
Assessment missing	1	(1.3%)	0		4	(10.3%)	0		5	(3.0%)

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System

Note: Exposed Not Randomized refers to all subjects exposed to THS 2.2 but not randomized. Overall Safety refers to all subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.1.10

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_pdev.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.1.4.1 Summary of Demographics and Other Baseline Characteristics - Safety Population**

Variable	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Sex						
Male	n (%)	39 (48.8%)	21 (51.2%)	20 (51.3%)	5 (55.6%)	85 (50.3%)
Female	n (%)	41 (51.3%)	20 (48.8%)	19 (48.7%)	4 (44.4%)	84 (49.7%)
Daily CC consumption at screening						
10-19 cig/day	n (%)	41 (51.3%)	21 (51.2%)	19 (48.7%)	3 (33.3%)	84 (49.7%)
>19 cig/day	n (%)	39 (48.8%)	20 (48.8%)	20 (51.3%)	6 (66.7%)	85 (50.3%)
Age (years)	n	80	41	39	9	169
	Mean (SD)	35.4 (9.40)	32.6 (10.06)	33.6 (11.51)	43.9 (13.60)	34.8 (10.51)
	Median	32.5	29.0	29.0	49.0	32.0
	Min, Max	22, 59	21, 59	21, 60	21, 58	21, 60
BMI (kg/m ²)	n	80	41	39	9	169
	Mean (SD)	24.46 (3.034)	25.80 (3.228)	24.81 (2.505)	26.18 (3.896)	24.96 (3.055)
	Median	24.35	26.00	24.60	25.80	24.70
	Min, Max	18.9, 31.6	18.8, 31.8	20.3, 31.6	21.4, 31.6	18.8, 31.8

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to all subjects exposed to THS 2.2.

Note: BMI = Body Mass Index.

Note: FTND = Fagerstrom Test for Nicotine Dependence.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.1.2, 15.3.1.6, 15.3.1.9, 15.3.6.15

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_demog.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.1.4.1 Summary of Demographics and Other Baseline Characteristics - Safety Population**

Variable	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
BMI classifications						
Underweight	n (%)	0	0	0	0	0
Normal weight	n (%)	46 (57.5%)	17 (41.5%)	22 (56.4%)	4 (44.4%)	89 (52.7%)
Overweight	n (%)	29 (36.3%)	19 (46.3%)	16 (41.0%)	3 (33.3%)	67 (39.6%)
Obese	n (%)	5 (6.3%)	5 (12.2%)	1 (2.6%)	2 (22.2%)	13 (7.7%)
Height (m)						
	n	80	41	39	9	169
	Mean (SD)	1.722 (0.0918)	1.710 (0.0926)	1.707 (0.1060)	1.678 (0.0828)	1.713 (0.0948)
	Median	1.710	1.700	1.720	1.660	1.710
	Min, Max	1.57, 2.00	1.52, 1.87	1.51, 1.88	1.56, 1.81	1.51, 2.00
Weight (kg)						
	n	80	41	39	9	169
	Mean (SD)	72.72 (11.708)	76.05 (14.579)	72.52 (11.614)	74.24 (15.708)	73.56 (12.627)
	Median	70.65	75.40	72.90	73.90	72.90
	Min, Max	49.7, 103.4	48.2, 106.0	52.3, 99.0	52.0, 102.8	48.2, 106.0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to all subjects exposed to THS 2.2.

Note: BMI = Body Mass Index.

Note: FTND = Fagerstrom Test for Nicotine Dependence.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.1.2, 15.3.1.6, 15.3.1.9, 15.3.6.15

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**Table 15.2.1.4.1 Summary of Demographics and Other Baseline Characteristics - Safety Population**

Variable	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Race						
White	n (%)	80 (100 %)	41 (100 %)	39 (100 %)	9 (100 %)	169 (100 %)
Black or African American	n (%)	0	0	0	0	0
American Indian or Alaska Native	n (%)	0	0	0	0	0
Asian	n (%)	0	0	0	0	0
Native Hawaiian or Other Pacific Islander	n (%)	0	0	0	0	0
Other	n (%)	0	0	0	0	0
ISO nicotine level						
<= 0.6 mg	n (%)	48 (60.0%)	32 (78.0%)	33 (84.6%)	4 (44.4%)	117 (69.2%)
> 0.6 - 1.0 mg	n (%)	32 (40.0%)	9 (22.0%)	6 (15.4%)	5 (55.6%)	52 (30.8%)

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to all subjects exposed to THS 2.2.

Note: BMI = Body Mass Index.

Note: FTND = Fagerstrom Test for Nicotine Dependence.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.1.2, 15.3.1.6, 15.3.1.9, 15.3.6.15

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**Table 15.2.1.4.1 Summary of Demographics and Other Baseline Characteristics - Safety Population**

Variable	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
CC nicotine yield at admission (mg)	n	80	41	39	9	169
	Mean (SD)	0.63 (0.134)	0.59 (0.121)	0.57 (0.111)	0.64 (0.151)	0.60 (0.129)
	Median	0.60	0.60	0.60	0.70	0.60
	Min, Max	0.1, 0.8	0.3, 0.8	0.3, 0.8	0.4, 0.8	0.1, 0.8
CC ISO tar yield at admission (mg)	n	80	41	39	9	169
	Mean (SD)	8.0 (1.81)	7.3 (1.70)	7.3 (1.46)	8.0 (2.24)	7.7 (1.75)
	Median	8.0	7.0	7.0	9.0	7.0
	Min, Max	1, 10	3, 10	4, 10	4, 10	1, 10
CC ISO tar yield at admission						
1-5 mg	n (%)	6 (7.5%)	7 (17.1%)	4 (10.3%)	2 (22.2%)	19 (11.2%)
6-8 mg	n (%)	43 (53.8%)	26 (63.4%)	29 (74.4%)	2 (22.2%)	100 (59.2%)
9-10 mg	n (%)	31 (38.8%)	8 (19.5%)	6 (15.4%)	5 (55.6%)	50 (29.6%)

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to all subjects exposed to THS 2.2.

Note: BMI = Body Mass Index.

Note: FTND = Fagerstrom Test for Nicotine Dependence.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.1.2, 15.3.1.6, 15.3.1.9, 15.3.6.15

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**Table 15.2.1.4.1 Summary of Demographics and Other Baseline Characteristics - Safety Population**

Variable	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
CYP2A6 activity (%)	n	80	41	39	1	161
	Mean (SD)	41.556 (15.3968)	44.313 (15.2036)	40.874 (15.7486)	NC	41.987 (15.4084)
	Median	40.501	40.794	38.310	NC	40.315
	Min, Max	6.68, 73.00	16.00, 73.57	14.65, 79.21	NC	6.68, 79.21
FTND total score classifications						
Missing	n (%)	4 (5.0%)	1 (2.4%)	1 (2.6%)	0	6 (3.6%)
Mild	n (%)	20 (25.0%)	7 (17.1%)	4 (10.3%)	2 (22.2%)	33 (19.5%)
Moderate	n (%)	36 (45.0%)	23 (56.1%)	22 (56.4%)	3 (33.3%)	84 (49.7%)
Severe	n (%)	20 (25.0%)	10 (24.4%)	12 (30.8%)	4 (44.4%)	46 (27.2%)
FTND total score	n	76	40	38	9	163
	Mean (SD)	5.0 (2.02)	5.1 (1.83)	5.5 (1.93)	5.7 (2.70)	5.2 (1.99)
	Median	5.0	5.0	5.0	6.0	5.0
	Min, Max	1, 9	1, 9	1, 9	2, 10	1, 10

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to all subjects exposed to THS 2.2.

Note: BMI = Body Mass Index.

Note: FTND = Fagerstrom Test for Nicotine Dependence.

Note: NC = Not calculated.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.1.2, 15.3.1.6, 15.3.1.9, 15.3.6.15

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_demog.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.1.4.2 Summary of Demographics and Other Baseline Characteristics - FAS Population**

Variable	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Overall (N=160)
Sex					
Male	n (%)	39 (48.8%)	21 (51.2%)	20 (51.3%)	80 (50.0%)
Female	n (%)	41 (51.3%)	20 (48.8%)	19 (48.7%)	80 (50.0%)
Daily CC consumption at screening					
10-19 cig/day	n (%)	41 (51.3%)	21 (51.2%)	19 (48.7%)	81 (50.6%)
>19 cig/day	n (%)	39 (48.8%)	20 (48.8%)	20 (51.3%)	79 (49.4%)
Age (years)	n	80	41	39	160
	Mean (SD)	35.4 (9.40)	32.6 (10.06)	33.6 (11.51)	34.3 (10.12)
	Median	32.5	29.0	29.0	31.5
	Min, Max	22, 59	21, 59	21, 60	21, 60
BMI (kg/m ²)	n	80	41	39	160
	Mean (SD)	24.46 (3.034)	25.80 (3.228)	24.81 (2.505)	24.89 (3.001)
	Median	24.35	26.00	24.60	24.70
	Min, Max	18.9, 31.6	18.8, 31.8	20.3, 31.6	18.8, 31.8

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: BMI = Body Mass Index.

Note: FTND = Fagerstrom Test for Nicotine Dependence.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.1.2, 15.3.1.6, 15.3.1.9, 15.3.6.15

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_demog2.sas

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**Table 15.2.1.4.2 Summary of Demographics and Other Baseline Characteristics - FAS Population**

Variable	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Overall (N=160)
BMI classifications					
Underweight	n (%)	0	0	0	0
Normal weight	n (%)	46 (57.5%)	17 (41.5%)	22 (56.4%)	85 (53.1%)
Overweight	n (%)	29 (36.3%)	19 (46.3%)	16 (41.0%)	64 (40.0%)
Obese	n (%)	5 (6.3%)	5 (12.2%)	1 (2.6%)	11 (6.9%)
Height (m)	n	80	41	39	160
	Mean (SD)	1.722 (0.0918)	1.710 (0.0926)	1.707 (0.1060)	1.715 (0.0953)
	Median	1.710	1.700	1.720	1.710
	Min, Max	1.57, 2.00	1.52, 1.87	1.51, 1.88	1.51, 2.00
Weight (kg)	n	80	41	39	160
	Mean (SD)	72.72 (11.708)	76.05 (14.579)	72.52 (11.614)	73.52 (12.491)
	Median	70.65	75.40	72.90	72.90
	Min, Max	49.7, 103.4	48.2, 106.0	52.3, 99.0	48.2, 106.0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: BMI = Body Mass Index.

Note: FTND = Fagerstrom Test for Nicotine Dependence.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.1.2, 15.3.1.6, 15.3.1.9, 15.3.6.15

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_demog2.sas

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**Table 15.2.1.4.2 Summary of Demographics and Other Baseline Characteristics - FAS Population**

Variable	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Overall (N=160)
Race					
White	n (%)	80 (100 %)	41 (100 %)	39 (100 %)	160 (100 %)
Black or African American	n (%)	0	0	0	0
American Indian or Alaska Native	n (%)	0	0	0	0
Asian	n (%)	0	0	0	0
Native Hawaiian or Other Pacific Islander	n (%)	0	0	0	0
Other	n (%)	0	0	0	0
ISO nicotine level					
<= 0.6 mg	n (%)	48 (60.0%)	32 (78.0%)	33 (84.6%)	113 (70.6%)
> 0.6 - 1.0 mg	n (%)	32 (40.0%)	9 (22.0%)	6 (15.4%)	47 (29.4%)

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: BMI = Body Mass Index.

Note: FTND = Fagerstrom Test for Nicotine Dependence.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.1.2, 15.3.1.6, 15.3.1.9, 15.3.6.15

Path: /cvm/projects/prj/development/000000106324/dev/tables/t_demog2.sas

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**Table 15.2.1.4.2 Summary of Demographics and Other Baseline Characteristics - FAS Population**

Variable	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Overall (N=160)
CC nicotine yield at admission (mg)	n	80	41	39	160
	Mean (SD)	0.63 (0.134)	0.59 (0.121)	0.57 (0.111)	0.60 (0.127)
	Median	0.60	0.60	0.60	0.60
	Min, Max	0.1, 0.8	0.3, 0.8	0.3, 0.8	0.1, 0.8
CC ISO tar yield at admission (mg)	n	80	41	39	160
	Mean (SD)	8.0 (1.81)	7.3 (1.70)	7.3 (1.46)	7.6 (1.72)
	Median	8.0	7.0	7.0	7.0
	Min, Max	1, 10	3, 10	4, 10	1, 10
CC ISO tar yield at admission					
1-5 mg	n (%)	6 (7.5%)	7 (17.1%)	4 (10.3%)	17 (10.6%)
6-8 mg	n (%)	43 (53.8%)	26 (63.4%)	29 (74.4%)	98 (61.3%)
9-10 mg	n (%)	31 (38.8%)	8 (19.5%)	6 (15.4%)	45 (28.1%)

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: BMI = Body Mass Index.

Note: FTND = Fagerstrom Test for Nicotine Dependence.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.1.2, 15.3.1.6, 15.3.1.9, 15.3.6.15

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_demog2.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.1.4.2 Summary of Demographics and Other Baseline Characteristics - FAS Population**

Variable	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Overall (N=160)
CYP2A6 activity (%)	n	80	41	39	160
	Mean (SD)	41.556 (15.3968)	44.313 (15.2036)	40.874 (15.7486)	42.096 (15.3945)
	Median	40.501	40.794	38.310	40.399
	Min, Max	6.68, 73.00	16.00, 73.57	14.65, 79.21	6.68, 79.21
FTND total score classifications					
Missing	n (%)	4 (5.0%)	1 (2.4%)	1 (2.6%)	6 (3.8%)
Mild	n (%)	20 (25.0%)	7 (17.1%)	4 (10.3%)	31 (19.4%)
Moderate	n (%)	36 (45.0%)	23 (56.1%)	22 (56.4%)	81 (50.6%)
Severe	n (%)	20 (25.0%)	10 (24.4%)	12 (30.8%)	42 (26.3%)
FTND total score					
	n	76	40	38	154
	Mean (SD)	5.0 (2.02)	5.1 (1.83)	5.5 (1.93)	5.2 (1.95)
	Median	5.0	5.0	5.0	5.0
	Min, Max	1, 9	1, 9	1, 9	1, 9

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: BMI = Body Mass Index.

Note: FTND = Fagerstrom Test for Nicotine Dependence.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.1.2, 15.3.1.6, 15.3.1.9, 15.3.6.15

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.1.4.2.1 Summary of Demographics and Other Baseline Characteristics by Sex - FAS Population**

Variable	Statistic	THS 2.2		CC		SA		Overall	
		Males (N=39)	Females (N=41)	Males (N=21)	Females (N=20)	Males (N=20)	Females (N=19)	Males (N=80)	Females (N=80)
Daily CC consumption at screening									
10-19 cig/day	n (%)	21 (53.8%)	20 (48.8%)	11 (52.4%)	10 (50.0%)	10 (50.0%)	9 (47.4%)	42 (52.5%)	39 (48.8%)
>19 cig/day	n (%)	18 (46.2%)	21 (51.2%)	10 (47.6%)	10 (50.0%)	10 (50.0%)	10 (52.6%)	38 (47.5%)	41 (51.3%)
Age (years)	n	39	41	21	20	20	19	80	80
	Mean (SD)	33.6 (8.11)	37.1 (10.29)	28.9 (5.72)	36.6 (12.09)	30.5 (9.10)	36.9 (13.02)	31.6 (8.01)	37.0 (11.29)
	Median	32.0	35.0	29.0	35.5	26.5	36.0	29.0	35.5
	Min, Max	22, 51	22, 59	21, 45	21, 59	21, 53	21, 60	21, 53	21, 60
BMI (kg/m ²)	n	39	41	21	20	20	19	80	80
	Mean (SD)	24.63 (3.190)	24.30 (2.909)	26.61 (2.542)	24.96 (3.695)	24.77 (2.430)	24.85 (2.649)	25.19 (2.948)	24.59 (3.043)
	Median	24.80	24.30	26.80	24.70	24.50	24.60	25.15	24.45
	Min, Max	19.1, 31.6	18.9, 31.4	21.6, 31.3	18.8, 31.8	20.3, 29.0	20.7, 31.6	19.1, 31.6	18.8, 31.8

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: BMI = Body Mass Index.

Note: FTND = Fagerstrom Test for Nicotine Dependence.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.1.2, 15.3.1.6, 15.3.1.9, 15.3.6.15

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**Table 15.2.1.4.2.1 Summary of Demographics and Other Baseline Characteristics by Sex - FAS Population**

Variable	Statistic	THS 2.2		CC		SA		Overall	
		Males (N=39)	Females (N=41)	Males (N=21)	Females (N=20)	Males (N=20)	Females (N=19)	Males (N=80)	Females (N=80)
BMI classifications									
Underweight	n (%)	0	0	0	0	0	0	0	0
Normal weight	n (%)	20 (51.3%)	26 (63.4%)	6 (28.6%)	11 (55.0%)	11 (55.0%)	11 (57.9%)	37 (46.3%)	48 (60.0%)
Overweight	n (%)	17 (43.6%)	12 (29.3%)	13 (61.9%)	6 (30.0%)	9 (45.0%)	7 (36.8%)	39 (48.8%)	25 (31.3%)
Obese	n (%)	2 (5.1%)	3 (7.3%)	2 (9.5%)	3 (15.0%)	0	1 (5.3%)	4 (5.0%)	7 (8.8%)
Height (m)	n	39	41	21	20	20	19	80	80
	Mean (SD)	1.792 (0.0673)	1.656 (0.0552)	1.780 (0.0614)	1.637 (0.0554)	1.796 (0.0435)	1.612 (0.0589)	1.790 (0.0602)	1.641 (0.0582)
	Median	1.774	1.640	1.780	1.635	1.805	1.610	1.780	1.630
	Min, Max	1.70, 2.00	1.57, 1.80	1.70, 1.87	1.52, 1.76	1.70, 1.88	1.51, 1.72	1.70, 2.00	1.51, 1.80
Weight (kg)	n	39	41	21	20	20	19	80	80
	Mean (SD)	79.07 (10.987)	66.67 (8.893)	84.50 (11.093)	67.18 (12.489)	79.93 (9.125)	64.72 (8.464)	80.71 (10.703)	66.33 (9.734)
	Median	77.00	67.00	84.00	62.95	77.45	63.60	78.55	65.75
	Min, Max	60.3, 103.4	49.7, 82.5	67.8, 106.0	48.2, 93.0	65.8, 99.0	52.3, 87.0	60.3, 106.0	48.2, 93.0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: BMI = Body Mass Index.

Note: FTND = Fagerstrom Test for Nicotine Dependence.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.1.2, 15.3.1.6, 15.3.1.9, 15.3.6.15

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**Table 15.2.1.4.2.1 Summary of Demographics and Other Baseline Characteristics by Sex - FAS Population**

Variable	Statistic	THS 2.2		CC		SA		Overall	
		Males (N=39)	Females (N=41)	Males (N=21)	Females (N=20)	Males (N=20)	Females (N=19)	Males (N=80)	Females (N=80)
Race									
White	n (%)	39 (100 %)	41 (100 %)	21 (100 %)	20 (100 %)	20 (100 %)	19 (100 %)	80 (100 %)	80 (100 %)
Black or African American	n (%)	0	0	0	0	0	0	0	0
American Indian or Alaska Native	n (%)	0	0	0	0	0	0	0	0
Asian	n (%)	0	0	0	0	0	0	0	0
Native Hawaiian or Other Pacific Islander	n (%)	0	0	0	0	0	0	0	0
Other	n (%)	0	0	0	0	0	0	0	0
ISO nicotine level									
<= 0.6 mg	n (%)	20 (51.3%)	28 (68.3%)	15 (71.4%)	17 (85.0%)	16 (80.0%)	17 (89.5%)	51 (63.8%)	62 (77.5%)
> 0.6 - 1.0 mg	n (%)	19 (48.7%)	13 (31.7%)	6 (28.6%)	3 (15.0%)	4 (20.0%)	2 (10.5%)	29 (36.3%)	18 (22.5%)

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: BMI = Body Mass Index.

Note: FTND = Fagerstrom Test for Nicotine Dependence.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.1.2, 15.3.1.6, 15.3.1.9, 15.3.6.15

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**Table 15.2.1.4.2.1 Summary of Demographics and Other Baseline Characteristics by Sex - FAS Population**

Variable	Statistic	THS 2.2		CC		SA		Overall	
		Males (N=39)	Females (N=41)	Males (N=21)	Females (N=20)	Males (N=20)	Females (N=19)	Males (N=80)	Females (N=80)
CC nicotine yield at admission (mg)	n	39	41	21	20	20	19	80	80
	Mean (SD)	0.65 (0.126)	0.60 (0.140)	0.62 (0.117)	0.56 (0.120)	0.57 (0.127)	0.56 (0.096)	0.62 (0.126)	0.58 (0.127)
	Median	0.60	0.60	0.60	0.50	0.50	0.60	0.60	0.60
	Min, Max	0.5, 0.8	0.1, 0.8	0.5, 0.8	0.3, 0.8	0.3, 0.8	0.4, 0.8	0.3, 0.8	0.1, 0.8
CC ISO tar yield at admission (mg)	n	39	41	21	20	20	19	80	80
	Mean (SD)	8.4 (1.45)	7.5 (2.01)	8.0 (1.33)	6.6 (1.74)	7.6 (1.51)	7.1 (1.40)	8.1 (1.46)	7.2 (1.83)
	Median	8.0	7.0	7.0	7.0	7.0	7.0	8.0	7.0
	Min, Max	5, 10	1, 10	7, 10	3, 10	4, 10	5, 10	4, 10	1, 10

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: BMI = Body Mass Index.

Note: FTND = Fagerstrom Test for Nicotine Dependence.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.1.2, 15.3.1.6, 15.3.1.9, 15.3.6.15

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**Table 15.2.1.4.2.1 Summary of Demographics and Other Baseline Characteristics by Sex - FAS Population**

Variable	Statistic	THS 2.2		CC		SA		Overall	
		Males (N=39)	Females (N=41)	Males (N=21)	Females (N=20)	Males (N=20)	Females (N=19)	Males (N=80)	Females (N=80)
CC ISO tar yield at admission									
1-5 mg	n (%)	1 (2.6%)	5 (12.2%)	0	7 (35.0%)	1 (5.0%)	3 (15.8%)	2 (2.5%)	15 (18.8%)
6-8 mg	n (%)	19 (48.7%)	24 (58.5%)	15 (71.4%)	11 (55.0%)	15 (75.0%)	14 (73.7%)	49 (61.3%)	49 (61.3%)
9-10 mg	n (%)	19 (48.7%)	12 (29.3%)	6 (28.6%)	2 (10.0%)	4 (20.0%)	2 (10.5%)	29 (36.3%)	16 (20.0%)
CYP2A6 activity (%)	n	39	41	21	20	20	19	80	80
	Mean (SD)	39.321 (16.1438)	43.682 (14.5292)	41.822 (14.6551)	46.929 (15.7001)	36.568 (12.4444)	45.408 (17.8276)	39.289 (14.8513)	44.904 (15.5067)
	Median	39.847	43.785	37.530	44.998	35.730	38.651	37.815	44.045
	Min, Max	6.68, 70.11	16.20, 73.00	23.45, 67.66	16.00, 73.57	18.67, 63.33	14.65, 79.21	6.68, 70.11	14.65, 79.21

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: BMI = Body Mass Index.

Note: FTND = Fagerstrom Test for Nicotine Dependence.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.1.2, 15.3.1.6, 15.3.1.9, 15.3.6.15

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**Table 15.2.1.4.2.1 Summary of Demographics and Other Baseline Characteristics by Sex - FAS Population**

Variable	Statistic	THS 2.2		CC		SA		Overall	
		Males (N=39)	Females (N=41)	Males (N=21)	Females (N=20)	Males (N=20)	Females (N=19)	Males (N=80)	Females (N=80)
FTND total score classifications									
Missing	n (%)	1 (2.6%)	3 (7.3%)	1 (4.8%)	0	0	1 (5.3%)	2 (2.5%)	4 (5.0%)
Mild	n (%)	12 (30.8%)	8 (19.5%)	3 (14.3%)	4 (20.0%)	4 (20.0%)	0	19 (23.8%)	12 (15.0%)
Moderate	n (%)	17 (43.6%)	19 (46.3%)	11 (52.4%)	12 (60.0%)	11 (55.0%)	11 (57.9%)	39 (48.8%)	42 (52.5%)
Severe	n (%)	9 (23.1%)	11 (26.8%)	6 (28.6%)	4 (20.0%)	5 (25.0%)	7 (36.8%)	20 (25.0%)	22 (27.5%)
FTND total score	n	38	38	20	20	20	18	78	76
	Mean (SD)	4.8 (2.09)	5.3 (1.95)	5.4 (1.61)	4.9 (2.04)	5.0 (2.13)	6.0 (1.58)	5.0 (1.98)	5.3 (1.91)
	Median	4.5	5.0	5.0	5.0	5.0	5.5	5.0	5.0
	Min, Max	1, 9	1, 9	3, 9	1, 8	1, 8	4, 9	1, 9	1, 9

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: BMI = Body Mass Index.

Note: FTND = Fagerstrom Test for Nicotine Dependence.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.1.2, 15.3.1.6, 15.3.1.9, 15.3.6.15

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**Table 15.2.1.4.2.2 Summary of Demographics and Other Baseline Characteristics by Cigarette Consumption - FAS Population**

Variable	Statistic	THS 2.2		CC		SA		Overall	
		10-19 cig/day (N=41)	>19 cig/day (N=39)	10-19 cig/day (N=21)	>19 cig/day (N=20)	10-19 cig/day (N=19)	>19 cig/day (N=20)	10-19 cig/day (N=81)	>19 cig/day (N=79)
Sex									
Male	n (%)	21 (51.2%)	18 (46.2%)	11 (52.4%)	10 (50.0%)	10 (52.6%)	10 (50.0%)	42 (51.9%)	38 (48.1%)
Female	n (%)	20 (48.8%)	21 (53.8%)	10 (47.6%)	10 (50.0%)	9 (47.4%)	10 (50.0%)	39 (48.1%)	41 (51.9%)
Age (years)	n	41	39	21	20	19	20	81	79
	Mean (SD)	33.0 (8.91)	37.9 (9.36)	32.2 (10.30)	33.1 (10.05)	31.2 (10.82)	35.9 (11.95)	32.4 (9.65)	36.2 (10.30)
	Median	32.0	37.0	29.0	29.5	27.0	36.5	30.0	36.0
	Min, Max	22, 59	22, 57	21, 57	21, 59	21, 53	21, 60	21, 59	21, 60
BMI (kg/m ²)	n	41	39	21	20	19	20	81	79
	Mean (SD)	24.48 (2.846)	24.44 (3.258)	25.66 (3.508)	25.96 (2.990)	24.67 (2.950)	24.94 (2.066)	24.83 (3.055)	24.95 (2.963)
	Median	24.30	24.50	25.60	26.20	24.40	24.90	24.60	24.90
	Min, Max	18.9, 30.2	19.1, 31.6	18.8, 31.3	20.9, 31.8	20.3, 31.6	21.1, 28.7	18.8, 31.6	19.1, 31.8

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: BMI = Body Mass Index.

Note: FTND = Fagerstrom Test for Nicotine Dependence.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.1.2, 15.3.1.6, 15.3.1.9, 15.3.6.15

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**Table 15.2.1.4.2.2 Summary of Demographics and Other Baseline Characteristics by Cigarette Consumption - FAS Population**

Variable	Statistic	THS 2.2		CC		SA		Overall	
		10-19 cig/day (N=41)	>19 cig/day (N=39)	10-19 cig/day (N=21)	>19 cig/day (N=20)	10-19 cig/day (N=19)	>19 cig/day (N=20)	10-19 cig/day (N=81)	>19 cig/day (N=79)
BMI classifications									
Underweight	n (%)	0	0	0	0	0	0	0	0
Normal weight	n (%)	24 (58.5%)	22 (56.4%)	9 (42.9%)	8 (40.0%)	11 (57.9%)	11 (55.0%)	44 (54.3%)	41 (51.9%)
Overweight	n (%)	15 (36.6%)	14 (35.9%)	9 (42.9%)	10 (50.0%)	7 (36.8%)	9 (45.0%)	31 (38.3%)	33 (41.8%)
Obese	n (%)	2 (4.9%)	3 (7.7%)	3 (14.3%)	2 (10.0%)	1 (5.3%)	0	6 (7.4%)	5 (6.3%)
Height (m)									
	n	41	39	21	20	19	20	81	79
	Mean (SD)	1.731 (0.1024)	1.713 (0.0793)	1.709 (0.0950)	1.711 (0.0924)	1.721 (0.1042)	1.693 (0.1085)	1.723 (0.1002)	1.707 (0.0899)
	Median	1.750	1.710	1.700	1.705	1.770	1.705	1.720	1.710
	Min, Max	1.57, 2.00	1.58, 1.85	1.52, 1.86	1.57, 1.87	1.56, 1.86	1.51, 1.88	1.52, 2.00	1.51, 1.88

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: BMI = Body Mass Index.

Note: FTND = Fagerstrom Test for Nicotine Dependence.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.1.2, 15.3.1.6, 15.3.1.9, 15.3.6.15

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**Table 15.2.1.4.2.2 Summary of Demographics and Other Baseline Characteristics by Cigarette Consumption - FAS Population**

Variable	Statistic	THS 2.2		CC		SA		Overall	
		10-19 cig/day (N=41)	>19 cig/day (N=39)	10-19 cig/day (N=21)	>19 cig/day (N=20)	10-19 cig/day (N=19)	>19 cig/day (N=20)	10-19 cig/day (N=81)	>19 cig/day (N=79)
Weight (kg)	n	41	39	21	20	19	20	81	79
	Mean (SD)	73.73 (12.813)	71.64 (10.481)	75.65 (15.735)	76.47 (13.656)	73.39 (12.772)	71.69 (10.665)	74.15 (13.472)	72.88 (11.449)
	Median	72.90	70.00	75.80	74.10	74.30	69.90	74.10	70.40
	Min, Max	49.7, 103.4	53.2, 98.1	48.2, 106.0	56.6, 99.8	52.3, 95.9	57.0, 99.0	48.2, 106.0	53.2, 99.8

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: BMI = Body Mass Index.

Note: FTND = Fagerstrom Test for Nicotine Dependence.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.1.2, 15.3.1.6, 15.3.1.9, 15.3.6.15

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**Table 15.2.1.4.2.2 Summary of Demographics and Other Baseline Characteristics by Cigarette Consumption - FAS Population**

Variable	Statistic	THS 2.2		CC		SA		Overall	
		10-19 cig/day (N=41)	>19 cig/day (N=39)	10-19 cig/day (N=21)	>19 cig/day (N=20)	10-19 cig/day (N=19)	>19 cig/day (N=20)	10-19 cig/day (N=81)	>19 cig/day (N=79)
Race									
White	n (%)	41 (100 %)	39 (100 %)	21 (100 %)	20 (100 %)	19 (100 %)	20 (100 %)	81 (100 %)	79 (100 %)
Black or African American	n (%)	0	0	0	0	0	0	0	0
American Indian or Alaska Native	n (%)	0	0	0	0	0	0	0	0
Asian	n (%)	0	0	0	0	0	0	0	0
Native Hawaiian or Other Pacific Islander	n (%)	0	0	0	0	0	0	0	0
Other	n (%)	0	0	0	0	0	0	0	0
ISO nicotine level									
<= 0.6 mg	n (%)	24 (58.5%)	24 (61.5%)	14 (66.7%)	18 (90.0%)	16 (84.2%)	17 (85.0%)	54 (66.7%)	59 (74.7%)
> 0.6 - 1.0 mg	n (%)	17 (41.5%)	15 (38.5%)	7 (33.3%)	2 (10.0%)	3 (15.8%)	3 (15.0%)	27 (33.3%)	20 (25.3%)

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: BMI = Body Mass Index.

Note: FTND = Fagerstrom Test for Nicotine Dependence.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.1.2, 15.3.1.6, 15.3.1.9, 15.3.6.15

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**Table 15.2.1.4.2.2 Summary of Demographics and Other Baseline Characteristics by Cigarette Consumption - FAS Population**

Variable	Statistic	THS 2.2		CC		SA		Overall	
		10-19 cig/day (N=41)	>19 cig/day (N=39)	10-19 cig/day (N=21)	>19 cig/day (N=20)	10-19 cig/day (N=19)	>19 cig/day (N=20)	10-19 cig/day (N=81)	>19 cig/day (N=79)
CC nicotine yield at admission (mg)	n	41	39	21	20	19	20	81	79
	Mean (SD)	0.61 (0.146)	0.64 (0.121)	0.63 (0.115)	0.55 (0.115)	0.56 (0.112)	0.57 (0.113)	0.61 (0.132)	0.60 (0.123)
	Median	0.60	0.60	0.60	0.50	0.50	0.60	0.60	0.60
	Min, Max	0.1, 0.8	0.4, 0.8	0.5, 0.8	0.3, 0.8	0.4, 0.8	0.3, 0.8	0.1, 0.8	0.3, 0.8
CC ISO tar yield at admission (mg)	n	41	39	21	20	19	20	81	79
	Mean (SD)	7.9 (1.97)	8.1 (1.64)	7.6 (1.81)	7.0 (1.56)	7.2 (1.54)	7.5 (1.40)	7.6 (1.84)	7.6 (1.61)
	Median	7.0	8.0	7.0	7.0	7.0	7.0	7.0	7.0
	Min, Max	1, 10	4, 10	5, 10	3, 10	5, 10	4, 10	1, 10	3, 10

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: BMI = Body Mass Index.

Note: FTND = Fagerstrom Test for Nicotine Dependence.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.1.2, 15.3.1.6, 15.3.1.9, 15.3.6.15

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**Table 15.2.1.4.2.2 Summary of Demographics and Other Baseline Characteristics by Cigarette Consumption - FAS Population**

Variable	Statistic	THS 2.2		CC		SA		Overall	
		10-19 cig/day (N=41)	>19 cig/day (N=39)	10-19 cig/day (N=21)	>19 cig/day (N=20)	10-19 cig/day (N=19)	>19 cig/day (N=20)	10-19 cig/day (N=81)	>19 cig/day (N=79)
CC ISO tar yield at admission									
1-5 mg	n (%)	3 (7.3%)	3 (7.7%)	4 (19.0%)	3 (15.0%)	3 (15.8%)	1 (5.0%)	10 (12.3%)	7 (8.9%)
6-8 mg	n (%)	22 (53.7%)	21 (53.8%)	11 (52.4%)	15 (75.0%)	13 (68.4%)	16 (80.0%)	46 (56.8%)	52 (65.8%)
9-10 mg	n (%)	16 (39.0%)	15 (38.5%)	6 (28.6%)	2 (10.0%)	3 (15.8%)	3 (15.0%)	25 (30.9%)	20 (25.3%)
CYP2A6 activity (%)	n	41	39	21	20	19	20	81	79
	Mean (SD)	38.209 (15.4190)	45.074 (14.7577)	43.456 (14.4744)	45.213 (16.2613)	42.894 (17.3352)	38.955 (14.2613)	40.668 (15.6582)	43.560 (15.0779)
	Median	36.884	44.537	39.248	44.998	38.651	37.521	38.737	43.061
	Min, Max	6.68, 70.11	17.38, 73.00	16.00, 73.57	23.45, 69.37	20.70, 79.21	14.65, 61.98	6.68, 79.21	14.65, 73.00

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: BMI = Body Mass Index.

Note: FTND = Fagerstrom Test for Nicotine Dependence.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.1.2, 15.3.1.6, 15.3.1.9, 15.3.6.15

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_demog2c.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.1.4.2.2 Summary of Demographics and Other Baseline Characteristics by Cigarette Consumption - FAS Population**

Variable	Statistic	THS 2.2		CC		SA		Overall	
		10-19 cig/day (N=41)	>19 cig/day (N=39)	10-19 cig/day (N=21)	>19 cig/day (N=20)	10-19 cig/day (N=19)	>19 cig/day (N=20)	10-19 cig/day (N=81)	>19 cig/day (N=79)
FTND total score classifications									
Missing	n (%)	2 (4.9%)	2 (5.1%)	0	1 (5.0%)	1 (5.3%)	0	3 (3.7%)	3 (3.8%)
Mild	n (%)	15 (36.6%)	5 (12.8%)	6 (28.6%)	1 (5.0%)	4 (21.1%)	0	25 (30.9%)	6 (7.6%)
Moderate	n (%)	19 (46.3%)	17 (43.6%)	11 (52.4%)	12 (60.0%)	10 (52.6%)	12 (60.0%)	40 (49.4%)	41 (51.9%)
Severe	n (%)	5 (12.2%)	15 (38.5%)	4 (19.0%)	6 (30.0%)	4 (21.1%)	8 (40.0%)	13 (16.0%)	29 (36.7%)
FTND total score									
	n	39	37	21	19	18	20	78	76
	Mean (SD)	4.1 (1.78)	6.0 (1.84)	4.5 (1.94)	5.8 (1.48)	4.7 (2.09)	6.2 (1.48)	4.4 (1.89)	6.0 (1.65)
	Median	4.0	6.0	5.0	6.0	4.5	6.0	4.0	6.0
	Min, Max	1, 7	2, 9	1, 8	3, 9	1, 8	4, 9	1, 8	2, 9

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: BMI = Body Mass Index.

Note: FTND = Fagerstrom Test for Nicotine Dependence.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.1.2, 15.3.1.6, 15.3.1.9, 15.3.6.15

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_demog2c.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.1.5 Summary of Current Cigarette Brands at Admission - Safety Population**

Brand	ISO Nicotine Yield (mg)	ISO Tar Yield (mg)	ISO CO Yield (mg)	THS 2.2 (N=80) n (%)	CC (N=41) n (%)	SA (N=39) n (%)	Enrolled Not Randomized (N=9) n (%)	Overall Safety (N=169) n (%)
Marlboro gold	0.5	7	9	11 (13.8%)	7 (17.1%)	7 (17.9%)	1 (11.1%)	26 (15.4%)
L&m blue	0.6	8	9	7 (8.8%)	2 (4.9%)	4 (10.3%)	1 (11.1%)	14 (8.3%)
L&m blue link	0.6	7	6	6 (7.5%)	1 (2.4%)	4 (10.3%)	0	11 (6.5%)
Ld pink	0.5	5	4	2 (2.5%)	5 (12.2%)	2 (5.1%)	0	9 (5.3%)
L&m blue light	0.6	8	9	4 (5.0%)	2 (4.9%)	0	0	6 (3.6%)
Marlboro red	0.7	9	9	5 (6.3%)	0	0	1 (11.1%)	6 (3.6%)
Chesterfield blue	0.6	7	7	2 (2.5%)	1 (2.4%)	1 (2.6%)	0	4 (2.4%)
L&m red	0.8	10	10	2 (2.5%)	0	2 (5.1%)	0	4 (2.4%)
Lucky strike	0.6	7	8	0	2 (4.9%)	2 (5.1%)	0	4 (2.4%)
Lucky strike red	0.8	10	10	2 (2.5%)	1 (2.4%)	0	1 (11.1%)	4 (2.4%)
Camel	0.8	10	10	3 (3.8%)	0	0	0	3 (1.8%)
Chesterfield red	0.8	10	10	2 (2.5%)	1 (2.4%)	0	0	3 (1.8%)
Ld blue	0.5	7	9	2 (2.5%)	0	1 (2.6%)	0	3 (1.8%)
Ld red	0.7	10	10	2 (2.5%)	0	1 (2.6%)	0	3 (1.8%)
Viceroy blue classic	0.5	7	8	2 (2.5%)	1 (2.4%)	0	0	3 (1.8%)

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.1.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_cigbrand.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.1.5 Summary of Current Cigarette Brands at Admission - Safety Population**

Brand	ISO Nicotine Yield (mg)	ISO Tar Yield (mg)	ISO CO Yield (mg)	THS 2.2 (N=80)		CC (N=41)		SA (N=39)		Enrolled Not Randomized (N=9)		Overall Safety (N=169)	
				n	(%)	n	(%)	n	(%)	n	(%)	n	(%)
Davidoff gold slim	0.6	7	6	1	(1.3%)	0		1	(2.6%)	0		2	(1.2%)
L&m red link	0.7	9	8	1	(1.3%)	0		0		1	(11.1%)	2	(1.2%)
Ld blue	0.3	3	2	1	(1.3%)	1	(2.4%)	0		0		2	(1.2%)
Lucky strike	0.8	10	10	2	(2.5%)	0		0		0		2	(1.2%)
Marlboro gold 100 prime edge	0.5	7	7	0		2	(4.9%)	0		0		2	(1.2%)
Red white	0.7	10	10	1	(1.3%)	1	(2.4%)	0		0		2	(1.2%)
Route 66	0.8	10	10	0		1	(2.4%)	1	(2.6%)	0		2	(1.2%)
She white	0.4	6	5	0		0		2	(5.1%)	0		2	(1.2%)
Vogue blue	0.7	7	5	1	(1.3%)	1	(2.4%)	0		0		2	(1.2%)
Winston red	0.8	10	10	1	(1.3%)	0		0		1	(11.1%)	2	(1.2%)
Austin	0.8	10	10	1	(1.3%)	0		0		0		1	(0.6%)
Black devil	0.8	10	10	1	(1.3%)	0		0		0		1	(0.6%)
Camel blue	0.6	8	9	0		1	(2.4%)	0		0		1	(0.6%)
Camel yellow	0.8	10	10	0		0		1	(2.6%)	0		1	(0.6%)
Chesterfield	0.8	10	10	1	(1.3%)	0		0		0		1	(0.6%)

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.1.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_cigbrand.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.1.5 Summary of Current Cigarette Brands at Admission - Safety Population**

Brand	ISO Nicotine Yield (mg)	ISO Tar Yield (mg)	ISO CO Yield (mg)	THS 2.2 (N=80)		CC (N=41)		SA (N=39)		Enrolled Not Randomized (N=9)		Overall Safety (N=169)	
				n	(%)	n	(%)	n	(%)	n	(%)	n	(%)
Chesterfield blue	0.5	7	9	0		0		1 (2.6%)		0		1 (0.6%)	
Chesterfield blue slims	0.6	7	7	0		1 (2.4%)		0		0		1 (0.6%)	
Golden fog red	0.8	10	10	0		1 (2.4%)		0		0		1 (0.6%)	
L&m	0.6	8	9	0		0		1 (2.6%)		0		1 (0.6%)	
L&m blue slim	0.6	7	6	1 (1.3%)		0		0		0		1 (0.6%)	
L&m link	0.7	9	8	1 (1.3%)		0		0		0		1 (0.6%)	
L&m link	0.6	7	6	1 (1.3%)		0		0		0		1 (0.6%)	
L&m orange link	0.4	4	3	1 (1.3%)		0		0		0		1 (0.6%)	
Ld	0.8	10	10	0		1 (2.4%)		0		0		1 (0.6%)	
Ld	0.5	7	9	0		0		1 (2.6%)		0		1 (0.6%)	
Ld blue 100	0.6	8	8	0		0		1 (2.6%)		0		1 (0.6%)	
Ld blue light 100	0.6	8	8	0		1 (2.4%)		0		0		1 (0.6%)	
Ld club platinum	0.5	5	4	0		1 (2.4%)		0		0		1 (0.6%)	
Ld club platinum slim	0.5	5	4	0		0		0		1 (11.1%)		1 (0.6%)	
Ld light 100	0.6	8	8	1 (1.3%)		0		0		0		1 (0.6%)	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.1.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_cigbrand.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.1.5 Summary of Current Cigarette Brands at Admission - Safety Population**

Brand	ISO Nicotine Yield (mg)	ISO Tar Yield (mg)	ISO CO Yield (mg)	THS 2.2 (N=80)		CC (N=41)		SA (N=39)		Enrolled Not Randomized (N=9)		Overall Safety (N=169)	
				n	(%)	n	(%)	n	(%)	n	(%)	n	(%)
Ld pink slim	0.5	5	4	0		0		1 (2.6%)		0		1 (0.6%)	
Ld red	0.8	10	10	0		1 (2.4%)		0		0		1 (0.6%)	
Ld red 100	0.8	10	10	1 (1.3%)		0		0		0		1 (0.6%)	
Ld silver super light	0.3	4	5	0		0		1 (2.6%)		0		1 (0.6%)	
Lucky strike blue	0.6	7	8	0		1 (2.4%)		0		0		1 (0.6%)	
Lucky strike blue light	0.6	7	8	0		1 (2.4%)		0		0		1 (0.6%)	
Lucky strike red original	0.8	10	10	0		1 (2.4%)		0		0		1 (0.6%)	
Marlboro gold 100	0.6	7	8	0		1 (2.4%)		0		0		1 (0.6%)	
Marlboro red	0.7	9	10	1 (1.3%)		0		0		0		1 (0.6%)	
Nevada light	0.5	7	8	0		0		1 (2.6%)		0		1 (0.6%)	
R1 super light	0.1	1	1	1 (1.3%)		0		0		0		1 (0.6%)	
Red white	0.5	6	6	0		0		1 (2.6%)		0		1 (0.6%)	
Red white	0.5	7	8	0		0		1 (2.6%)		0		1 (0.6%)	
Red white original	0.7	10	10	1 (1.3%)		0		0		0		1 (0.6%)	
Rgd blue super slim	0.5	6	6	1 (1.3%)		0		0		0		1 (0.6%)	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.1.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_cigbrand.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.1.5 Summary of Current Cigarette Brands at Admission - Safety Population**

Brand	ISO Nicotine Yield (mg)	ISO Tar Yield (mg)	ISO CO Yield (mg)	THS 2.2 (N=80)		CC (N=41)		SA (N=39)		Enrolled Not Randomized (N=9)		Overall Safety (N=169)	
				n	(%)	n	(%)	n	(%)	n	(%)	n	(%)
She delicious	0.4	6	5	0		1	(2.4%)	0		0		1	(0.6%)
Spike silver	0.4	4	4	1	(1.3%)	0		0		0		1	(0.6%)
Viceroy	0.8	10	10	1	(1.3%)	0		0		0		1	(0.6%)
Viceroy blue light superslim	0.6	7	6	1	(1.3%)	0		0		0		1	(0.6%)
Viceroy blue s-line	0.6	7	7	1	(1.3%)	0		0		0		1	(0.6%)
Viceroy red	0.7	10	10	1	(1.3%)	0		0		0		1	(0.6%)
Viceroy red classic	0.7	10	10	0		0		1	(2.6%)	0		1	(0.6%)
Viceroy red classic 100	0.8	10	10	0		0		0		1	(11.1%)	1	(0.6%)
Viceroy silver s-line	0.4	4	4	0		0		0		1	(11.1%)	1	(0.6%)
Viceroy super slim	0.6	7	6	1	(1.3%)	0		0		0		1	(0.6%)
Winston red c	0.8	10	10	1	(1.3%)	0		0		0		1	(0.6%)

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.1.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_cigbrand.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.1.6 Summary of Medical History - Safety Population**

System Organ Class Preferred Term	THS 2.2 (N=80)			CC (N=41)			SA (N=39)			Enrolled Not Randomized (N=9)			Overall Safety (N=169)		
	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events
Any medical history	33	(41.3%)	44	17	(41.5%)	27	18	(46.2%)	29	5	(55.6%)	11	73	(43.2%)	111
Injury, poisoning and procedural complications	19	(23.8%)	20	9	(22.0%)	9	5	(12.8%)	5	2	(22.2%)	4	35	(20.7%)	38
Clavicle fracture	1	(1.3%)	1	3	(7.3%)	3	1	(2.6%)	1	0			5	(3.0%)	5
Scar	4	(5.0%)	4	0			1	(2.6%)	1	0			5	(3.0%)	5
Ankle fracture	2	(2.5%)	2	2	(4.9%)	2	0			0			4	(2.4%)	4
Hand fracture	2	(2.5%)	2	0			0			1	(11.1%)	1	3	(1.8%)	3
Lower limb fracture	2	(2.5%)	2	1	(2.4%)	1	0			0			3	(1.8%)	3
Upper limb fracture	0			0			1	(2.6%)	1	1	(11.1%)	2	2	(1.2%)	3
Multiple fractures	1	(1.3%)	1	1	(2.4%)	1	0			0			2	(1.2%)	2
Wrist fracture	2	(2.5%)	2	0			0			0			2	(1.2%)	2
Concussion	1	(1.3%)	1	0			0			0			1	(0.6%)	1

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to all subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.1.8.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_medhis.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.1.6 Summary of Medical History - Safety Population**

System Organ Class Preferred Term	THS 2.2 (N=80)			CC (N=41)			SA (N=39)			Enrolled Not Randomized (N=9)			Overall Safety (N=169)		
	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events
Face injury	0			0			0			1 (11.1%)	1		1 (0.6%)	1	
Facial bones fracture	1 (1.3%)		1	0			0			0			1 (0.6%)	1	
Femur fracture	1 (1.3%)		1	0			0			0			1 (0.6%)	1	
Head injury	0			1 (2.4%)		1	0			0			1 (0.6%)	1	
Joint dislocation	1 (1.3%)		1	0			0			0			1 (0.6%)	1	
Joint injury	0			0			1 (2.6%)		1	0			1 (0.6%)	1	
Limb injury	0			0			1 (2.6%)		1	0			1 (0.6%)	1	
Tibia fracture	1 (1.3%)		1	0			0			0			1 (0.6%)	1	
Traumatic arthropathy	1 (1.3%)		1	0			0			0			1 (0.6%)	1	
Ulna fracture	0			1 (2.4%)		1	0			0			1 (0.6%)	1	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to all subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.1.8.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_medhis.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.1.6 Summary of Medical History - Safety Population**

System Organ Class Preferred Term	THS 2.2 (N=80)			CC (N=41)			SA (N=39)			Enrolled Not Randomized (N=9)			Overall Safety (N=169)		
	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events
Surgical and medical procedures	14	(17.5%)	15	8	(19.5%)	8	9	(23.1%)	10	3	(33.3%)	3	34	(20.1%)	36
Appendicectomy	6	(7.5%)	6	2	(4.9%)	2	4	(10.3%)	4	1	(11.1%)	1	13	(7.7%)	13
Caesarean section	3	(3.8%)	3	2	(4.9%)	2	0			0			5	(3.0%)	5
Nasal septal operation	1	(1.3%)	1	2	(4.9%)	2	1	(2.6%)	1	0			4	(2.4%)	4
Cholecystectomy	1	(1.3%)	1	0			1	(2.6%)	1	0			2	(1.2%)	2
Limb operation	1	(1.3%)	1	0			1	(2.6%)	1	0			2	(1.2%)	2
Ankle operation	0			1	(2.4%)	1	0			0			1	(0.6%)	1
Cleft palate repair	1	(1.3%)	1	0			0			0			1	(0.6%)	1
Ectopic pregnancy termination	0			0			1	(2.6%)	1	0			1	(0.6%)	1
Hernia hiatus repair	1	(1.3%)	1	0			0			0			1	(0.6%)	1
Hernia repair	0			0			0			1	(11.1%)	1	1	(0.6%)	1

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to all subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.1.8.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_medhis.sas

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**Table 15.2.1.6 Summary of Medical History - Safety Population**

System Organ Class Preferred Term	THS 2.2 (N=80)			CC (N=41)			SA (N=39)			Enrolled Not Randomized (N=9)			Overall Safety (N=169)		
	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events
Lesion excision	1	(1.3%)	1	0			0			0			1	(0.6%)	1
Meniscus removal	0			1	(2.4%)	1	0			0			1	(0.6%)	1
Ovarian cystectomy	0			0			1	(2.6%)	1	0			1	(0.6%)	1
Rhinoplasty	0			0			0			1	(11.1%)	1	1	(0.6%)	1
Sinus operation	0			0			1	(2.6%)	1	0			1	(0.6%)	1
Infections and infestations	2	(2.5%)	2	1	(2.4%)	1	3	(7.7%)	3	1	(11.1%)	1	7	(4.1%)	7
Appendicitis	1	(1.3%)	1	0			2	(5.1%)	2	0			3	(1.8%)	3
Ear infection	1	(1.3%)	1	0			0			0			1	(0.6%)	1
Hepatitis A	0			1	(2.4%)	1	0			0			1	(0.6%)	1
Respiratory tract infection viral	0			0			1	(2.6%)	1	0			1	(0.6%)	1
Tonsillitis	0			0			0			1	(11.1%)	1	1	(0.6%)	1

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to all subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.1.8.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_medhis.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.1.6 Summary of Medical History - Safety Population**

System Organ Class Preferred Term	THS 2.2 (N=80)			CC (N=41)			SA (N=39)			Enrolled Not Randomized (N=9)			Overall Safety (N=169)		
	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events
Reproductive system and breast disorders	1	(1.3%)	1	1	(2.4%)	1	2	(5.1%)	2	0			4	(2.4%)	4
Endometriosis	0			0			1	(2.6%)	1				1	(0.6%)	1
Fallopian tube obstruction	0			1	(2.4%)	1	0						1	(0.6%)	1
Genital tract inflammation	1	(1.3%)	1	0			0						1	(0.6%)	1
Menstruation irregular	0			0			1	(2.6%)	1				1	(0.6%)	1
Musculoskeletal and connective tissue disorders	0			1	(2.4%)	2	1	(2.6%)	1	1	(11.1%)	1	3	(1.8%)	4
Spinal osteoarthritis				1	(2.4%)	2	0			0			1	(0.6%)	2
Arthropathy				0			0			1	(11.1%)	1	1	(0.6%)	1
Intervertebral disc disorder				0			1	(2.6%)	1	0			1	(0.6%)	1

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to all subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.1.8.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_medhis.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.1.6 Summary of Medical History - Safety Population**

System Organ Class Preferred Term	THS 2.2 (N=80)			CC (N=41)			SA (N=39)			Enrolled Not Randomized (N=9)			Overall Safety (N=169)		
	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events
Eye disorders	0			1 (2.4%)	1		2 (5.1%)	2		0			3 (1.8%)	3	
Astigmatism				0			1 (2.6%)	1					1 (0.6%)	1	
Eyelid ptosis				1 (2.4%)	1		0						1 (0.6%)	1	
Strabismus				0			1 (2.6%)	1					1 (0.6%)	1	
Respiratory, thoracic and mediastinal disorders	0			1 (2.4%)	2		1 (2.6%)	1		0			2 (1.2%)	3	
Cough				0			1 (2.6%)	1					1 (0.6%)	1	
Nasal polyps				1 (2.4%)	1		0						1 (0.6%)	1	
Nasal septum deviation				1 (2.4%)	1		0						1 (0.6%)	1	
Immune system disorders	0			0			2 (5.1%)	2		0			2 (1.2%)	2	
Allergy to chemicals							1 (2.6%)	1					1 (0.6%)	1	
Food allergy							1 (2.6%)	1					1 (0.6%)	1	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to all subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.1.8.1

Path: /cvm/projects/prj/development/000000106324/dev/tables/t_medhis.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.1.6 Summary of Medical History - Safety Population**

System Organ Class Preferred Term	THS 2.2 (N=80)			CC (N=41)			SA (N=39)			Enrolled Not Randomized (N=9)			Overall Safety (N=169)		
	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events
Pregnancy, puerperium and perinatal conditions	0			2 (4.9%)	2		0			0			2 (1.2%)	2	
Gestational hypertension				1 (2.4%)	1								1 (0.6%)	1	
Previous caesarean section				1 (2.4%)	1								1 (0.6%)	1	
Renal and urinary disorders	1 (1.3%)	1		0			1 (2.6%)	1		0			2 (1.2%)	2	
Nephritis	0						1 (2.6%)	1					1 (0.6%)	1	
Nephrolithiasis	1 (1.3%)	1					0						1 (0.6%)	1	
Skin and subcutaneous tissue disorders	1 (1.3%)	1		0			1 (2.6%)	1		0			2 (1.2%)	2	
Acne	1 (1.3%)	1					0						1 (0.6%)	1	
Dermatitis atopic	0						1 (2.6%)	1					1 (0.6%)	1	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to all subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.1.8.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_medhis.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.1.6 Summary of Medical History - Safety Population**

System Organ Class Preferred Term	THS 2.2 (N=80)			CC (N=41)			SA (N=39)			Enrolled Not Randomized (N=9)			Overall Safety (N=169)		
	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events
Investigations	0			0			0			1 (11.1%)	2		1 (0.6%)	2	
Arthroscopy										1 (11.1%)	2		1 (0.6%)	2	
Cardiac disorders	1 (1.3%)		1	0			0			0			1 (0.6%)	1	
Arrhythmia	1 (1.3%)		1										1 (0.6%)	1	
Congenital, familial and genetic disorders	1 (1.3%)		1	0			0			0			1 (0.6%)	1	
Cryptorchism	1 (1.3%)		1										1 (0.6%)	1	
Endocrine disorders	0			0			1 (2.6%)	1		0			1 (0.6%)	1	
Hyperthyroidism							1 (2.6%)	1					1 (0.6%)	1	
Gastrointestinal disorders	1 (1.3%)		1	0			0			0			1 (0.6%)	1	
Constipation	1 (1.3%)		1										1 (0.6%)	1	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to all subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.1.8.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_medhis.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.1.6 Summary of Medical History - Safety Population**

System Organ Class Preferred Term	THS 2.2 (N=80)			CC (N=41)			SA (N=39)			Enrolled Not Randomized (N=9)			Overall Safety (N=169)		
	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events
Nervous system disorders	1	(1.3%)	1	0			0			0			1	(0.6%)	1
Brain stem haematoma	1	(1.3%)	1										1	(0.6%)	1
Psychiatric disorders	0			1	(2.4%)	1	0			0			1	(0.6%)	1
Depression				1	(2.4%)	1							1	(0.6%)	1

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to all subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.1.8.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_medhis.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.1.7 Summary of Concomitant Diseases - Safety Population**

System Organ Class Preferred Term	THS 2.2 (N=80)			CC (N=41)			SA (N=39)			Enrolled Not Randomized (N=9)			Overall Safety (N=169)		
	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events
Any concomitant disease	27	(33.8%)	32	9	(22.0%)	11	13	(33.3%)	15	3	(33.3%)	3	52	(30.8%)	61
Eye disorders	14	(17.5%)	14	3	(7.3%)	3	3	(7.7%)	3	1	(11.1%)	1	21	(12.4%)	21
Hypermetropia	7	(8.8%)	7	1	(2.4%)	1	1	(2.6%)	1	0			9	(5.3%)	9
Myopia	5	(6.3%)	5	1	(2.4%)	1	2	(5.1%)	2	1	(11.1%)	1	9	(5.3%)	9
Astigmatism	1	(1.3%)	1	1	(2.4%)	1	0			0			2	(1.2%)	2
Presbyopia	1	(1.3%)	1	0			0			0			1	(0.6%)	1
Immune system disorders	4	(5.0%)	4	3	(7.3%)	3	3	(7.7%)	3	1	(11.1%)	1	11	(6.5%)	11
Allergy to arthropod bite	1	(1.3%)	1	0			1	(2.6%)	1	0			2	(1.2%)	2
Allergy to chemicals	0			1	(2.4%)	1	1	(2.6%)	1	0			2	(1.2%)	2
Allergy to plants	1	(1.3%)	1	1	(2.4%)	1	0			0			2	(1.2%)	2
Multiple allergies	2	(2.5%)	2	0			0			0			2	(1.2%)	2

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to all subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.1.8.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_condis.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.1.7 Summary of Concomitant Diseases - Safety Population**

System Organ Class Preferred Term	THS 2.2 (N=80)			CC (N=41)			SA (N=39)			Enrolled Not Randomized (N=9)			Overall Safety (N=169)		
	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events
Allergy to metals	0			1 (2.4%)		1	0			0			1 (0.6%)		1
Drug hypersensitivity	0			0			0			1 (11.1%)		1	1 (0.6%)		1
Seasonal allergy	0			0			1 (2.6%)		1	0			1 (0.6%)		1
Injury, poisoning and procedural complications	3 (3.8%)		4	1 (2.4%)		2	5 (12.8%)		5	0			9 (5.3%)		11
Scar	3 (3.8%)		4	1 (2.4%)		2	3 (7.7%)		3				7 (4.1%)		9
Eye injury	0			0			1 (2.6%)		1				1 (0.6%)		1
Rib fracture	0			0			1 (2.6%)		1				1 (0.6%)		1
Skin and subcutaneous tissue disorders	3 (3.8%)		3	0			1 (2.6%)		1	0			4 (2.4%)		4
Acne	2 (2.5%)		2				1 (2.6%)		1				3 (1.8%)		3
Hyperkeratosis	1 (1.3%)		1				0						1 (0.6%)		1

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to all subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.1.8.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_condis.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.1.7 Summary of Concomitant Diseases - Safety Population**

System Organ Class Preferred Term	THS 2.2 (N=80)			CC (N=41)			SA (N=39)			Enrolled Not Randomized (N=9)			Overall Safety (N=169)		
	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events
Social circumstances	1	(1.3%)	2	1	(2.4%)	1	1	(2.6%)	1	0			3	(1.8%)	4
Tattoo	1	(1.3%)	2	1	(2.4%)	1	1	(2.6%)	1				3	(1.8%)	4
Respiratory, thoracic and mediastinal disorders	1	(1.3%)	1	0			1	(2.6%)	1	1	(11.1%)	1	3	(1.8%)	3
Cough	0						1	(2.6%)	1	0			1	(0.6%)	1
Dysphonia	1	(1.3%)	1				0			0			1	(0.6%)	1
Nasal septum deviation	0						0			1	(11.1%)	1	1	(0.6%)	1
Surgical and medical procedures	2	(2.5%)	2	0			0			0			2	(1.2%)	2
Bunion operation	1	(1.3%)	1										1	(0.6%)	1
Nasal septal operation	1	(1.3%)	1										1	(0.6%)	1

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to all subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.1.8.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_condis.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.1.7 Summary of Concomitant Diseases - Safety Population**

System Organ Class Preferred Term	THS 2.2 (N=80)			CC (N=41)			SA (N=39)			Enrolled Not Randomized (N=9)			Overall Safety (N=169)		
	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events
Blood and lymphatic system disorders	0			0			1 (2.6%)	1		0			1 (0.6%)	1	
Lymphadenopathy							1 (2.6%)	1					1 (0.6%)	1	
Gastrointestinal disorders	1 (1.3%)		1	0			0			0			1 (0.6%)	1	
Dental caries	1 (1.3%)		1										1 (0.6%)	1	
Infections and infestations	1 (1.3%)		1	0			0			0			1 (0.6%)	1	
Laryngitis	1 (1.3%)		1										1 (0.6%)	1	
Musculoskeletal and connective tissue disorders	0			1 (2.4%)		1	0			0			1 (0.6%)	1	
Scoliosis				1 (2.4%)		1							1 (0.6%)	1	
Neoplasms benign, malignant and unspecified (incl cysts and polyps)	0			1 (2.4%)		1	0			0			1 (0.6%)	1	
Uterine leiomyoma				1 (2.4%)		1							1 (0.6%)	1	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to all subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.1.8.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_condis.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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15.2.2 Product Use Tables

**Table 15.2.2.1 Descriptive Statistics of Use of THS 2.2 Product and CC - Safety Population**

Occasion	Study Day - Product Use	Number of Uses	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
THS 2.2 test		1	n (%)	62 (77.5%)	28 (68.3%)	29 (74.4%)	7 (77.8%)	126 (74.6%)
		2	n (%)	12 (15.0%)	11 (26.8%)	6 (15.4%)	2 (22.2%)	31 (18.3%)
		3	n (%)	6 (7.5%)	2 (4.9%)	4 (10.3%)	0	12 (7.1%)
	Day -2 - THS 2.2		n	80	41	39	9	169
			Mean (SD)	1.3 (0.61)	1.4 (0.59)	1.4 (0.67)	1.2 (0.45)	1.3 (0.61)
			Median	1.0	1.0	1.0	1.0	1.0
			Min, Max	1, 3	1, 3	1, 3	1, 2	1, 3
Prior to randomization	Day -1 - CC		n	80	41	39	9	169
			Mean (SD)	14.2 (3.46)	14.0 (3.11)	14.7 (3.65)	15.3 (4.00)	14.3 (3.44)
			Median	14.0	14.0	14.0	17.0	14.0
			Min, Max	8, 29	9, 23	7, 26	8, 20	7, 29

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.2.1.1, 15.3.2.1.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_ustatss.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.2.1 Descriptive Statistics of Use of THS 2.2 Product and CC - Safety Population**

Occasion	Study Day - Product Use	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Prior to randomization	Day -1 - THS 2.2	n	0	0	0	0	0
		Mean (SD)					
		Median					
		Min, Max					
	Day 0 - CC	n	80	41	39	9	169
		Mean (SD)	16.0 (3.45)	16.2 (4.05)	17.0 (4.17)	14.7 (4.00)	16.2 (3.81)
		Median	16.0	16.0	17.0	14.0	16.0
		Min, Max	9, 33	10, 32	8, 32	9, 21	8, 33
	Day 0 - THS 2.2	n	0	0	0	0	0
		Mean (SD)					
		Median					
		Min, Max					

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.2.1.1, 15.3.2.1.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_ustatss.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.2.1 Descriptive Statistics of Use of THS 2.2 Product and CC - Safety Population**

Occasion	Study Day - Product Use	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
After randomization	Day 1 - CC	n	0	41	0	0	41
		Mean (SD)		14.5 (3.63)			14.5 (3.63)
		Median		14.0			14.0
		Min, Max		6, 22			6, 22
	Day 1 - THS 2.2	n	80	0	0	0	80
		Mean (SD)	14.9 (6.14)				14.9 (6.14)
		Median	14.0				14.0
		Min, Max	5, 50				5, 50
	Day 2 - CC	n	0	41	0	0	41
		Mean (SD)		15.1 (3.79)			15.1 (3.79)
		Median		15.0			15.0
		Min, Max		8, 24			8, 24

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.2.1.1, 15.3.2.1.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_ustatss.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.2.1 Descriptive Statistics of Use of THS 2.2 Product and CC - Safety Population**

Occasion	Study Day - Product Use	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
After randomization	Day 2 - THS 2.2	n	80	0	0	0	80
		Mean (SD)	17.3 (6.25)				17.3 (6.25)
		Median	17.0				17.0
		Min, Max	6, 40				6, 40
	Day 3 - CC	n	0	41	0	0	41
		Mean (SD)		14.9 (3.49)			14.9 (3.49)
		Median		15.0			15.0
		Min, Max		8, 22			8, 22
	Day 3 - THS 2.2	n	80	0	0	0	80
		Mean (SD)	18.2 (5.94)				18.2 (5.94)
		Median	17.0				17.0
		Min, Max	6, 38				6, 38

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.2.1.1, 15.3.2.1.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_ustatss.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.2.1 Descriptive Statistics of Use of THS 2.2 Product and CC - Safety Population**

Occasion	Study Day - Product Use	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
After randomization	Day 4 - CC	n	0	41	0	0	41
		Mean (SD)		14.3 (3.43)			14.3 (3.43)
		Median		14.0			14.0
		Min, Max		8, 21			8, 21
	Day 4 - THS 2.2	n	79	0	0	0	79
		Mean (SD)	18.5 (6.69)				18.5 (6.69)
		Median	17.0				17.0
		Min, Max	9, 50				9, 50
	Day 5 - CC	n	0	41	0	0	41
		Mean (SD)		16.6 (3.79)			16.6 (3.79)
		Median		16.0			16.0
		Min, Max		10, 26			10, 26

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.2.1.1, 15.3.2.1.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_ustatss.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.2.1 Descriptive Statistics of Use of THS 2.2 Product and CC - Safety Population**

Occasion	Study Day - Product Use	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
After randomization	Day 5 - THS 2.2	n	79	0	0	0	79
		Mean (SD)	20.7 (8.09)				20.7 (8.09)
		Median	20.0				20.0
		Min, Max	9, 60				9, 60

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.2.1.1, 15.3.2.1.2

Path: /cvm/projects/prj/development/000000106324/dev/tables/t_ustatss.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL



15.2.3 Primary Endpoints Summary Tables

**Table 15.2.3.1 Analysis of Evening Blood COHb (%) versus CC on Day 5 - FAS**

Variable	Statistic	THS 2.2 (N=80)	CC (N=41)	THS : CC Ratio (%)
Blood COHb (%)	n	79	41	
	Geometric LS Mean (CV%)	1.06	4.53	23.45 (16.84)
	95% CI	1.02, 1.11	4.29, 4.77	22.00, 24.99
	p-value			<.001

Note: CC = Conventional cigarettes; THS = Tobacco Heating System.

Note: Adjusted geometric least squares (LS) means and confidence intervals (CIs) from an ANCOVA model conducted on log-transformed Day 5 values with log-transformed baseline value, study arm, sex and CC consumption reported at screening as fixed effect factors. Geometrical CV% of the ratio is estimated from the residual mean squares.

Baseline is defined as the last assessment prior to Day 1 product use for THS 2.2 and CC subjects and prior to 06:29 AM on Day 1 for SA subjects.

Note: p-value for one-sided test for comparison between products.

Appendix 15.4.3.1 and 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/tl_anlcohbfas.sas

Program Run: 27OCT14 cvn_ahedge Program Status: FINAL

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**Table 15.2.3.2 Analysis of MHBMA Urinary Concentration Adjusted for Creatinine (pg/mg creat) versus CC on Day 5 - FAS**

Variable	Statistic	THS 2.2 (N=80)	CC (N=41)	THS : CC Ratio (%)
Urinary MHBMA (pg/mg creat)	n	79	41	
	Geometric LS Mean (CV%)	194.05	2314.37	8.38 (54.57)
	95% CI	173.23, 217.38	1975.39, 2711.52	6.89, 10.20
	p-value			<.001

Note: CC = Conventional cigarettes; THS = Tobacco Heating System.

Note: Adjusted geometric least squares (LS) means and confidence intervals (CIs) from an ANCOVA model conducted on log-transformed Day 5 values with log-transformed baseline value, study arm, sex and CC consumption reported at screening as fixed effect factors. Geometrical CV% of the ratio is estimated from the residual mean squares.

Baseline is defined as the last assessment prior to Day 1 product use for THS 2.2 and CC subjects and prior to 06:29 AM on Day 1 for SA subjects.

Note: p-value for one-sided test for comparison between products.

Appendix 15.4.3.2 and 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/tl_anlmhbmafes.sas

Program Run: 27OCT14 cvn_ahedge Program Status: FINAL

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**Table 15.2.3.3 3-HPMA Urinary Concentration Adjusted for Creatinine (ng/mg creat) versus CC on Day 5 - FAS**

Variable	Statistic	THS 2.2 (N=80)	CC (N=41)	THS : CC Ratio (%)
Urinary 3-HPMA (ng/mg n creat)		79	41	
	Geometric LS Mean (CV%)	398.87	958.03	41.63 (26.10)
	95% CI	376.74, 422.30	884.99, 1037.09	37.75, 45.91
	p-value			<.001

Note: CC = Conventional cigarettes; THS = Tobacco Heating System.

Note: Adjusted geometric least squares (LS) means and confidence intervals (CIs) from an ANCOVA model conducted on log-transformed Day 5 values with log-transformed baseline value, study arm, sex and CC consumption reported at screening as fixed effect factors. Geometrical CV% of the ratio is estimated from the residual mean squares.

Baseline is defined as the last assessment prior to Day 1 product use for THS 2.2 and CC subjects and prior to 06:29 AM on Day 1 for SA subjects.

Note: p-value for one-sided test for comparison between products.

Appendix 15.4.3.3 and 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/tl_anl3hpmafas.sas

Program Run: 27OCT14 cvn_ahedge Program Status: FINAL

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**Table 15.2.3.4 S-PMA Urinary Concentration Adjusted for Creatinine (pg/mg creat) versus CC on Day 5 - FAS**

Variable	Statistic	THS 2.2 (N=80)	CC (N=41)	THS : CC Ratio (%)
Urinary S-PMA (pg/mg creat)	n	79	41	
	Geometric LS Mean (CV%)	164.51	2748.16	5.99 (37.30)
	95% CI	151.82, 178.25	2457.56, 3073.13	5.21, 6.87
	p-value			<.001

Note: CC = Conventional cigarettes; THS = Tobacco Heating System.

Note: Adjusted geometric least squares (LS) means and confidence intervals (CIs) from an ANCOVA model conducted on log-transformed Day 5 values with log-transformed baseline value, study arm, sex and CC consumption reported at screening as fixed effect factors. Geometrical CV% of the ratio is estimated from the residual mean squares.

Baseline is defined as the last assessment prior to Day 1 product use for THS 2.2 and CC subjects and prior to 06:29 AM on Day 1 for SA subjects.

Note: p-value for one-sided test for comparison between products.

Appendix 15.4.3.4 and 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/tl_anlspmaf.sas

Program Run: 27OCT14 cvn_ahedge Program Status: FINAL

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**Table 15.2.3.5 Descriptive Statistics of % Change from Baseline in Primary Biomarkers on Day 5 - FAS**

Variable	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)
Evening COHb (%)	n	79	41	39
	BLOQ - n (%)	0	0	0
	Mean (SD)	-76.20 (8.498)	-1.16 (22.649)	-79.24 (5.676)
	95% CI	-78.11, -74.29	-8.31, 5.99	-81.09, -77.40
	Median	-77.78	-9.38	-79.07
	Q25, Q75	-82.26, -72.97	-15.39, 12.91	-83.64, -75.60
	Min, Max	-90.3, -42.9	-42.9, 62.9	-88.6, -66.7

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_pbio.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.3.5 Descriptive Statistics of % Change from Baseline in Primary Biomarkers on Day 5 - FAS**

Variable	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)
3-HPMA Urinary Concentration Adjusted for Creatinine (ng/mg creat)	n	79	41	39
	BLOQ - n (%)	1 (1.3%)	0	0
	Mean (SD)	-49.68 (20.696)	20.28 (31.247)	-69.45 (11.761)
	95% CI	-54.32, -45.04	10.41, 30.14	-73.27, -65.63
	Median	-54.93	9.38	-72.54
	Q25, Q75	-62.28, -44.38	-1.19, 41.70	-77.67, -63.45
	Min, Max	-73.9, 44.6	-44.8, 115.1	-85.8, -44.5

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: BLOQ = number of observations imputed using half limit of quantification (20.0 ng/mL).

Appendix 15.3.3.1

Path: /cvm/projects/prj/development/000000106324/dev/tables/t_pbio.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.5 Descriptive Statistics of % Change from Baseline in Primary Biomarkers on Day 5 - FAS**

Variable	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)
MHBMA Urinary Concentration Adjusted for Creatinine (pg/mg creat)	n	79	41	39
	BLOQ - n (%)	34 (43.0%)	1 (2.4%)	16 (41.0%)
	Mean (SD)	-84.98 (14.906)	6.89 (26.158)	-79.56 (27.037)
	95% CI	-88.32, -81.63	-1.37, 15.15	-88.33, -70.79
	Median	-92.03	5.28	-92.83
	Q25, Q75	-94.62, -82.02	-6.85, 23.26	-96.69, -70.55
	Min, Max	-98.1, -41.6	-50.0, 55.1	-98.1, -6.4

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: BLOQ = number of observations imputed using half limit of quantification (0.100 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_pbio.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.5 Descriptive Statistics of % Change from Baseline in Primary Biomarkers on Day 5 - FAS**

Variable	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)
S-PMA Urinary Concentration Adjusted for Creatinine (pg/mg creat)	n	79	41	39
	BLOQ - n (%)	3 (3.8%)	0	0
	Mean (SD)	-92.03 (5.222)	9.48 (31.561)	-91.52 (6.754)
	95% CI	-93.20, -90.85	-0.48, 19.45	-93.72, -89.33
	Median	-93.30	4.24	-93.82
	Q25, Q75	-95.28, -91.56	-8.99, 21.22	-96.31, -88.82
	Min, Max	-97.7, -75.3	-46.4, 127.5	-97.9, -70.9

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: BLOQ = number of observations imputed using half limit of quantification (0.0250 ng/mL).

Appendix 15.3.3.1

Path: /cvm/projects/prj/development/000000106324/dev/tables/t_pbio.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.5.1 Descriptive Statistics of % Change from Baseline in Primary Biomarkers on Day 5 by Sex - FAS**

Variable	Statistic	THS 2.2		CC	
		Male (N=39)	Female (N=41)	Male (N=21)	Female (N=20)
Evening COHb (%)	n	39	40	21	20
	BLOQ - n (%)	0	0	0	0
	Mean (SD)	-78.34 (4.579)	-74.12 (10.723)	1.37 (24.851)	-3.82 (20.382)
	95% CI	-79.82, -76.85	-77.55, -70.68	-9.95, 12.68	-13.36, 5.73
	Median	-78.43	-75.58	-5.13	-10.94
	Q25, Q75	-82.26, -75.00	-82.39, -68.57	-15.00, 19.24	-21.09, 11.66
	Min, Max	-90.3, -69.4	-88.6, -42.9	-42.9, 62.9	-33.3, 38.1

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_pbios.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.3.5.1 Descriptive Statistics of % Change from Baseline in Primary Biomarkers on Day 5 by Sex - FAS**

Variable	Statistic	SA	
		Male (N=20)	Female (N=19)
Evening COHb (%)	n	20	19
	BLOQ - n (%)	0	0
	Mean (SD)	-80.09 (6.202)	-78.36 (5.078)
	95% CI	-82.99, -77.18	-80.81, -75.90
	Median	-81.42	-78.79
	Q25, Q75	-83.98, -75.79	-82.00, -75.60
	Min, Max	-88.6, -66.7	-87.5, -67.7

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_pbios.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.3.5.1 Descriptive Statistics of % Change from Baseline in Primary Biomarkers on Day 5 by Sex - FAS**

Variable	Statistic	THS 2.2		CC	
		Male (N=39)	Female (N=41)	Male (N=21)	Female (N=20)
3-HPMA Urinary Concentration Adjusted for Creatinine (ng/mg creat)	n	39	40	21	20
	BLOQ - n (%)	0	1 (2.5%)	0	0
	Mean (SD)	-51.26 (12.615)	-48.14 (26.394)	23.91 (35.202)	16.46 (26.854)
	95% CI	-55.35, -47.16	-56.58, -39.69	7.89, 39.94	3.89, 29.03
	Median	-54.41	-57.12	18.14	7.70
	Q25, Q75	-58.85, -44.29	-65.47, -46.13	2.26, 41.70	-4.83, 42.81
	Min, Max	-72.8, -23.0	-73.9, 44.6	-44.8, 115.1	-19.3, 69.8

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: BLOQ = number of observations imputed using half limit of quantification (20.0 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_pbios.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.3.5.1 Descriptive Statistics of % Change from Baseline in Primary Biomarkers on Day 5 by Sex - FAS**

Variable	Statistic	SA	
		Male (N=20)	Female (N=19)
3-HPMA Urinary Concentration Adjusted for Creatinine (ng/mg creat)	n	20	19
	BLOQ - n (%)	0	0
	Mean (SD)	-67.83 (12.954)	-71.16 (10.434)
	95% CI	-73.89, -61.76	-76.19, -66.13
	Median	-73.44	-72.54
	Q25, Q75	-77.63, -57.49	-78.68, -67.04
	Min, Max	-84.4, -44.9	-85.8, -44.5

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: BLOQ = number of observations imputed using half limit of quantification (20.0 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_pbios.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.3.5.1 Descriptive Statistics of % Change from Baseline in Primary Biomarkers on Day 5 by Sex - FAS**

Variable	Statistic	THS 2.2		CC	
		Male (N=39)	Female (N=41)	Male (N=21)	Female (N=20)
MHBMA Urinary Concentration Adjusted for Creatinine (pg/mg creat)	n	39	40	21	20
	BLOQ - n (%)	13 (33.3%)	21 (52.5%)	0	1 (5.0%)
	Mean (SD)	-83.89 (16.343)	-86.04 (13.482)	6.03 (27.582)	7.79 (25.261)
	95% CI	-89.19, -78.58	-90.36, -81.72	-6.53, 18.59	-4.04, 19.61
	Median	-92.26	-91.84	1.88	8.29
	Q25, Q75	-94.39, -77.09	-94.81, -84.91	-10.44, 23.26	-3.87, 27.27
	Min, Max	-98.1, -41.6	-97.6, -52.1	-34.3, 55.1	-50.0, 44.6

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: BLOQ = number of observations imputed using half limit of quantification (0.100 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_pbios.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.3.5.1 Descriptive Statistics of % Change from Baseline in Primary Biomarkers on Day 5 by Sex - FAS**

Variable	Statistic	SA	
		Male (N=20)	Female (N=19)
MHBMA Urinary Concentration Adjusted for Creatinine (pg/mg creat)	n	20	19
	BLOQ - n (%)	5 (25.0%)	11 (57.9%)
	Mean (SD)	-77.02 (28.371)	-82.23 (26.057)
	95% CI	-90.30, -63.74	-94.79, -69.66
	Median	-91.22	-93.04
	Q25, Q75	-96.76, -63.78	-96.28, -87.37
	Min, Max	-98.1, -6.4	-97.7, -13.8

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: BLOQ = number of observations imputed using half limit of quantification (0.100 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_pbios.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.3.5.1 Descriptive Statistics of % Change from Baseline in Primary Biomarkers on Day 5 by Sex - FAS**

Variable	Statistic	THS 2.2		CC	
		Male (N=39)	Female (N=41)	Male (N=21)	Female (N=20)
S-PMA Urinary Concentration Adjusted for Creatinine (pg/mg creat)	n	39	40	21	20
	BLOQ - n (%)	1 (2.6%)	2 (5.0%)	0	0
	Mean (SD)	-92.54 (4.893)	-91.53 (5.539)	16.33 (40.737)	2.29 (15.618)
	95% CI	-94.13, -90.95	-93.30, -89.75	-2.21, 34.88	-5.02, 9.61
	Median	-93.81	-93.05	5.83	3.70
	Q25, Q75	-95.86, -92.05	-95.13, -90.66	-3.56, 31.45	-11.12, 13.69
	Min, Max	-97.7, -75.6	-97.7, -75.3	-46.4, 127.5	-27.2, 25.0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: BLOQ = number of observations imputed using half limit of quantification (0.0250 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_pbios.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.3.5.1 Descriptive Statistics of % Change from Baseline in Primary Biomarkers on Day 5 by Sex - FAS**

Variable	Statistic	SA	
		Male (N=20)	Female (N=19)
S-PMA Urinary Concentration Adjusted for Creatinine (pg/mg creat) n		20	19
BLOQ - n (%)		0	0
Mean (SD)		-91.46 (7.303)	-91.59 (6.325)
95% CI		-94.89, -88.04	-94.64, -88.53
Median		-93.67	-94.22
Q25, Q75		-96.61, -90.09	-94.83, -88.73
Min, Max		-97.9, -74.0	-96.6, -70.9

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: BLOQ = number of observations imputed using half limit of quantification (0.0250 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_pbios.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.3.5.2 Descriptive Statistics of % Change from Baseline in Primary Biomarkers on Day 5 by Cigarette Consumption - FAS**

Variable	Statistic	THS 2.2		CC	
		10-19 cig/day (N=41)	>19 cig/day (N=39)	10-19 cig/day (N=21)	>19 cig/day (N=20)
Evening COHb (%)	n	40	39	21	20
	BLOQ - n (%)	0	0	0	0
	Mean (SD)	-74.36 (9.349)	-78.09 (7.163)	0.49 (27.797)	-2.90 (16.139)
	95% CI	-77.35, -71.36	-80.41, -75.76	-12.16, 13.15	-10.46, 4.66
	Median	-75.28	-80.65	-9.43	-3.37
	Q25, Q75	-79.46, -72.36	-82.82, -74.35	-15.16, 24.08	-16.67, 10.05
	Min, Max	-88.6, -42.9	-90.3, -58.3	-42.9, 62.9	-29.7, 25.0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_pbioc.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.3.5.2 Descriptive Statistics of % Change from Baseline in Primary Biomarkers on Day 5 by Cigarette Consumption - FAS**

Variable	Statistic	SA	
		10-19 cig/day (N=19)	>19 cig/day (N=20)
Evening COHb (%)	n	19	20
	BLOQ - n (%)	0	0
	Mean (SD)	-78.06 (5.189)	-80.36 (6.017)
	95% CI	-80.57, -75.56	-83.18, -77.54
	Median	-78.79	-79.53
	Q25, Q75	-82.00, -75.00	-86.60, -75.90
	Min, Max	-83.9, -66.7	-88.6, -68.0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_pbioc.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.3.5.2 Descriptive Statistics of % Change from Baseline in Primary Biomarkers on Day 5 by Cigarette Consumption - FAS**

Variable	Statistic	THS 2.2		CC	
		10-19 cig/day (N=41)	>19 cig/day (N=39)	10-19 cig/day (N=21)	>19 cig/day (N=20)
3-HPMA Urinary Concentration Adjusted for Creatinine (ng/mg creat)	n	40	39	21	20
	BLOQ - n (%)	0	1 (2.6%)	0	0
	Mean (SD)	-45.64 (22.353)	-53.82 (18.208)	22.32 (35.359)	18.13 (27.017)
	95% CI	-52.79, -38.49	-59.73, -47.92	6.22, 38.42	5.48, 30.78
	Median	-50.59	-57.61	7.99	18.85
	Q25, Q75	-58.05, -40.50	-68.35, -47.02	-1.19, 47.18	-1.41, 37.69
	Min, Max	-73.9, 44.6	-73.7, 16.8	-19.3, 115.1	-44.8, 62.4

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: BLOQ = number of observations imputed using half limit of quantification (20.0 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_pbioc.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.3.5.2 Descriptive Statistics of % Change from Baseline in Primary Biomarkers on Day 5 by Cigarette Consumption - FAS**

Variable	Statistic	SA	
		10-19 cig/day (N=19)	>19 cig/day (N=20)
3-HPMA Urinary Concentration Adjusted for Creatinine (ng/mg creat)	n	19	20
	BLOQ - n (%)	0	0
	Mean (SD)	-64.31 (12.076)	-74.34 (9.321)
	95% CI	-70.13, -58.48	-78.71, -69.97
	Median	-67.05	-75.04
	Q25, Q75	-74.99, -52.00	-79.88, -71.85
	Min, Max	-84.4, -44.5	-85.8, -44.9

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: BLOQ = number of observations imputed using half limit of quantification (20.0 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_pbioc.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.3.5.2 Descriptive Statistics of % Change from Baseline in Primary Biomarkers on Day 5 by Cigarette Consumption - FAS**

Variable	Statistic	THS 2.2		CC	
		10-19 cig/day (N=41)	>19 cig/day (N=39)	10-19 cig/day (N=21)	>19 cig/day (N=20)
MHBMA Urinary Concentration Adjusted for Creatinine (pg/mg creat)	n	40	39	21	20
	BLOQ - n (%)	22 (55.0%)	12 (30.8%)	0	1 (5.0%)
	Mean (SD)	-82.98 (16.664)	-87.03 (12.749)	3.08 (25.410)	10.89 (26.983)
	95% CI	-88.31, -77.64	-91.16, -82.89	-8.49, 14.65	-1.75, 23.52
	Median	-92.39	-90.79	-0.71	17.83
	Q25, Q75	-94.75, -70.59	-94.45, -85.60	-10.44, 15.25	-6.28, 30.01
	Min, Max	-97.6, -41.6	-98.1, -42.3	-35.5, 55.1	-50.0, 51.3

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: BLOQ = number of observations imputed using half limit of quantification (0.100 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_pbioc.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.3.5.2 Descriptive Statistics of % Change from Baseline in Primary Biomarkers on Day 5 by Cigarette Consumption - FAS**

Variable	Statistic	SA	
		10-19 cig/day (N=19)	>19 cig/day (N=20)
MHBMA Urinary Concentration Adjusted for Creatinine (pg/mg creat)	n	19	20
	BLOQ - n (%)	10 (52.6%)	6 (30.0%)
	Mean (SD)	-84.22 (22.683)	-75.13 (30.532)
	95% CI	-95.16, -73.28	-89.42, -60.83
	Median	-93.62	-91.28
	Q25, Q75	-96.83, -86.33	-96.26, -62.77
	Min, Max	-98.1, -13.8	-98.0, -6.4

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: BLOQ = number of observations imputed using half limit of quantification (0.100 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_pbioc.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.3.5.2 Descriptive Statistics of % Change from Baseline in Primary Biomarkers on Day 5 by Cigarette Consumption - FAS**

Variable	Statistic	THS 2.2		CC	
		10-19 cig/day (N=41)	>19 cig/day (N=39)	10-19 cig/day (N=21)	>19 cig/day (N=20)
S-PMA Urinary Concentration Adjusted for Creatinine (pg/mg creat)	n	40	39	21	20
	BLOQ - n (%)	1 (2.5%)	2 (5.1%)	0	0
	Mean (SD)	-90.89 (5.577)	-93.19 (4.615)	7.98 (35.239)	11.07 (28.014)
	95% CI	-92.68, -89.11	-94.69, -91.69	-8.07, 24.02	-2.05, 24.18
	Median	-92.87	-94.51	1.02	8.19
	Q25, Q75	-94.19, -89.00	-95.86, -92.35	-9.18, 13.56	0.35, 25.37
	Min, Max	-97.7, -75.3	-97.7, -75.6	-33.4, 127.5	-46.4, 83.8

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: BLOQ = number of observations imputed using half limit of quantification (0.0250 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_pbioc.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.3.5.2 Descriptive Statistics of % Change from Baseline in Primary Biomarkers on Day 5 by Cigarette Consumption - FAS**

Variable	Statistic	SA	
		10-19 cig/day (N=19)	>19 cig/day (N=20)
S-PMA Urinary Concentration Adjusted for Creatinine (pg/mg creat) n		19	20
BLOQ - n (%)		0	0
Mean (SD)		-92.42 (6.023)	-90.67 (7.436)
95% CI		-95.33, -89.51	-94.16, -87.19
Median		-94.18	-93.67
Q25, Q75		-96.49, -91.49	-96.24, -88.46
Min, Max		-97.0, -70.9	-97.9, -74.0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: BLOQ = number of observations imputed using half limit of quantification (0.0250 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_pbioc.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.3.6 Descriptive Statistics of COHb (%) - FAS**

Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)	
		Raw value	%Change (*)	Raw value	%Change (*)	Raw value	%Change (*)
Baseline	n	80		41		39	
	BLOQ - n (%)	0		0		0	
	Geometric Mean (CV%)	4.65 (36.82%)		4.68 (33.09%)		4.96 (29.60%)	
	Geometric 95% CI	4.29, 5.04		4.22, 5.18		4.51, 5.46	
	Median	4.55		4.70		5.00	
	Q25, Q75	3.70, 6.15		3.80, 5.60		4.20, 6.00	
	Min, Max	1.6, 9.3		2.6, 11.2		2.5, 8.8	
	Mean (SD)	4.94 (1.677)		4.93 (1.779)		5.17 (1.501)	
	95% CI						

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_cohb.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.6 Descriptive Statistics of COHb (%) - FAS**

Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)	
		Raw value	%Change (*)	Raw value	%Change (*)	Raw value	%Change (*)
Day 1	n	80	80	40	40	39	39
	BLOQ - n (%)	0		0		0	
	Geometric Mean (CV%)	1.20 (14.65%)		4.22 (35.05%)		1.21 (15.87%)	
	Geometric 95% CI	1.16, 1.25		3.78, 4.72		1.14, 1.27	
	Median	1.20	-74.75	3.90	-10.39	1.20	-76.00
	Q25, Q75	1.10, 1.30	-78.58, -69.33	3.15, 5.50	-20.84, -1.78	1.10, 1.40	-80.00, -70.96
	Min, Max	0.9, 1.9	-87.5, -37.5	2.7, 10.8	-44.9, 39.0	0.8, 1.6	-84.1, -56.0
	Mean (SD)	1.22 (0.186)	-72.99 (8.641)	4.49 (1.725)	-8.86 (17.945)	1.22 (0.190)	-74.94 (6.434)
	95% CI		-74.91, -71.06		-14.60, -3.11		-77.03, -72.85

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_cohb.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.6 Descriptive Statistics of COHb (%) - FAS**

Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)	
		Raw value	%Change (*)	Raw value	%Change (*)	Raw value	%Change (*)
Day 2	n	80	80	41	41	39	39
	BLOQ - n (%)	0		0		0	
	Geometric Mean (CV%)	1.09 (9.51%)		4.08 (31.99%)		1.01 (11.08%)	
	Geometric 95% CI	1.06, 1.12		3.69, 4.51		0.97, 1.06	
	Median	1.10	-76.34	4.10	-14.58	1.00	-79.59
	Q25, Q75	1.00, 1.20	-81.76, -71.26	3.10, 5.00	-24.40, -2.63	1.00, 1.10	-83.34, -76.19
	Min, Max	0.8, 1.4	-88.6, -37.5	2.3, 7.7	-41.0, 30.8	0.7, 1.2	-89.8, -60.0
	Mean (SD)	1.09 (0.102)	-75.18 (9.330)	4.28 (1.404)	-11.13 (17.306)	1.02 (0.109)	-78.76 (6.140)
	95% CI		-77.26, -73.10		-16.60, -5.66		-80.75, -76.76

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_cohb.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.6 Descriptive Statistics of COHb (%) - FAS**

Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)	
		Raw value	%Change (*)	Raw value	%Change (*)	Raw value	%Change (*)
Day 3	n	79	79	41	41	39	39
	BLOQ - n (%)	0		0		0	
	Geometric Mean (CV%)	1.08 (10.15%)		4.28 (31.49%)		0.99 (9.39%)	
	Geometric 95% CI	1.05, 1.11		3.88, 4.72		0.95, 1.02	
	Median	1.10	-76.47	4.30	-10.42	1.00	-79.59
	Q25, Q75	1.00, 1.20	-81.82, -71.79	3.40, 4.90	-16.67, 2.64	0.90, 1.00	-83.34, -76.19
	Min, Max	0.8, 1.3	-88.6, -42.9	2.6, 10.2	-41.9, 51.4	0.7, 1.2	-89.8, -63.3
	Mean (SD)	1.09 (0.108)	-75.62 (8.825)	4.49 (1.550)	-6.73 (18.804)	0.99 (0.090)	-79.29 (5.928)
	95% CI		-77.60, -73.64		-12.67, -0.79		-81.22, -77.37

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_cohb.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.6 Descriptive Statistics of COHb (%) - FAS**

Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)	
		Raw value	%Change (*)	Raw value	%Change (*)	Raw value	%Change (*)
Day 4	n	79	79	41	41	39	39
	BLOQ - n (%)	0		0		0	
	Geometric Mean (CV%)	1.10 (8.42%)		4.29 (31.56%)		1.03 (9.28%)	
	Geometric 95% CI	1.07, 1.13		3.89, 4.73		0.99, 1.07	
	Median	1.10	-76.19	4.20	-8.00	1.00	-79.07
	Q25, Q75	1.00, 1.20	-81.54, -72.22	3.30, 5.40	-19.41, 2.09	1.00, 1.10	-83.06, -75.00
	Min, Max	0.9, 1.3	-88.6, -37.5	2.7, 8.1	-37.8, 37.1	0.8, 1.2	-88.6, -60.0
	Mean (SD)	1.10 (0.093)	-75.17 (9.165)	4.50 (1.454)	-7.02 (15.513)	1.03 (0.096)	-78.39 (6.307)
	95% CI		-77.23, -73.12		-11.92, -2.12		-80.44, -76.34

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_cohb.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.6 Descriptive Statistics of COHb (%) - FAS**

Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)	
		Raw value	%Change (*)	Raw value	%Change (*)	Raw value	%Change (*)
Day 5, 15 min < T ₀ n		79	79	41	41		
	BLOQ - n (%)	0		0			
	Geometric Mean (CV%)	1.11 (10.62%)		2.65 (30.15%)			
	Geometric 95% CI	1.08, 1.14		2.41, 2.91			
	Median	1.10	-76.19	2.50	-44.74		
	Q25, Q75	1.00, 1.20	-81.36, -71.79	2.20, 2.80	-48.98, -36.58		
	Min, Max	0.9, 1.4	-89.2, -38.1	1.8, 5.8	-63.5, -5.7		
	Mean (SD)	1.12 (0.119)	-74.98 (9.036)	2.77 (0.936)	-42.07 (12.675)		
	95% CI		-77.01, -72.95		-46.07, -38.06		

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_cohb.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.6 Descriptive Statistics of COHb (%) - FAS**

Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)	
		Raw value	%Change (*)	Raw value	%Change (*)	Raw value	%Change (*)
Day 5, 08:00-10:00 n AM						39	39
	BLOQ - n (%)					0	
	Geometric Mean (CV%)					1.08 (9.67%)	
	Geometric 95% CI					1.04, 1.12	
	Median					1.10	-77.78
	Q25, Q75					1.00, 1.20	-82.15, -73.80
	Min, Max					0.8, 1.2	-88.6, -63.3
	Mean (SD)					1.08 (0.100)	-77.37 (6.523)
	95% CI						-79.49, -75.25

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_cohb.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.6 Descriptive Statistics of COHb (%) - FAS**

Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)	
		Raw value	%Change (*)	Raw value	%Change (*)	Raw value	%Change (*)
Day 5, 12:00-02:00 n PM		79	79	41	41	39	39
	BLOQ - n (%)	0		0		0	
	Geometric Mean (CV%)	1.11 (10.47%)		4.24 (36.43%)		1.03 (10.81%)	
	Geometric 95% CI	1.08, 1.14		3.78, 4.74		0.99, 1.07	
	Median	1.10	-76.09	4.40	-11.76	1.00	-78.43
	Q25, Q75	1.00, 1.20	-81.25, -71.79	3.20, 5.30	-19.36, 0.00	1.00, 1.10	-83.06, -75.00
	Min, Max	0.9, 1.5	-89.2, -42.9	2.4, 9.4	-40.0, 52.4	0.8, 1.3	-88.6, -64.0
	Mean (SD)	1.12 (0.118)	-75.02 (8.591)	4.51 (1.660)	-7.56 (19.283)	1.04 (0.112)	-78.36 (6.148)
	95% CI		-76.95, -73.09		-13.65, -1.47		-80.36, -76.36

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_cohb.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.6 Descriptive Statistics of COHb (%) - FAS**

Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)	
		Raw value	%Change (*)	Raw value	%Change (*)	Raw value	%Change (*)
Day 5, 04:00-06:00 n PM		79	79	41	41	39	39
	BLOQ - n (%)	0		0		0	
	Geometric Mean (CV%)	1.07 (10.38%)		4.55 (29.88%)		1.02 (9.67%)	
	Geometric 95% CI	1.04, 1.10		4.14, 4.99		0.98, 1.05	
	Median	1.10	-77.55	4.70	-5.26	1.00	-79.59
	Q25, Q75	1.00, 1.10	-81.82, -72.22	3.60, 5.40	-16.67, 6.25	1.00, 1.10	-83.34, -74.35
	Min, Max	0.8, 1.4	-89.2, -43.8	2.7, 8.9	-32.4, 51.4	0.7, 1.2	-88.6, -64.0
	Mean (SD)	1.08 (0.112)	-75.82 (8.649)	4.74 (1.451)	-0.98 (19.661)	1.02 (0.093)	-78.74 (5.867)
	95% CI		-77.76, -73.87		-7.19, 5.23		-80.64, -76.83

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_cohb.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.6 Descriptive Statistics of COHb (%) - FAS**

Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)	
		Raw value	%Change (*)	Raw value	%Change (*)	Raw value	%Change (*)
Day 5, 08:00-10:00 n PM		79	79	41	41	39	39
	BLOQ - n (%)	0		0		0	
	Geometric Mean (CV%)	1.06 (9.44%)		4.51 (34.27%)		0.99 (10.39%)	
	Geometric 95% CI	1.03, 1.08		4.05, 5.01		0.95, 1.03	
	Median	1.10	-77.78	4.80	-9.38	1.00	-79.07
	Q25, Q75	1.00, 1.10	-82.26, -72.97	3.40, 5.70	-15.39, 12.91	0.90, 1.10	-83.64, -75.60
	Min, Max	0.9, 1.4	-90.3, -42.9	2.7, 10.1	-42.9, 62.9	0.7, 1.2	-88.6, -66.7
	Mean (SD)	1.06 (0.102)	-76.20 (8.498)	4.76 (1.655)	-1.16 (22.649)	1.00 (0.099)	-79.24 (5.676)
	95% CI		-78.11, -74.29		-8.31, 5.99		-81.09, -77.40

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_cohb.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.6.1 Descriptive Statistics of COHb (%) by Sex - FAS**

Timepoint	Statistic	THS 2.2			
		Male (N=39)		Female (N=41)	
		Raw value	%Change (*)	Raw value	%Change (*)
Baseline	n	39		41	
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	5.06 (26.58%)		4.30 (43.48%)	
	Geometric 95% CI	4.64, 5.51		3.77, 4.91	
	Median	4.70		4.10	
	Q25, Q75	4.10, 6.30		3.50, 6.10	
	Min, Max	3.6, 9.3		1.6, 8.8	
	Mean (SD)	5.23 (1.446)		4.66 (1.845)	
	95% CI				

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_cohbs.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.6.1 Descriptive Statistics of COHb (%) by Sex - FAS**

Timepoint	Statistic	THS 2.2			
		Male (N=39)		Female (N=41)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 1	n	39	39	41	41
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	1.28 (14.43%)		1.13 (12.20%)	
	Geometric 95% CI	1.22, 1.35		1.09, 1.18	
	Median	1.20	-74.47	1.10	-75.56
	Q25, Q75	1.20, 1.40	-78.13, -70.27	1.00, 1.20	-78.69, -65.51
	Min, Max	1.0, 1.9	-83.9, -65.9	0.9, 1.5	-87.5, -37.5
	Mean (SD)	1.29 (0.195)	-74.25 (4.543)	1.14 (0.142)	-71.78 (11.171)
	95% CI		-75.73, -72.78		-75.31, -68.25

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_cohbs.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.6.1 Descriptive Statistics of COHb (%) by Sex - FAS**

Timepoint	Statistic	THS 2.2			
		Male (N=39)		Female (N=41)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 2	n	39	39	41	41
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	1.12 (9.15%)		1.06 (9.17%)	
	Geometric 95% CI	1.08, 1.16		1.02, 1.10	
	Median	1.10	-76.60	1.10	-75.61
	Q25, Q75	1.10, 1.20	-81.25, -73.80	1.00, 1.10	-82.26, -67.56
	Min, Max	0.9, 1.4	-87.1, -70.0	0.8, 1.2	-88.6, -37.5
	Mean (SD)	1.12 (0.102)	-77.38 (4.667)	1.06 (0.095)	-73.09 (11.917)
	95% CI		-78.90, -75.86		-76.85, -69.32

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_cohbs.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.6.1 Descriptive Statistics of COHb (%) by Sex - FAS**

Timepoint	Statistic	THS 2.2			
		Male (N=39)		Female (N=41)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 3	n	39	39	40	40
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	1.09 (9.06%)		1.07 (11.14%)	
	Geometric 95% CI	1.06, 1.13		1.03, 1.11	
	Median	1.10	-77.55	1.10	-75.64
	Q25, Q75	1.00, 1.20	-82.26, -72.97	1.00, 1.20	-81.59, -69.20
	Min, Max	0.9, 1.3	-88.2, -70.0	0.8, 1.3	-88.6, -42.9
	Mean (SD)	1.10 (0.099)	-77.83 (4.872)	1.08 (0.117)	-73.47 (11.091)
	95% CI		-79.42, -76.25		-77.02, -69.92

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_cohbs.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.6.1 Descriptive Statistics of COHb (%) by Sex - FAS**

Timepoint	Statistic	THS 2.2			
		Male (N=39)		Female (N=41)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 4	n	39	39	40	40
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	1.10 (9.54%)		1.10 (7.31%)	
	Geometric 95% CI	1.06, 1.14		1.07, 1.13	
	Median	1.10	-78.05	1.10	-75.56
	Q25, Q75	1.00, 1.20	-81.97, -72.97	1.00, 1.20	-81.15, -66.64
	Min, Max	0.9, 1.3	-88.2, -68.4	1.0, 1.2	-88.6, -37.5
	Mean (SD)	1.11 (0.106)	-77.64 (5.115)	1.10 (0.081)	-72.77 (11.422)
	95% CI		-79.31, -75.98		-76.42, -69.11

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_cohbs.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.6.1 Descriptive Statistics of COHb (%) by Sex - FAS**

Timepoint	Statistic	THS 2.2			
		Male (N=39)		Female (N=41)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 5, 15 min < T ₀	n	39	39	40	40
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	1.13 (10.86%)		1.09 (10.25%)	
	Geometric 95% CI	1.09, 1.17		1.05, 1.13	
	Median	1.10	-76.92	1.10	-75.28
	Q25, Q75	1.00, 1.20	-81.36, -72.72	1.00, 1.20	-81.50, -67.04
	Min, Max	0.9, 1.4	-89.2, -67.6	0.9, 1.3	-88.6, -38.1
	Mean (SD)	1.14 (0.123)	-77.09 (5.009)	1.10 (0.113)	-72.93 (11.408)
	95% CI		-78.71, -75.46		-76.59, -69.28

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_cohbs.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.6.1 Descriptive Statistics of COHb (%) by Sex - FAS**

Timepoint	Statistic	THS 2.2			
		Male (N=39)		Female (N=41)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 5, 12:00-02:00 PM	n	39	39	40	40
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	1.15 (10.58%)		1.08 (9.57%)	
	Geometric 95% CI	1.10, 1.19		1.04, 1.12	
	Median	1.10	-76.47	1.10	-75.58
	Q25, Q75	1.10, 1.20	-81.25, -72.50	1.00, 1.15	-80.85, -68.21
	Min, Max	0.9, 1.5	-89.2, -64.9	0.9, 1.3	-88.6, -42.9
	Mean (SD)	1.15 (0.124)	-76.73 (5.076)	1.09 (0.103)	-73.36 (10.805)
	95% CI		-78.38, -75.08		-76.82, -69.90

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_cohbs.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.6.1 Descriptive Statistics of COHb (%) by Sex - FAS**

Timepoint	Statistic	THS 2.2			
		Male (N=39)		Female (N=41)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 5, 04:00-06:00 PM	n	39	39	40	40
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	1.09 (10.71%)		1.06 (9.98%)	
	Geometric 95% CI	1.05, 1.13		1.02, 1.10	
	Median	1.10	-78.43	1.10	-75.58
	Q25, Q75	1.00, 1.10	-81.97, -73.17	1.00, 1.10	-81.75, -68.06
	Min, Max	0.8, 1.4	-89.2, -70.3	0.9, 1.3	-88.6, -43.8
	Mean (SD)	1.09 (0.117)	-77.90 (4.849)	1.06 (0.106)	-73.78 (10.868)
	95% CI		-79.48, -76.33		-77.26, -70.30

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_cohbs.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.6.1 Descriptive Statistics of COHb (%) by Sex - FAS**

Timepoint	Statistic	THS 2.2			
		Male (N=39)		Female (N=41)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 5, 08:00-10:00 PM	n	39	39	40	40
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	1.07 (10.18%)		1.05 (8.67%)	
	Geometric 95% CI	1.03, 1.11		1.01, 1.08	
	Median	1.10	-78.43	1.00	-75.58
	Q25, Q75	1.00, 1.10	-82.26, -75.00	1.00, 1.10	-82.39, -68.57
	Min, Max	0.9, 1.4	-90.3, -69.4	0.9, 1.2	-88.6, -42.9
	Mean (SD)	1.07 (0.112)	-78.34 (4.579)	1.05 (0.091)	-74.12 (10.723)
	95% CI		-79.82, -76.85		-77.55, -70.68

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_cohbs.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.6.1 Descriptive Statistics of COHb (%) by Sex - FAS**

Timepoint	Statistic	CC			
		Male (N=21)		Female (N=20)	
		Raw value	%Change (*)	Raw value	%Change (*)
Baseline	n	21		20	
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	5.04 (39.43%)		4.32 (23.53%)	
	Geometric 95% CI	4.24, 6.00		3.87, 4.82	
	Median	4.90		4.15	
	Q25, Q75	3.90, 6.20		3.70, 4.80	
	Min, Max	2.6, 11.2		3.1, 6.8	
	Mean (SD)	5.41 (2.177)		4.44 (1.079)	
	95% CI				

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_cohbs.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.3.6.1 Descriptive Statistics of COHb (%) by Sex - FAS**

Timepoint	Statistic	CC			
		Male (N=21)		Female (N=20)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 1	n	21	21	19	19
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	4.75 (40.21%)		3.71 (22.89%)	
	Geometric 95% CI	3.98, 5.67		3.33, 4.15	
	Median	5.20	-7.69	3.80	-16.67
	Q25, Q75	3.50, 5.90	-18.19, 13.47	2.90, 4.60	-20.84, -6.45
	Min, Max	2.7, 10.8	-44.9, 39.0	2.7, 5.6	-32.5, 14.3
	Mean (SD)	5.10 (2.067)	-3.58 (21.182)	3.81 (0.876)	-14.69 (11.435)
	95% CI		-13.23, 6.07		-20.21, -9.18

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_cohbs.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.3.6.1 Descriptive Statistics of COHb (%) by Sex - FAS**

Timepoint	Statistic	CC			
		Male (N=21)		Female (N=20)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 2	n	21	21	20	20
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	4.40 (35.63%)		3.76 (26.14%)	
	Geometric 95% CI	3.76, 5.16		3.33, 4.25	
	Median	4.40	-15.00	3.65	-11.00
	Q25, Q75	3.70, 5.50	-26.42, -1.78	3.05, 4.45	-22.73, -3.17
	Min, Max	2.3, 7.7	-41.0, 30.8	2.6, 7.7	-35.0, 16.1
	Mean (SD)	4.65 (1.547)	-10.70 (19.324)	3.90 (1.149)	-11.58 (15.397)
	95% CI		-19.50, -1.90		-18.79, -4.37

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_cohbs.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.3.6.1 Descriptive Statistics of COHb (%) by Sex - FAS**

Timepoint	Statistic	CC			
		Male (N=21)		Female (N=20)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 3	n	21	21	20	20
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	4.59 (38.08%)		3.97 (21.45%)	
	Geometric 95% CI	3.88, 5.44		3.59, 4.39	
	Median	4.50	-8.93	4.00	-10.53
	Q25, Q75	3.30, 5.60	-21.22, 7.15	3.40, 4.60	-16.15, 2.36
	Min, Max	2.6, 10.2	-41.9, 51.4	2.7, 6.1	-34.0, 29.0
	Mean (SD)	4.91 (1.931)	-6.52 (21.990)	4.06 (0.858)	-6.96 (15.340)
	95% CI		-16.53, 3.50		-14.14, 0.22

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_cohbs.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.3.6.1 Descriptive Statistics of COHb (%) by Sex - FAS**

Timepoint	Statistic	CC			
		Male (N=21)		Female (N=20)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 4	n	21	21	20	20
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	4.62 (35.30%)		3.97 (25.65%)	
	Geometric 95% CI	3.95, 5.41		3.52, 4.47	
	Median	4.60	-8.00	3.75	-7.54
	Q25, Q75	3.60, 6.00	-20.76, 6.46	3.15, 4.85	-16.85, 1.05
	Min, Max	2.8, 8.1	-37.8, 37.1	2.7, 6.2	-26.8, 14.3
	Mean (SD)	4.89 (1.684)	-6.67 (18.286)	4.09 (1.060)	-7.39 (12.423)
	95% CI		-15.00, 1.66		-13.21, -1.57

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_cohbs.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.3.6.1 Descriptive Statistics of COHb (%) by Sex - FAS**

Timepoint	Statistic	CC			
		Male (N=21)		Female (N=20)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 5, 15 min < T ₀	n	21	21	20	20
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	3.06 (34.37%)		2.27 (13.20%)	
	Geometric 95% CI	2.63, 3.57		2.13, 2.42	
	Median	2.80	-41.67	2.30	-46.09
	Q25, Q75	2.50, 4.10	-47.17, -29.03	2.05, 2.50	-52.09, -41.28
	Min, Max	1.8, 5.8	-63.5, -5.7	1.8, 2.7	-63.2, -22.6
	Mean (SD)	3.23 (1.101)	-37.80 (13.718)	2.29 (0.294)	-46.55 (9.941)
	95% CI		-44.05, -31.55		-51.21, -41.89

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_cohbs.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.6.1 Descriptive Statistics of COHb (%) by Sex - FAS**

Timepoint	Statistic	CC			
		Male (N=21)		Female (N=20)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 5, 12:00-02:00 PM	n	21	21	20	20
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	4.59 (40.09%)		3.89 (30.82%)	
	Geometric 95% CI	3.85, 5.48		3.38, 4.49	
	Median	4.50	-14.63	3.70	-10.27
	Q25, Q75	3.50, 6.50	-19.36, 6.39	3.05, 5.20	-19.38, -1.21
	Min, Max	2.5, 9.4	-33.9, 33.3	2.4, 6.4	-40.0, 52.4
	Mean (SD)	4.93 (1.915)	-6.99 (19.708)	4.07 (1.240)	-8.16 (19.320)
	95% CI		-15.97, 1.99		-17.20, 0.89

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_cohbs.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.6.1 Descriptive Statistics of COHb (%) by Sex - FAS**

Timepoint	Statistic	CC			
		Male (N=21)		Female (N=20)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 5, 04:00-06:00 PM	n	21	21	20	20
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	4.92 (33.07%)		4.19 (24.23%)	
	Geometric 95% CI	4.24, 5.70		3.74, 4.69	
	Median	5.00	-6.45	3.95	-3.67
	Q25, Q75	4.30, 6.00	-10.26, 4.88	3.50, 5.10	-18.94, 7.52
	Min, Max	2.7, 8.9	-32.4, 51.4	2.8, 6.3	-29.8, 38.1
	Mean (SD)	5.16 (1.667)	-0.49 (21.396)	4.31 (1.055)	-1.49 (18.204)
	95% CI		-10.23, 9.26		-10.01, 7.03

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_cohbs.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.6.1 Descriptive Statistics of COHb (%) by Sex - FAS**

Timepoint	Statistic	CC			
		Male (N=21)		Female (N=20)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 5, 08:00-10:00 PM	n	21	21	20	20
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	4.97 (34.55%)		4.07 (31.32%)	
	Geometric 95% CI	4.26, 5.79		3.52, 4.70	
	Median	5.10	-5.13	3.80	-10.94
	Q25, Q75	4.10, 6.10	-15.00, 19.24	3.15, 5.30	-21.09, 11.66
	Min, Max	2.8, 10.1	-42.9, 62.9	2.7, 7.2	-33.3, 38.1
	Mean (SD)	5.24 (1.795)	1.37 (24.851)	4.26 (1.361)	-3.82 (20.382)
	95% CI		-9.95, 12.68		-13.36, 5.73

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_cohbs.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.3.6.1 Descriptive Statistics of COHb (%) by Sex - FAS**

Timepoint	Statistic	SA			
		Male (N=20)		Female (N=19)	
		Raw value	%Change (*)	Raw value	%Change (*)
Baseline	n	20		19	
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	5.31 (31.78%)		4.62 (25.98%)	
	Geometric 95% CI	4.59, 6.15		4.08, 5.23	
	Median	5.30		4.40	
	Q25, Q75	4.70, 6.55		3.90, 5.50	
	Min, Max	2.5, 8.8		3.1, 8.1	
	Mean (SD)	5.55 (1.606)		4.77 (1.307)	
	95% CI				

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_cohbs.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.3.6.1 Descriptive Statistics of COHb (%) by Sex - FAS**

Timepoint	Statistic	SA			
		Male (N=20)		Female (N=19)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 1	n	20	20	19	19
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	1.28 (11.52%)		1.14 (17.78%)	
	Geometric 95% CI	1.21, 1.35		1.04, 1.24	
	Median	1.30	-76.14	1.20	-75.76
	Q25, Q75	1.15, 1.40	-82.36, -71.54	1.00, 1.20	-77.78, -70.96
	Min, Max	1.1, 1.5	-84.1, -56.0	0.8, 1.6	-83.9, -65.7
	Mean (SD)	1.29 (0.147)	-74.93 (7.723)	1.15 (0.210)	-74.95 (4.943)
	95% CI		-78.54, -71.31		-77.34, -72.56

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_cohbs.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.3.6.1 Descriptive Statistics of COHb (%) by Sex - FAS**

Timepoint	Statistic	SA			
		Male (N=20)		Female (N=19)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 2	n	20	20	19	19
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	1.04 (7.18%)		0.99 (13.72%)	
	Geometric 95% CI	1.00, 1.08		0.92, 1.06	
	Median	1.00	-80.58	1.00	-78.18
	Q25, Q75	1.00, 1.10	-83.98, -76.38	0.90, 1.10	-81.40, -76.19
	Min, Max	0.9, 1.2	-89.8, -60.0	0.7, 1.2	-85.5, -70.7
	Mean (SD)	1.05 (0.076)	-79.29 (7.366)	0.99 (0.132)	-78.19 (4.655)
	95% CI		-82.75, -75.84		-80.44, -75.94

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_cohbs.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.6.1 Descriptive Statistics of COHb (%) by Sex - FAS**

Timepoint	Statistic	SA			
		Male (N=20)		Female (N=19)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 3	n	20	20	19	19
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	1.00 (8.13%)		0.97 (10.60%)	
	Geometric 95% CI	0.96, 1.05		0.92, 1.03	
	Median	1.00	-81.42	1.00	-79.07
	Q25, Q75	0.95, 1.05	-84.72, -77.37	0.90, 1.00	-80.77, -75.60
	Min, Max	0.9, 1.2	-89.8, -63.3	0.7, 1.2	-86.1, -71.0
	Mean (SD)	1.01 (0.083)	-80.07 (7.059)	0.98 (0.098)	-78.47 (4.495)
	95% CI		-83.38, -76.76		-80.64, -76.30

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_cohbs.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.6.1 Descriptive Statistics of COHb (%) by Sex - FAS**

Timepoint	Statistic	SA			
		Male (N=20)		Female (N=19)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 4	n	20	20	19	19
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	1.04 (9.94%)		1.02 (8.64%)	
	Geometric 95% CI	0.99, 1.09		0.97, 1.07	
	Median	1.00	-79.97	1.00	-77.78
	Q25, Q75	1.00, 1.15	-84.57, -76.23	1.00, 1.10	-80.00, -73.80
	Min, Max	0.9, 1.2	-88.6, -60.0	0.8, 1.2	-86.1, -67.7
	Mean (SD)	1.05 (0.106)	-79.30 (7.290)	1.02 (0.086)	-77.43 (5.098)
	95% CI		-82.72, -75.89		-79.90, -74.97

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_cohbs.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.6.1 Descriptive Statistics of COHb (%) by Sex - FAS**

Timepoint	Statistic	SA			
		Male (N=20)		Female (N=19)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 5, 08:00-10:00 AM	n	20	20	19	19
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	1.09 (9.63%)		1.07 (9.92%)	
	Geometric 95% CI	1.03, 1.14		1.01, 1.13	
	Median	1.10	-79.22	1.10	-75.76
	Q25, Q75	1.00, 1.20	-83.81, -75.23	1.00, 1.10	-80.00, -73.17
	Min, Max	0.9, 1.2	-88.6, -63.3	0.8, 1.2	-86.1, -64.5
	Mean (SD)	1.09 (0.103)	-78.54 (6.943)	1.07 (0.100)	-76.15 (5.987)
	95% CI		-81.79, -75.29		-79.04, -73.26

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_cohbs.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.6.1 Descriptive Statistics of COHb (%) by Sex - FAS**

Timepoint	Statistic	SA			
		Male (N=20)		Female (N=19)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 5, 12:00-02:00 PM	n	20	20	19	19
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	1.05 (9.29%)		1.01 (12.20%)	
	Geometric 95% CI	1.00, 1.10		0.95, 1.08	
	Median	1.00	-80.56	1.00	-77.27
	Q25, Q75	1.00, 1.10	-83.58, -75.79	1.00, 1.10	-81.40, -74.35
	Min, Max	0.9, 1.3	-88.6, -64.0	0.8, 1.3	-87.5, -64.5
	Mean (SD)	1.06 (0.100)	-79.25 (6.686)	1.02 (0.123)	-77.42 (5.549)
	95% CI		-82.39, -76.12		-80.10, -74.74

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_cohbs.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.3.6.1 Descriptive Statistics of COHb (%) by Sex - FAS**

Timepoint	Statistic	SA			
		Male (N=20)		Female (N=19)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 5, 04:00-06:00 PM	n	20	20	19	19
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	1.03 (7.41%)		1.00 (11.57%)	
	Geometric 95% CI	0.99, 1.07		0.94, 1.06	
	Median	1.05	-80.18	1.00	-78.79
	Q25, Q75	1.00, 1.10	-83.98, -76.59	1.00, 1.10	-80.77, -73.17
	Min, Max	0.9, 1.1	-88.6, -64.0	0.7, 1.2	-87.5, -71.0
	Mean (SD)	1.04 (0.075)	-79.64 (6.548)	1.01 (0.108)	-77.79 (5.055)
	95% CI		-82.71, -76.57		-80.23, -75.34

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_cohbs.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.6.1 Descriptive Statistics of COHb (%) by Sex - FAS**

Timepoint	Statistic	SA			
		Male (N=20)		Female (N=19)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 5, 08:00-10:00 PM	n	20	20	19	19
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	1.01 (9.44%)		0.97 (11.23%)	
	Geometric 95% CI	0.96, 1.06		0.92, 1.03	
	Median	1.00	-81.42	1.00	-78.79
	Q25, Q75	1.00, 1.10	-83.98, -75.79	0.90, 1.10	-82.00, -75.60
	Min, Max	0.8, 1.2	-88.6, -66.7	0.7, 1.1	-87.5, -67.7
	Mean (SD)	1.02 (0.094)	-80.09 (6.202)	0.98 (0.104)	-78.36 (5.078)
	95% CI		-82.99, -77.18		-80.81, -75.90

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_cohbs.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.6.2 Descriptive Statistics of COHb (%) by Cigarette Consumption - FAS**

Timepoint	Statistic	THS 2.2			
		10-19 cig/day (N=41)		>19 cig/day (N=39)	
		Raw value	%Change (*)	Raw value	%Change (*)
Baseline	n	41		39	
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	4.20 (36.50%)		5.18 (34.00%)	
	Geometric 95% CI	3.75, 4.70		4.65, 5.77	
	Median	4.10		5.70	
	Q25, Q75	3.60, 5.10		4.00, 6.60	
	Min, Max	1.6, 8.8		2.4, 9.3	
	Mean (SD)	4.45 (1.533)		5.45 (1.689)	
	95% CI				

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_cohbc.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.6.2 Descriptive Statistics of COHb (%) by Cigarette Consumption - FAS**

Timepoint	Statistic	THS 2.2			
		10-19 cig/day (N=41)		>19 cig/day (N=39)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 1	n	41	41	39	39
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	1.15 (12.17%)		1.26 (15.50%)	
	Geometric 95% CI	1.10, 1.20		1.20, 1.33	
	Median	1.20	-73.33	1.20	-76.60
	Q25, Q75	1.00, 1.20	-77.56, -69.23	1.10, 1.40	-79.67, -69.44
	Min, Max	0.9, 1.5	-87.5, -37.5	1.0, 1.9	-87.3, -54.2
	Mean (SD)	1.16 (0.142)	-71.37 (9.724)	1.28 (0.206)	-74.68 (7.065)
	95% CI		-74.45, -68.30		-76.98, -72.39

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_cohbc.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.6.2 Descriptive Statistics of COHb (%) by Cigarette Consumption - FAS**

Timepoint	Statistic	THS 2.2			
		10-19 cig/day (N=41)		>19 cig/day (N=39)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 2	n	41	41	39	39
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	1.07 (8.25%)		1.11 (10.41%)	
	Geometric 95% CI	1.03, 1.10		1.07, 1.15	
	Median	1.10	-75.00	1.10	-79.10
	Q25, Q75	1.00, 1.10	-78.44, -70.73	1.10, 1.20	-83.10, -72.22
	Min, Max	0.9, 1.2	-88.6, -37.5	0.8, 1.4	-88.6, -57.1
	Mean (SD)	1.07 (0.088)	-73.01 (10.427)	1.12 (0.112)	-77.46 (7.490)
	95% CI		-76.30, -69.71		-79.90, -75.03

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_cohbc.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.6.2 Descriptive Statistics of COHb (%) by Cigarette Consumption - FAS**

Timepoint	Statistic	THS 2.2			
		10-19 cig/day (N=41)		>19 cig/day (N=39)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 3	n	40	40	39	39
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	1.06 (9.76%)		1.10 (10.33%)	
	Geometric 95% CI	1.03, 1.10		1.06, 1.14	
	Median	1.10	-75.64	1.10	-79.59
	Q25, Q75	1.00, 1.15	-79.43, -71.61	1.00, 1.20	-83.10, -72.22
	Min, Max	0.9, 1.2	-87.5, -42.9	0.8, 1.3	-88.6, -54.2
	Mean (SD)	1.07 (0.103)	-73.74 (9.406)	1.11 (0.111)	-77.55 (7.845)
	95% CI		-76.76, -70.73		-80.10, -75.00

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_cohbc.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.6.2 Descriptive Statistics of COHb (%) by Cigarette Consumption - FAS**

Timepoint	Statistic	THS 2.2			
		10-19 cig/day (N=41)		>19 cig/day (N=39)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 4	n	40	40	39	39
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	1.07 (7.94%)		1.14 (7.79%)	
	Geometric 95% CI	1.04, 1.10		1.10, 1.17	
	Median	1.10	-74.96	1.10	-79.66
	Q25, Q75	1.00, 1.10	-79.61, -71.12	1.10, 1.20	-82.26, -72.72
	Min, Max	0.9, 1.3	-88.6, -37.5	1.0, 1.3	-88.2, -54.2
	Mean (SD)	1.07 (0.086)	-73.39 (10.292)	1.14 (0.088)	-77.01 (7.543)
	95% CI		-76.68, -70.09		-79.46, -74.56

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_cohbc.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.6.2 Descriptive Statistics of COHb (%) by Cigarette Consumption - FAS**

Timepoint	Statistic	THS 2.2			
		10-19 cig/day (N=41)		>19 cig/day (N=39)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 5, 15 min < T ₀	n	40	40	39	39
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	1.08 (10.14%)		1.14 (10.58%)	
	Geometric 95% CI	1.04, 1.12		1.10, 1.18	
	Median	1.10	-73.57	1.10	-79.10
	Q25, Q75	1.00, 1.20	-80.86, -70.73	1.10, 1.20	-81.97, -72.50
	Min, Max	0.9, 1.3	-88.6, -38.1	0.9, 1.4	-89.2, -58.3
	Mean (SD)	1.09 (0.110)	-73.07 (10.104)	1.15 (0.122)	-76.94 (7.420)
	95% CI		-76.31, -69.84		-79.35, -74.53

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_cohbc.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.6.2 Descriptive Statistics of COHb (%) by Cigarette Consumption - FAS**

Timepoint	Statistic	THS 2.2			
		10-19 cig/day (N=41)		>19 cig/day (N=39)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 5, 12:00-02:00 PM	n	40	40	39	39
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	1.08 (9.86%)		1.15 (10.13%)	
	Geometric 95% CI	1.04, 1.12		1.11, 1.19	
	Median	1.10	-73.57	1.10	-78.85
	Q25, Q75	1.00, 1.20	-80.00, -71.42	1.10, 1.20	-81.70, -71.79
	Min, Max	0.9, 1.3	-88.6, -42.9	0.9, 1.5	-89.2, -54.2
	Mean (SD)	1.08 (0.106)	-73.33 (9.493)	1.16 (0.119)	-76.76 (7.273)
	95% CI		-76.37, -70.29		-79.12, -74.40

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_cohbc.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.6.2 Descriptive Statistics of COHb (%) by Cigarette Consumption - FAS**

Timepoint	Statistic	THS 2.2			
		10-19 cig/day (N=41)		>19 cig/day (N=39)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 5, 04:00-06:00 PM	n	40	40	39	39
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	1.04 (9.81%)		1.11 (9.75%)	
	Geometric 95% CI	1.00, 1.07		1.07, 1.15	
	Median	1.00	-74.14	1.10	-79.10
	Q25, Q75	1.00, 1.10	-81.11, -72.22	1.10, 1.20	-82.82, -72.50
	Min, Max	0.8, 1.3	-88.6, -43.8	0.9, 1.4	-89.2, -54.2
	Mean (SD)	1.04 (0.101)	-74.34 (9.269)	1.12 (0.110)	-77.34 (7.790)
	95% CI		-77.31, -71.37		-79.87, -74.81

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_cohbc.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.6.2 Descriptive Statistics of COHb (%) by Cigarette Consumption - FAS**

Timepoint	Statistic	THS 2.2			
		10-19 cig/day (N=41)		>19 cig/day (N=39)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 5, 08:00-10:00 PM	n	40	40	39	39
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	1.03 (7.80%)		1.08 (10.50%)	
	Geometric 95% CI	1.00, 1.07		1.04, 1.12	
	Median	1.00	-75.28	1.10	-80.65
	Q25, Q75	1.00, 1.10	-79.46, -72.36	1.00, 1.20	-82.82, -74.35
	Min, Max	0.9, 1.2	-88.6, -42.9	0.9, 1.4	-90.3, -58.3
	Mean (SD)	1.04 (0.081)	-74.36 (9.349)	1.09 (0.116)	-78.09 (7.163)
	95% CI		-77.35, -71.36		-80.41, -75.76

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_cohbc.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.6.2 Descriptive Statistics of COHb (%) by Cigarette Consumption - FAS**

Timepoint	Statistic	CC			
		10-19 cig/day (N=21)		>19 cig/day (N=20)	
		Raw value	%Change (*)	Raw value	%Change (*)
Baseline	n	21		20	
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	4.47 (26.00%)		4.90 (39.56%)	
	Geometric 95% CI	3.97, 5.03		4.09, 5.86	
	Median	4.20		4.75	
	Q25, Q75	3.50, 5.40		3.80, 6.50	
	Min, Max	3.1, 7.5		2.6, 11.2	
	Mean (SD)	4.61 (1.220)		5.27 (2.206)	
	95% CI				

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_cohbc.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.3.6.2 Descriptive Statistics of COHb (%) by Cigarette Consumption - FAS**

Timepoint	Statistic	CC			
		10-19 cig/day (N=21)		>19 cig/day (N=20)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 1	n	20	20	20	20
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	4.18 (30.19%)		4.27 (40.28%)	
	Geometric 95% CI	3.64, 4.81		3.55, 5.12	
	Median	4.55	-10.26	3.80	-10.39
	Q25, Q75	3.10, 5.30	-20.72, 12.50	3.15, 5.60	-20.84, -5.01
	Min, Max	2.7, 6.9	-44.9, 39.0	2.7, 10.8	-32.5, 19.2
	Mean (SD)	4.36 (1.250)	-5.80 (21.206)	4.62 (2.124)	-11.91 (13.843)
	95% CI		-15.73, 4.13		-18.40, -5.43

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_cohbc.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.6.2 Descriptive Statistics of COHb (%) by Cigarette Consumption - FAS**

Timepoint	Statistic	CC			
		10-19 cig/day (N=21)		>19 cig/day (N=20)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 2	n	21	21	20	20
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	4.02 (23.95%)		4.14 (39.66%)	
	Geometric 95% CI	3.61, 4.49		3.46, 4.95	
	Median	4.30	-6.38	3.80	-16.03
	Q25, Q75	3.60, 4.50	-24.40, 2.39	3.00, 5.55	-24.04, -4.71
	Min, Max	2.3, 6.1	-36.7, 25.7	2.3, 7.7	-41.0, 30.8
	Mean (SD)	4.13 (0.928)	-8.32 (17.725)	4.45 (1.785)	-14.08 (16.791)
	95% CI		-16.39, -0.25		-21.95, -6.22

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_cohbc.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.6.2 Descriptive Statistics of COHb (%) by Cigarette Consumption - FAS**

Timepoint	Statistic	CC			
		10-19 cig/day (N=21)		>19 cig/day (N=20)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 3	n	21	21	20	20
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	4.20 (24.86%)		4.36 (38.04%)	
	Geometric 95% CI	3.75, 4.70		3.67, 5.19	
	Median	4.50	-10.53	4.15	-9.99
	Q25, Q75	3.60, 4.90	-15.63, 7.15	3.20, 5.65	-16.67, -1.43
	Min, Max	2.6, 6.6	-38.8, 51.4	2.8, 10.2	-41.9, 22.6
	Mean (SD)	4.31 (0.993)	-3.95 (21.547)	4.68 (1.986)	-9.65 (15.434)
	95% CI		-13.77, 5.86		-16.88, -2.42

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_cohbc.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.6.2 Descriptive Statistics of COHb (%) by Cigarette Consumption - FAS**

Timepoint	Statistic	CC			
		10-19 cig/day (N=21)		>19 cig/day (N=20)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 4	n	21	21	20	20
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	4.22 (27.85%)		4.37 (35.81%)	
	Geometric 95% CI	3.72, 4.78		3.71, 5.14	
	Median	4.60	-6.25	3.95	-9.41
	Q25, Q75	3.30, 4.80	-15.16, 3.51	3.40, 6.00	-23.14, 0.41
	Min, Max	2.8, 6.9	-26.8, 37.1	2.7, 8.1	-37.8, 15.4
	Mean (SD)	4.37 (1.173)	-4.47 (15.952)	4.64 (1.721)	-9.70 (14.967)
	95% CI		-11.74, 2.79		-16.71, -2.69

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_cohbc.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.6.2 Descriptive Statistics of COHb (%) by Cigarette Consumption - FAS**

Timepoint	Statistic	CC			
		10-19 cig/day (N=21)		>19 cig/day (N=20)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 5, 15 min < T ₀	n	21	21	20	20
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	2.58 (23.65%)		2.72 (36.41%)	
	Geometric 95% CI	2.32, 2.88		2.30, 3.21	
	Median	2.50	-42.42	2.40	-45.77
	Q25, Q75	2.30, 2.80	-47.17, -36.58	2.10, 3.45	-51.87, -33.10
	Min, Max	1.8, 4.1	-62.7, -5.7	1.8, 5.8	-63.5, -15.4
	Mean (SD)	2.65 (0.635)	-41.08 (12.211)	2.90 (1.177)	-43.10 (13.381)
	95% CI		-46.65, -35.52		-49.37, -36.84

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_cohbc.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.6.2 Descriptive Statistics of COHb (%) by Cigarette Consumption - FAS**

Timepoint	Statistic	CC			
		10-19 cig/day (N=21)		>19 cig/day (N=20)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 5, 12:00-02:00 PM	n	21	21	20	20
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	4.20 (28.70%)		4.27 (44.26%)	
	Geometric 95% CI	3.69, 4.78		3.50, 5.21	
	Median	4.40	-8.77	4.15	-12.83
	Q25, Q75	3.30, 5.00	-15.79, 6.39	2.95, 6.25	-22.84, -3.84
	Min, Max	2.8, 7.5	-33.9, 52.4	2.4, 9.4	-40.0, 25.0
	Mean (SD)	4.37 (1.252)	-3.83 (21.612)	4.66 (2.026)	-11.48 (16.113)
	95% CI		-13.67, 6.01		-19.02, -3.93

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_cohbc.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.6.2 Descriptive Statistics of COHb (%) by Cigarette Consumption - FAS**

Timepoint	Statistic	CC			
		10-19 cig/day (N=21)		>19 cig/day (N=20)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 5, 04:00-06:00 PM	n	21	21	20	20
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	4.52 (26.31%)		4.57 (34.03%)	
	Geometric 95% CI	4.02, 5.09		3.91, 5.34	
	Median	4.80	0.00	4.60	-5.90
	Q25, Q75	3.90, 5.30	-16.67, 16.67	3.45, 5.75	-14.69, 0.81
	Min, Max	2.7, 7.3	-26.5, 51.4	2.9, 8.9	-32.4, 25.8
	Mean (SD)	4.67 (1.173)	3.36 (22.378)	4.83 (1.724)	-5.53 (15.621)
	95% CI		-6.83, 13.55		-12.85, 1.78

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_cohbc.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.6.2 Descriptive Statistics of COHb (%) by Cigarette Consumption - FAS**

Timepoint	Statistic	CC			
		10-19 cig/day (N=21)		>19 cig/day (N=20)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 5, 08:00-10:00 PM	n	21	21	20	20
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	4.34 (32.81%)		4.70 (36.14%)	
	Geometric 95% CI	3.74, 5.02		3.98, 5.54	
	Median	4.80	-9.43	4.70	-3.37
	Q25, Q75	3.20, 5.30	-15.16, 24.08	3.45, 6.25	-16.67, 10.05
	Min, Max	2.7, 7.8	-42.9, 62.9	2.9, 10.1	-29.7, 25.0
	Mean (SD)	4.55 (1.419)	0.49 (27.797)	4.99 (1.882)	-2.90 (16.139)
	95% CI		-12.16, 13.15		-10.46, 4.66

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_cohbc.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.6.2 Descriptive Statistics of COHb (%) by Cigarette Consumption - FAS**

Timepoint	Statistic	SA			
		10-19 cig/day (N=19)		>19 cig/day (N=20)	
		Raw value	%Change (*)	Raw value	%Change (*)
Baseline	n	19		20	
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	4.59 (24.14%)		5.34 (32.88%)	
	Geometric 95% CI	4.09, 5.16		4.59, 6.21	
	Median	4.70		5.30	
	Q25, Q75	3.90, 5.50		4.30, 7.00	
	Min, Max	3.0, 6.8		2.5, 8.8	
	Mean (SD)	4.72 (1.084)		5.60 (1.729)	
	95% CI				

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_cohbc.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.3.6.2 Descriptive Statistics of COHb (%) by Cigarette Consumption - FAS**

Timepoint	Statistic	SA			
		10-19 cig/day (N=19)		>19 cig/day (N=20)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 1	n	19	19	20	20
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	1.18 (18.77%)		1.24 (12.56%)	
	Geometric 95% CI	1.07, 1.29		1.16, 1.32	
	Median	1.20	-75.76	1.20	-76.37
	Q25, Q75	1.00, 1.40	-76.75, -70.90	1.15, 1.40	-82.45, -71.42
	Min, Max	0.8, 1.6	-83.9, -60.0	1.0, 1.6	-84.1, -56.0
	Mean (SD)	1.19 (0.220)	-73.85 (5.605)	1.25 (0.158)	-75.97 (7.123)
	95% CI		-76.56, -71.15		-79.31, -72.63

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_cohbc.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.6.2 Descriptive Statistics of COHb (%) by Cigarette Consumption - FAS**

Timepoint	Statistic	SA			
		10-19 cig/day (N=19)		>19 cig/day (N=20)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 2	n	19	19	20	20
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	0.99 (12.17%)		1.04 (9.81%)	
	Geometric 95% CI	0.93, 1.06		0.98, 1.09	
	Median	1.00	-78.79	1.00	-81.08
	Q25, Q75	0.90, 1.10	-80.77, -75.00	1.00, 1.10	-84.96, -76.73
	Min, Max	0.7, 1.2	-85.5, -63.3	0.8, 1.2	-89.8, -60.0
	Mean (SD)	1.00 (0.116)	-77.86 (5.198)	1.04 (0.100)	-79.61 (6.945)
	95% CI		-80.37, -75.35		-82.86, -76.35

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_cohbc.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.3.6.2 Descriptive Statistics of COHb (%) by Cigarette Consumption - FAS**

Timepoint	Statistic	SA			
		10-19 cig/day (N=19)		>19 cig/day (N=20)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 3	n	19	19	20	20
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	0.99 (11.56%)		0.99 (7.07%)	
	Geometric 95% CI	0.93, 1.05		0.95, 1.03	
	Median	1.00	-78.79	1.00	-81.13
	Q25, Q75	0.90, 1.10	-81.36, -74.46	0.95, 1.00	-85.65, -77.00
	Min, Max	0.7, 1.2	-85.5, -63.3	0.9, 1.2	-89.8, -64.0
	Mean (SD)	0.99 (0.108)	-77.91 (5.389)	0.99 (0.072)	-80.61 (6.247)
	95% CI		-80.51, -75.30		-83.54, -77.68

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_cohbc.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.6.2 Descriptive Statistics of COHb (%) by Cigarette Consumption - FAS**

Timepoint	Statistic	SA			
		10-19 cig/day (N=19)		>19 cig/day (N=20)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 4	n	19	19	20	20
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	1.01 (8.93%)		1.05 (9.54%)	
	Geometric 95% CI	0.96, 1.06		0.99, 1.10	
	Median	1.00	-77.78	1.00	-79.80
	Q25, Q75	1.00, 1.00	-80.77, -75.00	1.00, 1.10	-85.65, -74.90
	Min, Max	0.8, 1.2	-85.0, -66.7	0.9, 1.2	-88.6, -60.0
	Mean (SD)	1.02 (0.090)	-77.47 (5.008)	1.05 (0.100)	-79.27 (7.358)
	95% CI		-79.89, -75.05		-82.72, -75.82

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_cohbc.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.6.2 Descriptive Statistics of COHb (%) by Cigarette Consumption - FAS**

Timepoint	Statistic	SA			
		10-19 cig/day (N=19)		>19 cig/day (N=20)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 5, 08:00-10:00 AM	n	19	19	20	20
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	1.09 (9.11%)		1.06 (10.28%)	
	Geometric 95% CI	1.04, 1.14		1.01, 1.12	
	Median	1.10	-76.00	1.10	-78.96
	Q25, Q75	1.10, 1.10	-79.67, -72.50	1.00, 1.20	-84.66, -74.40
	Min, Max	0.8, 1.2	-83.9, -63.3	0.9, 1.2	-88.6, -64.0
	Mean (SD)	1.09 (0.092)	-75.67 (5.659)	1.07 (0.109)	-78.99 (7.008)
	95% CI		-78.40, -72.94		-82.28, -75.71

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_cohbc.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.6.2 Descriptive Statistics of COHb (%) by Cigarette Consumption - FAS**

Timepoint	Statistic	SA			
		10-19 cig/day (N=19)		>19 cig/day (N=20)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 5, 12:00-02:00 PM	n	19	19	20	20
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	1.02 (10.07%)		1.05 (11.52%)	
	Geometric 95% CI	0.96, 1.07		0.99, 1.11	
	Median	1.00	-78.43	1.05	-79.18
	Q25, Q75	1.00, 1.10	-81.64, -74.35	1.00, 1.10	-84.63, -76.55
	Min, Max	0.8, 1.3	-83.9, -64.5	0.8, 1.3	-88.6, -64.0
	Mean (SD)	1.02 (0.104)	-77.28 (5.587)	1.06 (0.120)	-79.39 (6.613)
	95% CI		-79.98, -74.58		-82.49, -76.29

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_cohbc.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.3.6.2 Descriptive Statistics of COHb (%) by Cigarette Consumption - FAS**

Timepoint	Statistic	SA			
		10-19 cig/day (N=19)		>19 cig/day (N=20)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 5, 04:00-06:00 PM	n	19	19	20	20
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	1.01 (11.10%)		1.02 (8.36%)	
	Geometric 95% CI	0.95, 1.07		0.98, 1.07	
	Median	1.00	-78.43	1.00	-80.18
	Q25, Q75	1.00, 1.10	-81.64, -74.35	1.00, 1.10	-85.86, -74.11
	Min, Max	0.7, 1.1	-83.9, -66.7	0.9, 1.2	-88.6, -64.0
	Mean (SD)	1.02 (0.102)	-77.53 (4.844)	1.03 (0.086)	-79.88 (6.617)
	95% CI		-79.87, -75.19		-82.98, -76.78

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_cohbc.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.3.6.2 Descriptive Statistics of COHb (%) by Cigarette Consumption - FAS**

Timepoint	Statistic	SA			
		10-19 cig/day (N=19)		>19 cig/day (N=20)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 5, 08:00-10:00 PM	n	19	19	20	20
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	0.98 (11.76%)		1.00 (9.14%)	
	Geometric 95% CI	0.92, 1.05		0.95, 1.05	
	Median	1.00	-78.79	1.00	-79.53
	Q25, Q75	0.90, 1.10	-82.00, -75.00	0.95, 1.10	-86.60, -75.90
	Min, Max	0.7, 1.2	-83.9, -66.7	0.8, 1.1	-88.6, -68.0
	Mean (SD)	0.99 (0.111)	-78.06 (5.189)	1.01 (0.089)	-80.36 (6.017)
	95% CI		-80.57, -75.56		-83.18, -77.54

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_cohbc.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.7 Descriptive Statistics of MHBMA Urinary Concentration Adjusted for Creatinine (pg/mg creat) - FAS**

Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
		Raw value	%Change (*)	Raw value	%Change (*)
Baseline	n	80		41	
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	1888.27 (113.08%)		2317.31 (78.67%)	
	Geometric 95% CI	1542.95, 2310.86		1861.41, 2884.85	
	Median	2310.34		2500.00	
	Q25, Q75	1159.37, 3586.02		1761.90, 3758.63	
	Min, Max	231.9, 7218.5		311.7, 7533.3	
	Mean (SD)	2592.45 (1780.578)		2823.91 (1672.323)	
	95% CI				

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.100 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_mhbma.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.7 Descriptive Statistics of MHBMA Urinary Concentration Adjusted for Creatinine (pg/mg creat) - FAS**

Timepoint	Statistic	SA (N=39)	
		Raw value	%Change (*)
Baseline	n	39	
	BLOQ - n (%)	0	
	Geometric Mean (CV%)	1699.39 (147.05%)	
	Geometric 95% CI	1200.16, 2406.29	
	Median	2455.66	
	Q25, Q75	701.94, 3992.10	
	Min, Max	221.7, 6886.5	
	Mean (SD)	2583.12 (1902.737)	
	95% CI		

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.100 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_mhbma.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.7 Descriptive Statistics of MHBMA Urinary Concentration Adjusted for Creatinine (pg/mg creat) - FAS**

Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 1	n	80	80	41	41
	BLOQ - n (%)	4 (5.0%)		0	
	Geometric Mean (CV%)	586.07 (83.32%)		2459.59 (84.32%)	
	Geometric 95% CI	498.62, 688.86		1951.66, 3099.72	
	Median	617.09	-70.95	2581.30	5.21
	Q25, Q75	456.05, 947.89	-77.56, -61.41	1968.83, 3869.16	-9.20, 24.87
	Min, Max	52.9, 3777.4	-84.4, -9.6	327.9, 6974.4	-34.0, 102.2
	Mean (SD)	737.97 (539.998)	-66.19 (15.575)	3047.36 (1826.902)	9.20 (27.237)
	95% CI		-69.66, -62.72		0.59, 17.80

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.100 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_mhbma.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.7 Descriptive Statistics of MHBMA Urinary Concentration Adjusted for Creatinine (pg/mg creat) - FAS**

Timepoint	Statistic	SA (N=39)	
		Raw value	%Change (*)
Day 1	n	39	39
	BLOQ - n (%)	1 (2.6%)	
	Geometric Mean (CV%)	563.81 (75.86%)	
	Geometric 95% CI	453.13, 701.53	
	Median	614.82	-70.64
	Q25, Q75	349.01, 947.11	-78.01, -50.35
	Min, Max	133.7, 1677.1	-82.0, 35.8
	Mean (SD)	686.01 (407.681)	-61.84 (24.692)
	95% CI		-69.85, -53.83

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.100 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_mhbma.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.7 Descriptive Statistics of MHBMA Urinary Concentration Adjusted for Creatinine (pg/mg creat) - FAS**

Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 2	n	80	80	41	41
	BLOQ - n (%)	7 (8.8%)		0	
	Geometric Mean (CV%)	288.75 (48.27%)		2147.45 (91.56%)	
	Geometric 95% CI	260.78, 319.71		1678.65, 2747.17	
	Median	297.32	-87.39	2413.79	-2.26
	Q25, Q75	220.83, 364.42	-90.74, -72.96	1720.93, 3202.10	-21.34, 11.90
	Min, Max	84.9, 2363.6	-95.9, -9.3	208.0, 6588.6	-87.2, 97.4
	Mean (SD)	327.86 (256.164)	-79.01 (19.183)	2696.91 (1606.435)	-1.07 (31.864)
	95% CI		-83.28, -74.74		-11.13, 8.99

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.100 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_mhbma.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.7 Descriptive Statistics of MHBMA Urinary Concentration Adjusted for Creatinine (pg/mg creat) - FAS**

Timepoint	Statistic	SA (N=39)	
		Raw value	%Change (*)
Day 2	n	39	39
	BLOQ - n (%)	3 (7.7%)	
	Geometric Mean (CV%)	278.79 (36.08%)	
	Geometric 95% CI	248.90, 312.27	
	Median	278.11	-87.06
	Q25, Q75	239.63, 340.87	-91.98, -59.52
	Min, Max	106.6, 517.1	-97.2, 24.1
	Mean (SD)	294.79 (96.616)	-72.07 (32.632)
	95% CI		-82.65, -61.48

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.100 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_mhbma.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.7 Descriptive Statistics of MHBMA Urinary Concentration Adjusted for Creatinine (pg/mg creat) - FAS**

Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 3	n	79	79	41	41
	BLOQ - n (%)	9 (11.4%)		1 (2.4%)	
	Geometric Mean (CV%)	240.84 (59.39%)		2147.10 (90.66%)	
	Geometric 95% CI	212.94, 272.40		1681.44, 2741.72	
	Median	257.38	-89.97	2442.59	1.05
	Q25, Q75	195.79, 304.60	-92.71, -77.91	1913.46, 3118.09	-11.61, 21.38
	Min, Max	54.9, 5461.0	-98.3, 13.4	141.2, 7064.1	-97.3, 77.5
	Mean (SD)	315.13 (592.382)	-80.74 (20.935)	2651.08 (1455.587)	2.35 (31.773)
	95% CI		-85.43, -76.04		-7.68, 12.39

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.100 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_mhbma.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.7 Descriptive Statistics of MHBMA Urinary Concentration Adjusted for Creatinine (pg/mg creat) - FAS**

Timepoint	Statistic	SA (N=39)	
		Raw value	%Change (*)
Day 3	n	39	39
	BLOQ - n (%)	9 (23.1%)	
	Geometric Mean (CV%)	216.01 (48.38%)	
	Geometric 95% CI	186.17, 250.63	
	Median	223.88	-90.22
	Q25, Q75	164.70, 281.25	-93.85, -61.15
	Min, Max	79.2, 571.8	-98.1, 11.0
	Mean (SD)	238.40 (107.576)	-76.13 (29.126)
	95% CI		-85.57, -66.68

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.100 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_mhbm.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.7 Descriptive Statistics of MHBMA Urinary Concentration Adjusted for Creatinine (pg/mg creat) - FAS**

Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 4	n	79	79	41	41
	BLOQ - n (%)	16 (20.3%)		0	
	Geometric Mean (CV%)	215.96 (47.00%)		2387.21 (78.35%)	
	Geometric 95% CI	195.40, 238.70		1918.95, 2969.75	
	Median	250.61	-90.31	2551.96	1.86
	Q25, Q75	159.89, 298.28	-93.90, -77.05	1901.54, 3209.55	-15.00, 27.09
	Min, Max	61.3, 412.2	-98.6, -32.5	341.9, 7171.1	-51.9, 54.4
	Mean (SD)	235.82 (90.620)	-83.20 (16.333)	2901.19 (1694.657)	6.32 (26.218)
	95% CI		-86.87, -79.54		-1.96, 14.60

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.100 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_mhbma.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.7 Descriptive Statistics of MHBMA Urinary Concentration Adjusted for Creatinine (pg/mg creat) - FAS**

Timepoint	Statistic	SA (N=39)	
		Raw value	%Change (*)
Day 4	n	39	39
	BLOQ - n (%)	13 (33.3%)	
	Geometric Mean (CV%)	189.81 (51.44%)	
	Geometric 95% CI	162.22, 222.09	
	Median	202.03	-91.24
	Q25, Q75	124.06, 262.23	-95.66, -69.15
	Min, Max	71.9, 439.1	-98.0, 17.8
	Mean (SD)	211.18 (94.610)	-75.76 (33.253)
	95% CI		-86.55, -64.98

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.100 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_mhbm.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.7 Descriptive Statistics of MHBMA Urinary Concentration Adjusted for Creatinine (pg/mg creat) - FAS**

Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 5	n	79	79	41	41
	BLOQ - n (%)	34 (43.0%)		1 (2.4%)	
	Geometric Mean (CV%)	192.93 (46.00%)		2399.40 (89.21%)	
	Geometric 95% CI	174.90, 212.83		1884.60, 3054.83	
	Median	203.08	-92.03	2547.17	5.28
	Q25, Q75	145.87, 276.67	-94.62, -82.02	1981.81, 3692.67	-6.85, 23.26
	Min, Max	62.8, 384.6	-98.1, -41.6	155.8, 7163.6	-50.0, 55.1
	Mean (SD)	210.18 (80.697)	-84.98 (14.906)	2960.30 (1671.373)	6.89 (26.158)
	95% CI		-88.32, -81.63		-1.37, 15.15

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.100 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_mhbma.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.7 Descriptive Statistics of MHBMA Urinary Concentration Adjusted for Creatinine (pg/mg creat) - FAS**

Timepoint	Statistic	SA (N=39)	
		Raw value	%Change (*)
Day 5	n	39	39
	BLOQ - n (%)	16 (41.0%)	
	Geometric Mean (CV%)	163.17 (54.22%)	
	Geometric 95% CI	138.41, 192.36	
	Median	183.12	-92.83
	Q25, Q75	107.99, 239.76	-96.69, -70.55
	Min, Max	68.7, 431.1	-98.1, -6.4
	Mean (SD)	184.15 (90.464)	-79.56 (27.037)
	95% CI		-88.33, -70.79

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.100 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_mhbm.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.7.1 Descriptive Statistics of MHBMA Urinary Concentration Adjusted for Creatinine (pg/mg creat) by Sex - FAS**

Timepoint	Statistic	THS 2.2			
		Male (N=39)		Female (N=41)	
		Raw value	%Change (*)	Raw value	%Change (*)
Baseline	n	39		41	
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	1561.31 (123.46%)		2262.61 (98.51%)	
	Geometric 95% CI	1142.94, 2132.82		1744.68, 2934.28	
	Median	2165.61		3059.94	
	Q25, Q75	846.60, 2947.26		1261.24, 3981.91	
	Min, Max	231.9, 6258.9		252.4, 7218.5	
	Mean (SD)	2222.49 (1625.871)		2944.37 (1868.147)	
	95% CI				

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.100 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_mhbm.as

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.7.1 Descriptive Statistics of MHBMA Urinary Concentration Adjusted for Creatinine (pg/mg creat) by Sex - FAS**

Timepoint	Statistic	THS 2.2			
		Male (N=39)		Female (N=41)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 1	n	39	39	41	41
	BLOQ - n (%)	2 (5.1%)		2 (4.9%)	
	Geometric Mean (CV%)	492.21 (76.19%)		691.92 (85.48%)	
	Geometric 95% CI	395.28, 612.91		547.68, 874.13	
	Median	584.51	-71.21	748.09	-70.25
	Q25, Q75	371.76, 687.69	-77.51, -51.06	522.72, 1052.18	-77.61, -62.77
	Min, Max	97.1, 1921.7	-81.4, -36.5	52.9, 3777.4	-84.4, -9.6
	Mean (SD)	599.56 (371.948)	-66.02 (14.038)	869.63 (638.774)	-66.34 (17.084)
	95% CI		-70.58, -61.47		-71.74, -60.95

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.100 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_mhbm.as

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.7.1 Descriptive Statistics of MHBMA Urinary Concentration Adjusted for Creatinine (pg/mg creat) by Sex - FAS**

Timepoint	Statistic	THS 2.2			
		Male (N=39)		Female (N=41)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 2	n	39	39	41	41
	BLOQ - n (%)	2 (5.1%)		5 (12.2%)	
	Geometric Mean (CV%)	279.45 (52.40%)		297.88 (44.57%)	
	Geometric 95% CI	238.21, 327.83		260.43, 340.71	
	Median	265.76	-87.27	317.07	-88.89
	Q25, Q75	213.19, 342.08	-89.88, -57.66	254.27, 387.53	-91.96, -82.97
	Min, Max	84.9, 2363.6	-94.5, -9.3	107.1, 669.1	-95.9, -31.7
	Mean (SD)	332.51 (345.163)	-74.92 (22.222)	323.45 (127.974)	-82.90 (15.026)
	95% CI		-82.13, -67.71		-87.65, -78.15

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.100 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_mhbm.as

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.7.1 Descriptive Statistics of MHBMA Urinary Concentration Adjusted for Creatinine (pg/mg creat) by Sex - FAS**

Timepoint	Statistic	THS 2.2			
		Male (N=39)		Female (N=41)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 3	n	39	39	40	40
	BLOQ - n (%)	3 (7.7%)		6 (15.0%)	
	Geometric Mean (CV%)	215.84 (40.06%)		268.01 (73.88%)	
	Geometric 95% CI	190.47, 244.60		217.01, 330.99	
	Median	231.03	-89.97	285.37	-89.98
	Q25, Q75	172.41, 264.61	-91.77, -76.30	243.41, 323.27	-93.53, -80.56
	Min, Max	54.9, 433.0	-96.8, -31.3	71.4, 5461.0	-98.3, 13.4
	Mean (SD)	229.83 (75.852)	-79.66 (20.255)	398.30 (825.745)	-81.79 (21.783)
	95% CI		-86.23, -73.08		-88.77, -74.82

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.100 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_mhbm.as

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.7.1 Descriptive Statistics of MHBMA Urinary Concentration Adjusted for Creatinine (pg/mg creat) by Sex - FAS**

Timepoint	Statistic	THS 2.2			
		Male (N=39)		Female (N=41)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 4	n	39	39	40	40
	BLOQ - n (%)	9 (23.1%)		7 (17.5%)	
	Geometric Mean (CV%)	196.56 (47.81%)		236.73 (44.55%)	
	Geometric 95% CI	169.67, 227.70		206.61, 271.24	
	Median	201.85	-90.19	274.24	-90.54
	Q25, Q75	134.78, 278.05	-93.69, -73.32	190.65, 310.66	-93.93, -85.02
	Min, Max	83.1, 409.8	-98.2, -32.5	61.3, 412.2	-98.6, -47.1
	Mean (SD)	216.27 (92.611)	-80.76 (18.842)	254.88 (85.513)	-85.58 (13.263)
	95% CI		-86.88, -74.65		-89.83, -81.33

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.100 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_mhbm.as

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.7.1 Descriptive Statistics of MHBMA Urinary Concentration Adjusted for Creatinine (pg/mg creat) by Sex - FAS**

Timepoint	Statistic	THS 2.2			
		Male (N=39)		Female (N=41)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 5	n	39	39	40	40
	BLOQ - n (%)	13 (33.3%)		21 (52.5%)	
	Geometric Mean (CV%)	165.95 (49.54%)		223.46 (36.52%)	
	Geometric 95% CI	142.57, 193.17		199.55, 250.24	
	Median	178.05	-92.26	253.70	-91.84
	Q25, Q75	110.86, 253.26	-94.39, -77.09	174.87, 288.20	-94.81, -84.91
	Min, Max	62.8, 357.9	-98.1, -41.6	89.9, 384.6	-97.6, -52.1
	Mean (SD)	183.36 (78.809)	-83.89 (16.343)	236.33 (74.511)	-86.04 (13.482)
	95% CI		-89.19, -78.58		-90.36, -81.72

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.100 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_mhbm.as

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.7.1 Descriptive Statistics of MHBMA Urinary Concentration Adjusted for Creatinine (pg/mg creat) by Sex - FAS**

Timepoint	Statistic	CC			
		Male (N=21)		Female (N=20)	
		Raw value	%Change (*)	Raw value	%Change (*)
Baseline	n	21		20	
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	2458.12 (82.17%)		2178.12 (76.76%)	
	Geometric 95% CI	1772.62, 3408.72		1583.98, 2995.13	
	Median	2632.11		2495.58	
	Q25, Q75	1621.42, 4601.36		1850.85, 3279.11	
	Min, Max	493.5, 7533.3		311.7, 5147.1	
	Mean (SD)	3066.59 (1996.756)		2569.09 (1247.955)	
	95% CI				

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.100 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_mhbm.as

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.7.1 Descriptive Statistics of MHBMA Urinary Concentration Adjusted for Creatinine (pg/mg creat) by Sex - FAS**

Timepoint	Statistic	CC			
		Male (N=21)		Female (N=20)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 1	n	21	21	20	20
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	2663.82 (84.95%)		2261.99 (85.07%)	
	Geometric 95% CI	1904.56, 3725.75		1601.47, 3194.92	
	Median	2675.39	12.35	2415.56	3.79
	Q25, Q75	1968.83, 4992.87	-6.29, 26.11	1882.61, 3529.98	-9.51, 22.64
	Min, Max	415.2, 6974.4	-34.0, 102.2	327.9, 6197.2	-31.0, 40.2
	Mean (SD)	3313.00 (2041.028)	12.72 (33.436)	2768.44 (1575.288)	5.49 (18.888)
	95% CI		-2.50, 27.95		-3.35, 14.34

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.100 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_mhbm.as

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.7.1 Descriptive Statistics of MHBMA Urinary Concentration Adjusted for Creatinine (pg/mg creat) by Sex - FAS**

Timepoint	Statistic	CC			
		Male (N=21)		Female (N=20)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 2	n	21	21	20	20
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	2162.11 (104.37%)		2132.17 (81.17%)	
	Geometric 95% CI	1462.79, 3195.74		1528.37, 2974.51	
	Median	2311.76	-11.38	2453.41	2.92
	Q25, Q75	1720.93, 3961.17	-21.81, 11.82	1889.12, 2947.44	-20.02, 15.95
	Min, Max	208.0, 6588.6	-87.2, 97.4	339.3, 6528.1	-34.3, 41.5
	Mean (SD)	2822.80 (1808.717)	-2.25 (39.689)	2564.73 (1397.628)	0.18 (21.824)
	95% CI		-20.33, 15.82		-10.04, 10.40

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.100 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_mhbm.as

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.7.1 Descriptive Statistics of MHBMA Urinary Concentration Adjusted for Creatinine (pg/mg creat) by Sex - FAS**

Timepoint	Statistic	CC			
		Male (N=21)		Female (N=20)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 3	n	21	21	20	20
	BLOQ - n (%)	0		1 (5.0%)	
	Geometric Mean (CV%)	2377.40 (70.16%)		1929.25 (112.53%)	
	Geometric 95% CI	1782.55, 3170.76		1263.41, 2945.99	
	Median	2434.87	-3.90	2541.65	3.33
	Q25, Q75	1913.46, 3485.99	-21.85, 21.38	2037.46, 3084.24	-7.85, 19.55
	Min, Max	599.0, 7064.1	-50.3, 77.5	141.2, 5017.7	-97.3, 36.4
	Mean (SD)	2817.20 (1605.172)	2.38 (35.220)	2476.66 (1298.376)	2.33 (28.629)
	95% CI		-13.66, 18.41		-11.07, 15.73

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.100 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_mhbm.as

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.3.7.1 Descriptive Statistics of MHBMA Urinary Concentration Adjusted for Creatinine (pg/mg creat) by Sex - FAS**

Timepoint	Statistic	CC			
		Male (N=21)		Female (N=20)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 4	n	21	21	20	20
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	2545.55 (73.18%)		2231.55 (85.30%)	
	Geometric 95% CI	1889.42, 3429.54		1578.75, 3154.26	
	Median	2369.75	-3.05	2741.23	2.59
	Q25, Q75	1901.54, 3621.40	-13.97, 36.69	1925.83, 3179.33	-16.22, 19.43
	Min, Max	627.2, 7171.1	-51.9, 54.4	341.9, 5096.8	-28.7, 54.2
	Mean (SD)	3092.14 (1969.722)	7.80 (29.550)	2700.70 (1370.916)	4.76 (22.868)
	95% CI		-5.66, 21.25		-5.95, 15.47

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.100 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_mhbm.as

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.7.1 Descriptive Statistics of MHBMA Urinary Concentration Adjusted for Creatinine (pg/mg creat) by Sex - FAS**

Timepoint	Statistic	CC			
		Male (N=21)		Female (N=20)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 5	n	21	21	20	20
	BLOQ - n (%)	0		1 (5.0%)	
	Geometric Mean (CV%)	2523.29 (79.16%)		2275.85 (102.16%)	
	Geometric 95% CI	1836.91, 3466.15		1532.20, 3380.44	
	Median	2456.31	1.88	2701.38	8.29
	Q25, Q75	1857.75, 4440.30	-10.44, 23.26	2080.77, 3608.56	-3.87, 27.27
	Min, Max	502.8, 7163.6	-34.3, 55.1	155.8, 5608.3	-50.0, 44.6
	Mean (SD)	3100.99 (1942.576)	6.03 (27.582)	2812.57 (1365.270)	7.79 (25.261)
	95% CI		-6.53, 18.59		-4.04, 19.61

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.100 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_mhbm.as

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.7.1 Descriptive Statistics of MHBMA Urinary Concentration Adjusted for Creatinine (pg/mg creat) by Sex - FAS**

Timepoint	Statistic	SA			
		Male (N=20)		Female (N=19)	
		Raw value	%Change (*)	Raw value	%Change (*)
Baseline	n	20		19	
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	1475.86 (160.21%)		1971.36 (135.55%)	
	Geometric 95% CI	870.64, 2501.77		1205.06, 3224.96	
	Median	2015.66		2812.50	
	Q25, Q75	537.52, 3918.82		1204.95, 3992.10	
	Min, Max	233.3, 5712.2		221.7, 6886.5	
	Mean (SD)	2372.99 (1915.833)		2804.31 (1915.215)	
	95% CI				

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.100 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_mhbm.as

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.7.1 Descriptive Statistics of MHBMA Urinary Concentration Adjusted for Creatinine (pg/mg creat) by Sex - FAS**

Timepoint	Statistic	SA			
		Male (N=20)		Female (N=19)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 1	n	20	20	19	19
	BLOQ - n (%)	0		1 (5.3%)	
	Geometric Mean (CV%)	526.82 (69.84%)		605.56 (83.57%)	
	Geometric 95% CI	392.24, 707.57		426.40, 860.00	
	Median	571.20	-69.27	628.38	-70.70
	Q25, Q75	331.47, 855.40	-76.31, -48.12	411.37, 1154.53	-78.39, -64.30
	Min, Max	168.8, 1677.1	-82.0, 35.8	133.7, 1649.4	-79.7, -34.1
	Mean (SD)	629.69 (382.124)	-56.94 (31.336)	745.30 (435.326)	-67.01 (13.992)
	95% CI		-71.61, -42.27		-73.76, -60.26

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.100 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_mhbm.as

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.3.7.1 Descriptive Statistics of MHBMA Urinary Concentration Adjusted for Creatinine (pg/mg creat) by Sex - FAS**

Timepoint	Statistic	SA			
		Male (N=20)		Female (N=19)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 2	n	20	20	19	19
	BLOQ - n (%)	0		3 (15.8%)	
	Geometric Mean (CV%)	282.24 (28.33%)		275.20 (43.94%)	
	Geometric 95% CI	247.82, 321.44		224.75, 336.97	
	Median	268.92	-86.75	299.26	-87.77
	Q25, Q75	244.11, 331.71	-92.74, -49.09	239.63, 349.57	-91.76, -83.04
	Min, Max	159.2, 490.4	-93.6, 16.9	106.6, 517.1	-97.2, 24.1
	Mean (SD)	292.86 (83.443)	-67.88 (34.220)	296.82 (111.123)	-76.47 (31.177)
	95% CI		-83.90, -51.86		-91.50, -61.44

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.100 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_mhbm.as

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.7.1 Descriptive Statistics of MHBMA Urinary Concentration Adjusted for Creatinine (pg/mg creat) by Sex - FAS**

Timepoint	Statistic	SA			
		Male (N=20)		Female (N=19)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 3	n	20	20	19	19
	BLOQ - n (%)	2 (10.0%)		7 (36.8%)	
	Geometric Mean (CV%)	223.00 (45.45%)		208.88 (52.50%)	
	Geometric 95% CI	182.07, 273.14		164.67, 264.96	
	Median	245.62	-87.57	206.93	-90.26
	Q25, Q75	183.06, 298.34	-94.24, -51.30	139.66, 275.07	-93.85, -83.25
	Min, Max	79.2, 395.5	-98.1, 11.0	97.8, 571.8	-97.5, -19.2
	Mean (SD)	240.97 (87.421)	-69.52 (34.773)	235.70 (127.862)	-83.08 (20.371)
	95% CI		-85.80, -53.24		-92.90, -73.26

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.100 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_mhbm.as

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.7.1 Descriptive Statistics of MHBMA Urinary Concentration Adjusted for Creatinine (pg/mg creat) by Sex - FAS**

Timepoint	Statistic	SA			
		Male (N=20)		Female (N=19)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 4	n	20	20	19	19
	BLOQ - n (%)	6 (30.0%)		7 (36.8%)	
	Geometric Mean (CV%)	186.86 (50.76%)		192.97 (53.65%)	
	Geometric 95% CI	149.34, 233.80		151.43, 245.91	
	Median	187.45	-90.70	208.33	-91.24
	Q25, Q75	141.10, 261.51	-95.26, -62.87	114.41, 290.84	-96.23, -85.29
	Min, Max	71.9, 399.2	-98.0, 17.8	77.4, 439.1	-97.6, 6.8
	Mean (SD)	206.84 (92.132)	-72.75 (35.310)	215.75 (99.468)	-78.94 (31.584)
	95% CI		-89.28, -56.22		-94.16, -63.71

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.100 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_mhbm.as

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.7.1 Descriptive Statistics of MHBMA Urinary Concentration Adjusted for Creatinine (pg/mg creat) by Sex - FAS**

Timepoint	Statistic	SA			
		Male (N=20)		Female (N=19)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 5	n	20	20	19	19
	BLOQ - n (%)	5 (25.0%)		11 (57.9%)	
	Geometric Mean (CV%)	156.65 (56.19%)		170.33 (53.37%)	
	Geometric 95% CI	122.59, 200.17		133.81, 216.81	
	Median	196.00	-91.22	151.52	-93.04
	Q25, Q75	99.61, 233.52	-96.76, -63.78	107.99, 259.91	-96.28, -87.37
	Min, Max	68.7, 303.5	-98.1, -6.4	89.6, 431.1	-97.7, -13.8
	Mean (SD)	176.11 (79.033)	-77.02 (28.371)	192.61 (102.646)	-82.23 (26.057)
	95% CI		-90.30, -63.74		-94.79, -69.66

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.100 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_mhbm.as

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.7.2 Descriptive Statistics of MHBMA Urinary Concentration Adjusted for Creatinine (pg/mg creat) by Cigarette Consumption - FAS**

Timepoint	Statistic	THS 2.2			
		10-19 cig/day (N=41)		>19 cig/day (N=39)	
		Raw value	%Change (*)	Raw value	%Change (*)
Baseline	n	41		39	
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	1553.92 (118.15%)		2317.60 (101.54%)	
	Geometric 95% CI	1156.90, 2087.18		1764.16, 3044.65	
	Median	1961.65		2450.47	
	Q25, Q75	913.72, 3142.86		1545.60, 4400.00	
	Min, Max	231.9, 6355.3		295.9, 7218.5	
	Mean (SD)	2163.31 (1480.414)		3043.59 (1968.923)	
	95% CI				

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.100 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_mhbmacc.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.7.2 Descriptive Statistics of MHBMA Urinary Concentration Adjusted for Creatinine (pg/mg creat) by Cigarette Consumption - FAS**

Timepoint	Statistic	THS 2.2			
		10-19 cig/day (N=41)		>19 cig/day (N=39)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 1	n	41	41	39	39
	BLOQ - n (%)	3 (7.3%)		1 (2.6%)	
	Geometric Mean (CV%)	475.75 (79.57%)		729.74 (78.85%)	
	Geometric 95% CI	381.39, 593.45		582.48, 914.23	
	Median	554.17	-71.05	712.69	-70.18
	Q25, Q75	376.14, 751.56	-77.94, -59.82	519.69, 1065.64	-76.86, -62.77
	Min, Max	52.9, 1211.0	-84.4, -26.0	116.3, 3777.4	-81.7, -9.6
	Mean (SD)	573.89 (306.153)	-66.73 (14.590)	910.47 (669.303)	-65.62 (16.722)
	95% CI		-71.34, -62.12		-71.05, -60.20

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.100 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_mhbmasc.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.7.2 Descriptive Statistics of MHBMA Urinary Concentration Adjusted for Creatinine (pg/mg creat) by Cigarette Consumption - FAS**

Timepoint	Statistic	THS 2.2			
		10-19 cig/day (N=41)		>19 cig/day (N=39)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 2	n	41	41	39	39
	BLOQ - n (%)	5 (12.2%)		2 (5.1%)	
	Geometric Mean (CV%)	256.03 (55.25%)		327.65 (36.19%)	
	Geometric 95% CI	217.54, 301.33		292.43, 367.12	
	Median	264.26	-87.60	333.83	-87.27
	Q25, Q75	196.53, 317.08	-90.60, -65.29	265.75, 434.60	-91.08, -81.86
	Min, Max	84.9, 2363.6	-95.9, -9.3	126.9, 669.1	-94.5, -32.5
	Mean (SD)	309.64 (340.282)	-76.33 (21.871)	347.03 (117.469)	-81.83 (15.671)
	95% CI		-83.23, -69.42		-86.91, -76.75

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.100 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_mhbmasc.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.7.2 Descriptive Statistics of MHBMA Urinary Concentration Adjusted for Creatinine (pg/mg creat) by Cigarette Consumption - FAS**

Timepoint	Statistic	THS 2.2			
		10-19 cig/day (N=41)		>19 cig/day (N=39)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 3	n	40	40	39	39
	BLOQ - n (%)	6 (15.0%)		3 (7.7%)	
	Geometric Mean (CV%)	208.10 (49.92%)		279.78 (64.33%)	
	Geometric 95% CI	178.96, 241.99		231.19, 338.59	
	Median	240.44	-90.49	283.89	-89.72
	Q25, Q75	159.64, 297.32	-92.84, -64.05	234.96, 317.81	-92.54, -82.13
	Min, Max	54.9, 433.0	-98.3, -31.4	80.5, 5461.0	-96.8, 13.4
	Mean (SD)	228.54 (90.279)	-78.86 (21.033)	403.94 (834.236)	-82.66 (20.931)
	95% CI		-85.60, -72.13		-89.45, -75.87

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.100 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_mhbmasc.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.7.2 Descriptive Statistics of MHBMA Urinary Concentration Adjusted for Creatinine (pg/mg creat) by Cigarette Consumption - FAS**

Timepoint	Statistic	THS 2.2			
		10-19 cig/day (N=41)		>19 cig/day (N=39)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 4	n	40	40	39	39
	BLOQ - n (%)	10 (25.0%)		6 (15.4%)	
	Geometric Mean (CV%)	194.74 (52.20%)		240.14 (38.70%)	
	Geometric 95% CI	166.44, 227.84		212.75, 271.05	
	Median	210.17	-90.99	264.89	-90.19
	Q25, Q75	131.51, 285.16	-93.84, -67.80	193.98, 308.42	-93.94, -84.89
	Min, Max	61.3, 409.8	-98.6, -32.5	93.6, 412.2	-98.2, -47.1
	Mean (SD)	216.96 (95.779)	-80.34 (19.489)	255.16 (81.751)	-86.14 (11.847)
	95% CI		-86.58, -74.10		-89.98, -82.29

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.100 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_mhbmasc.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.7.2 Descriptive Statistics of MHBMA Urinary Concentration Adjusted for Creatinine (pg/mg creat) by Cigarette Consumption - FAS**

Timepoint	Statistic	THS 2.2			
		10-19 cig/day (N=41)		>19 cig/day (N=39)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 5	n	40	40	39	39
	BLOQ - n (%)	22 (55.0%)		12 (30.8%)	
	Geometric Mean (CV%)	170.55 (54.14%)		218.95 (32.03%)	
	Geometric 95% CI	145.02, 200.57		197.85, 242.30	
	Median	176.69	-92.39	241.55	-90.79
	Q25, Q75	111.73, 264.12	-94.75, -70.59	178.04, 279.17	-94.45, -85.60
	Min, Max	62.8, 384.6	-97.6, -41.6	87.9, 359.9	-98.1, -42.3
	Mean (SD)	192.11 (91.579)	-82.98 (16.664)	228.71 (63.741)	-87.03 (12.749)
	95% CI		-88.31, -77.64		-91.16, -82.89

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.100 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_mhbmacc.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.7.2 Descriptive Statistics of MHBMA Urinary Concentration Adjusted for Creatinine (pg/mg creat) by Cigarette Consumption - FAS**

Timepoint	Statistic	CC			
		10-19 cig/day (N=21)		>19 cig/day (N=20)	
		Raw value	%Change (*)	Raw value	%Change (*)
Baseline	n	21		20	
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	2491.59 (55.97%)		2147.42 (102.22%)	
	Geometric 95% CI	1964.63, 3159.88		1445.46, 3190.27	
	Median	2500.00		2355.71	
	Q25, Q75	1939.79, 3507.37		1405.97, 4216.27	
	Min, Max	616.1, 6092.1		311.7, 7533.3	
	Mean (SD)	2798.49 (1322.176)		2850.60 (2011.491)	
	95% CI				

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.100 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_mhbmacc.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.7.2 Descriptive Statistics of MHBMA Urinary Concentration Adjusted for Creatinine (pg/mg creat) by Cigarette Consumption - FAS**

Timepoint	Statistic	CC			
		10-19 cig/day (N=21)		>19 cig/day (N=20)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 1	n	21	21	20	20
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	2619.41 (64.34%)		2302.27 (106.40%)	
	Geometric 95% CI	2003.79, 3424.16		1532.16, 3459.46	
	Median	2675.39	0.72	2423.86	8.02
	Q25, Q75	2044.64, 3815.03	-10.41, 26.11	1791.86, 4891.94	-5.97, 23.55
	Min, Max	551.9, 6844.7	-34.0, 102.2	327.9, 6974.4	-33.6, 43.1
	Mean (SD)	3019.61 (1548.848)	9.25 (32.850)	3076.51 (2121.210)	9.14 (20.636)
	95% CI		-5.71, 24.20		-0.52, 18.81

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.100 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_mhbmacc.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.7.2 Descriptive Statistics of MHBMA Urinary Concentration Adjusted for Creatinine (pg/mg creat) by Cigarette Consumption - FAS**

Timepoint	Statistic	CC			
		10-19 cig/day (N=21)		>19 cig/day (N=20)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 2	n	21	21	20	20
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	2170.29 (91.46%)		2123.73 (94.86%)	
	Geometric 95% CI	1521.91, 3094.89		1459.76, 3089.71	
	Median	2692.31	-8.88	2142.80	0.65
	Q25, Q75	1990.13, 3049.51	-25.33, 8.59	1657.39, 3948.82	-17.75, 27.79
	Min, Max	208.0, 5399.0	-87.2, 97.4	339.3, 6588.6	-47.4, 41.9
	Mean (SD)	2637.80 (1283.347)	-4.33 (36.524)	2758.97 (1921.284)	2.36 (26.631)
	95% CI		-20.96, 12.30		-10.11, 14.83

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.100 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_mhbmacc.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.7.2 Descriptive Statistics of MHBMA Urinary Concentration Adjusted for Creatinine (pg/mg creat) by Cigarette Consumption - FAS**

Timepoint	Statistic	CC			
		10-19 cig/day (N=21)		>19 cig/day (N=20)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 3	n	21	21	20	20
	BLOQ - n (%)	0		1 (5.0%)	
	Geometric Mean (CV%)	2512.06 (54.21%)		1820.82 (124.48%)	
	Geometric 95% CI	1993.88, 3164.90		1157.79, 2863.53	
	Median	2742.35	-1.62	2192.80	3.33
	Q25, Q75	2382.97, 3118.09	-11.61, 22.02	1111.40, 3524.83	-10.75, 20.14
	Min, Max	622.5, 5017.7	-50.3, 77.5	141.2, 7064.1	-97.3, 59.6
	Mean (SD)	2771.00 (1079.016)	5.03 (30.592)	2525.17 (1789.513)	-0.45 (33.526)
	95% CI		-8.90, 18.96		-16.15, 15.24

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.100 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_mhbmacc.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.7.2 Descriptive Statistics of MHBMA Urinary Concentration Adjusted for Creatinine (pg/mg creat) by Cigarette Consumption - FAS**

Timepoint	Statistic	CC			
		10-19 cig/day (N=21)		>19 cig/day (N=20)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 4	n	21	21	20	20
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	2468.62 (64.99%)		2304.62 (94.10%)	
	Geometric 95% CI	1884.17, 3234.36		1587.62, 3345.44	
	Median	2543.40	-3.05	2706.71	6.33
	Q25, Q75	2047.37, 3195.07	-16.35, 17.82	1645.93, 3905.47	-7.80, 37.11
	Min, Max	439.4, 5748.0	-36.3, 41.7	341.9, 7171.1	-51.9, 54.4
	Mean (SD)	2842.18 (1431.774)	1.80 (24.173)	2963.15 (1969.838)	11.06 (28.032)
	95% CI		-9.21, 12.81		-2.07, 24.18

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.100 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_mhbmacc.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.7.2 Descriptive Statistics of MHBMA Urinary Concentration Adjusted for Creatinine (pg/mg creat) by Cigarette Consumption - FAS**

Timepoint	Statistic	CC			
		10-19 cig/day (N=21)		>19 cig/day (N=20)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 5	n	21	21	20	20
	BLOQ - n (%)	0		1 (5.0%)	
	Geometric Mean (CV%)	2495.89 (65.82%)		2302.10 (115.76%)	
	Geometric 95% CI	1899.45, 3279.61		1495.27, 3544.28	
	Median	2793.18	-0.71	2492.50	17.83
	Q25, Q75	2089.28, 3305.42	-10.44, 15.25	1851.81, 4775.49	-6.28, 30.01
	Min, Max	397.3, 6266.2	-35.5, 55.1	155.8, 7163.6	-50.0, 51.3
	Mean (SD)	2853.01 (1319.796)	3.08 (25.410)	3072.95 (2005.344)	10.89 (26.983)
	95% CI		-8.49, 14.65		-1.75, 23.52

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.100 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_mhbmacc.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.7.2 Descriptive Statistics of MHBMA Urinary Concentration Adjusted for Creatinine (pg/mg creat) by Cigarette Consumption - FAS**

Timepoint	Statistic	SA			
		10-19 cig/day (N=19)		>19 cig/day (N=20)	
		Raw value	%Change (*)	Raw value	%Change (*)
Baseline	n	19		20	
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	1718.75 (142.62%)		1681.21 (157.90%)	
	Geometric 95% CI	1034.40, 2855.83		996.08, 2837.55	
	Median	2375.00		2550.91	
	Q25, Q75	1150.53, 3649.46		622.00, 4227.06	
	Min, Max	221.7, 5370.4		263.5, 6886.5	
	Mean (SD)	2478.67 (1639.317)		2682.34 (2161.857)	
	95% CI				

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.100 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_mhbmacc.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.7.2 Descriptive Statistics of MHBMA Urinary Concentration Adjusted for Creatinine (pg/mg creat) by Cigarette Consumption - FAS**

Timepoint	Statistic	SA			
		10-19 cig/day (N=19)		>19 cig/day (N=20)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 1	n	19	19	20	20
	BLOQ - n (%)	1 (5.3%)		0	
	Geometric Mean (CV%)	548.15 (82.93%)		579.11 (71.37%)	
	Geometric 95% CI	386.78, 776.83		428.90, 781.93	
	Median	628.38	-69.76	581.74	-70.69
	Q25, Q75	476.42, 763.70	-75.87, -56.73	331.47, 1032.99	-78.20, -48.12
	Min, Max	133.7, 1677.1	-82.0, -27.6	193.0, 1649.4	-82.0, 35.8
	Mean (SD)	672.66 (396.551)	-65.33 (15.445)	698.70 (427.884)	-58.53 (31.141)
	95% CI		-72.78, -57.88		-73.11, -43.95

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.100 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_mhbmasc.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.7.2 Descriptive Statistics of MHBMA Urinary Concentration Adjusted for Creatinine (pg/mg creat) by Cigarette Consumption - FAS**

Timepoint	Statistic	SA			
		10-19 cig/day (N=19)		>19 cig/day (N=20)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 2	n	19	19	20	20
	BLOQ - n (%)	1 (5.3%)		2 (10.0%)	
	Geometric Mean (CV%)	276.04 (27.45%)		281.42 (43.84%)	
	Geometric 95% CI	242.42, 314.33		231.28, 342.43	
	Median	272.98	-86.46	280.17	-87.41
	Q25, Q75	247.25, 322.54	-91.90, -71.96	236.65, 385.75	-92.30, -49.11
	Min, Max	159.2, 490.4	-93.3, 24.1	106.6, 517.1	-97.2, 16.9
	Mean (SD)	285.53 (76.519)	-74.50 (31.041)	303.59 (113.820)	-69.76 (34.717)
	95% CI		-89.47, -59.54		-86.01, -53.50

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.100 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_mhbmacc.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.7.2 Descriptive Statistics of MHBMA Urinary Concentration Adjusted for Creatinine (pg/mg creat) by Cigarette Consumption - FAS**

Timepoint	Statistic	SA			
		10-19 cig/day (N=19)		>19 cig/day (N=20)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 3	n	19	19	20	20
	BLOQ - n (%)	7 (36.8%)		2 (10.0%)	
	Geometric Mean (CV%)	184.43 (50.84%)		251.00 (40.81%)	
	Geometric 95% CI	146.37, 232.39		208.88, 301.62	
	Median	185.83	-90.61	250.72	-89.75
	Q25, Q75	116.27, 271.48	-94.80, -71.83	196.66, 298.34	-93.75, -53.13
	Min, Max	79.2, 393.2	-98.1, -19.2	125.3, 571.8	-97.5, 11.0
	Mean (SD)	204.39 (91.122)	-81.16 (22.243)	270.72 (114.116)	-71.35 (34.331)
	95% CI		-91.88, -70.43		-87.42, -55.27

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.100 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_mhbmacc.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.7.2 Descriptive Statistics of MHBMA Urinary Concentration Adjusted for Creatinine (pg/mg creat) by Cigarette Consumption - FAS**

Timepoint	Statistic	SA			
		10-19 cig/day (N=19)		>19 cig/day (N=20)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 4	n	19	19	20	20
	BLOQ - n (%)	10 (52.6%)		3 (15.0%)	
	Geometric Mean (CV%)	159.32 (44.86%)		224.17 (51.59%)	
	Geometric 95% CI	129.61, 195.84		178.58, 281.39	
	Median	173.31	-91.24	255.15	-90.60
	Q25, Q75	114.41, 223.46	-95.96, -85.29	170.14, 317.85	-95.24, -53.39
	Min, Max	71.9, 306.7	-98.0, 6.8	77.4, 439.1	-97.6, 17.8
	Mean (SD)	172.81 (68.323)	-81.00 (28.969)	247.63 (103.000)	-70.79 (36.919)
	95% CI		-94.97, -67.03		-88.07, -53.51

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.100 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_mhbmacc.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.7.2 Descriptive Statistics of MHBMA Urinary Concentration Adjusted for Creatinine (pg/mg creat) by Cigarette Consumption - FAS**

Timepoint	Statistic	SA			
		10-19 cig/day (N=19)		>19 cig/day (N=20)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 5	n	19	19	20	20
	BLOQ - n (%)	10 (52.6%)		6 (30.0%)	
	Geometric Mean (CV%)	136.67 (45.46%)		193.09 (56.40%)	
	Geometric 95% CI	110.90, 168.43		150.99, 246.93	
	Median	146.44	-93.62	227.72	-91.28
	Q25, Q75	92.08, 201.62	-96.83, -86.33	113.23, 278.15	-96.26, -62.77
	Min, Max	68.7, 298.4	-98.1, -13.8	71.7, 431.1	-98.0, -6.4
	Mean (SD)	148.99 (62.738)	-84.22 (22.683)	217.55 (101.135)	-75.13 (30.532)
	95% CI		-95.16, -73.28		-89.42, -60.83

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.100 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_mhbmacc.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.8 Descriptive Statistics of 3-HPMA Urinary Concentration Adjusted for Creatinine (ng/mg creat) - FAS**

Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)	
		Raw value	%Change (*)	Raw value	%Change (*)	Raw value	%Change (*)
Baseline	n	80		41		39	
	BLOQ - n (%)	0		0		0	
	Geometric Mean (CV%)	841.84 (58.66%)		799.37 (47.28%)		861.88 (47.34%)	
	Geometric 95% CI	745.88, 950.14		693.71, 921.13		744.98, 997.12	
	Median	827.47		837.67		899.03	
	Q25, Q75	638.81, 1210.72		609.04, 1051.34		657.52, 1247.57	
	Min, Max	94.6, 2495.7		287.9, 2246.7		352.7, 2530.7	
	Mean (SD)	957.14 (467.226)		879.91 (397.995)		950.69 (442.010)	
	95% CI						

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (20.0 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hpma.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.8 Descriptive Statistics of 3-HPMA Urinary Concentration Adjusted for Creatinine (ng/mg creat) - FAS**

Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)	
		Raw value	%Change (*)	Raw value	%Change (*)	Raw value	%Change (*)
Day 1	n	80	80	41	41	39	39
	BLOQ - n (%)	0		0		0	
	Geometric Mean (CV%)	472.09 (39.66%)		964.92 (44.15%)		367.06 (33.84%)	
	Geometric 95% CI	433.60, 514.00		844.59, 1102.39		329.90, 408.40	
	Median	489.90	-47.28	973.03	15.91	380.74	-57.06
	Q25, Q75	381.36, 595.73	-55.01, -37.27	720.10, 1341.47	8.39, 37.56	310.07, 447.69	-64.74, -45.70
	Min, Max	82.2, 953.7	-70.1, 77.1	427.8, 2102.3	-26.6, 114.1	174.5, 794.7	-80.8, -24.2
	Mean (SD)	503.03 (169.415)	-39.99 (25.202)	1048.97 (425.844)	24.37 (32.093)	386.74 (128.870)	-55.29 (13.894)
	95% CI	-45.60, -34.38		14.23, 34.50		-59.79, -50.78	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (20.0 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hpma.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.8 Descriptive Statistics of 3-HPMA Urinary Concentration Adjusted for Creatinine (ng/mg creat) - FAS**

Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)	
		Raw value	%Change (*)	Raw value	%Change (*)	Raw value	%Change (*)
Day 2	n	80	80	41	41	39	39
	BLOQ - n (%)	0		0		0	
	Geometric Mean (CV%)	316.19 (44.03%)		781.80 (50.32%)		202.82 (29.97%)	
	Geometric 95% CI	287.92, 347.24		672.93, 908.28		184.42, 223.05	
	Median	335.41	-62.25	818.53	-2.52	198.10	-75.16
	Q25, Q75	253.10, 392.48	-70.79, -53.06	558.41, 1127.10	-14.54, 15.64	177.64, 237.48	-82.79, -65.79
	Min, Max	53.7, 730.6	-86.3, 13.1	191.2, 1646.6	-67.8, 66.2	66.1, 351.1	-91.5, -55.3
	Mean (SD)	341.33 (126.302)	-59.83 (15.846)	866.28 (383.749)	1.68 (27.193)	210.60 (54.964)	-74.40 (9.906)
	95% CI		-63.36, -56.30		-6.91, 10.27		-77.62, -71.19

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (20.0 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hpma.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.8 Descriptive Statistics of 3-HPMA Urinary Concentration Adjusted for Creatinine (ng/mg creat) - FAS**

Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)	
		Raw value	%Change (*)	Raw value	%Change (*)	Raw value	%Change (*)
Day 3	n	79	79	41	41	39	39
	BLOQ - n (%)	0		0		0	
	Geometric Mean (CV%)	438.42 (43.74%)		917.63 (42.58%)		292.06 (39.26%)	
	Geometric 95% CI	399.20, 481.49		806.71, 1043.79		258.33, 330.20	
	Median	442.88	-49.14	1001.95	15.50	304.24	-65.40
	Q25, Q75	351.63, 527.92	-59.01, -39.01	637.54, 1131.21	0.89, 31.69	250.26, 365.95	-71.91, -56.94
	Min, Max	38.8, 1671.4	-83.4, 83.6	493.0, 1932.0	-57.9, 132.4	74.1, 532.2	-90.4, -36.5
	Mean (SD)	472.49 (196.719)	-44.78 (24.495)	996.39 (420.904)	20.14 (36.678)	310.09 (99.388)	-63.74 (12.889)
	95% CI	-50.27, -39.29		8.56, 31.73		-67.92, -59.55	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (20.0 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hpma.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.8 Descriptive Statistics of 3-HPMA Urinary Concentration Adjusted for Creatinine (ng/mg creat) - FAS**

Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)	
		Raw value	%Change (*)	Raw value	%Change (*)	Raw value	%Change (*)
Day 4	n	79	79	41	41	39	39
	BLOQ - n (%)	0		0		0	
	Geometric Mean (CV%)	280.73 (45.11%)		791.20 (48.33%)		152.31 (41.31%)	
	Geometric 95% CI	254.93, 309.15		684.67, 914.31		133.92, 173.23	
	Median	271.63	-68.07	854.77	-0.67	152.17	-81.89
	Q25, Q75	229.09, 375.98	-74.12, -60.15	555.26, 1078.79	-13.79, 12.01	111.95, 193.23	-85.28, -77.66
	Min, Max	33.7, 597.0	-84.1, 35.7	318.5, 1806.9	-45.1, 62.3	67.2, 354.1	-94.1, -65.5
	Mean (SD)	303.71 (114.118)	-63.96 (19.119)	873.50 (388.498)	2.12 (25.821)	164.28 (65.270)	-81.04 (6.873)
	95% CI						

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (20.0 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hpma.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.8 Descriptive Statistics of 3-HPMA Urinary Concentration Adjusted for Creatinine (ng/mg creat) - FAS**

Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)	
		Raw value	%Change (*)	Raw value	%Change (*)	Raw value	%Change (*)
Day 5	n	79	79	41	41	39	39
	BLOQ - n (%)	1 (1.3%)		0		0	
	Geometric Mean (CV%)	402.26 (43.36%)		931.01 (39.44%)		245.69 (25.98%)	
	Geometric 95% CI	366.55, 441.45		825.73, 1049.72		226.15, 266.91	
	Median	413.67	-54.93	941.04	9.38	240.26	-72.54
	Q25, Q75	310.58, 496.82	-62.28, -44.38	666.66, 1187.82	-1.19, 41.70	213.26, 288.38	-77.67, -63.45
	Min, Max	36.8, 1261.8	-73.9, 44.6	451.6, 1896.6	-44.8, 115.1	126.9, 436.5	-85.8, -44.5
	Mean (SD)	431.93 (162.496)	-49.68 (20.696)	998.95 (384.992)	20.28 (31.247)	253.60 (65.507)	-69.45 (11.761)
	95% CI		-54.32, -45.04		10.41, 30.14		-73.27, -65.63

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (20.0 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hpma.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.8.1 Descriptive Statistics of 3-HPMA Urinary Concentration Adjusted for Creatinine (ng/mg creat) by Sex - FAS**

Timepoint	Statistic	THS 2.2			
		Male (N=39)		Female (N=41)	
		Raw value	%Change (*)	Raw value	%Change (*)
Baseline	n	39		41	
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	819.72 (36.48%)		863.43 (76.84%)	
	Geometric 95% CI	731.00, 919.22		696.38, 1070.55	
	Median	764.76		987.03	
	Q25, Q75	638.79, 1105.32		646.34, 1415.79	
	Min, Max	406.9, 1807.7		94.6, 2495.7	
	Mean (SD)	873.29 (332.064)		1036.91 (559.474)	
	95% CI				

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (20.0 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hpmas.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.8.1 Descriptive Statistics of 3-HPMA Urinary Concentration Adjusted for Creatinine (ng/mg creat) by Sex - FAS**

Timepoint	Statistic	THS 2.2			
		Male (N=39)		Female (N=41)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 1	n	39	39	41	41
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	442.52 (29.63%)		502.05 (47.13%)	
	Geometric 95% CI	402.81, 486.16		435.87, 578.28	
	Median	416.07	-48.29	524.77	-46.77
	Q25, Q75	359.59, 563.64	-53.62, -39.30	426.28, 677.42	-58.09, -15.60
	Min, Max	188.7, 776.3	-70.1, 5.5	82.2, 953.7	-68.0, 77.1
	Mean (SD)	460.56 (130.829)	-44.05 (16.131)	543.43 (192.335)	-36.13 (31.238)
	95% CI		-49.29, -38.82		-45.99, -26.26

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (20.0 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hpmas.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.8.1 Descriptive Statistics of 3-HPMA Urinary Concentration Adjusted for Creatinine (ng/mg creat) by Sex - FAS**

Timepoint	Statistic	THS 2.2			
		Male (N=39)		Female (N=41)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 2	n	39	39	41	41
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	330.28 (33.46%)		303.35 (52.75%)	
	Geometric 95% CI	297.18, 367.06		259.43, 354.71	
	Median	332.07	-60.64	338.75	-64.91
	Q25, Q75	275.45, 388.34	-67.52, -52.31	247.17, 396.63	-73.08, -54.39
	Min, Max	141.7, 730.6	-77.1, 13.1	53.7, 639.8	-86.3, -12.3
	Mean (SD)	347.85 (117.458)	-57.57 (15.783)	335.13 (135.345)	-61.98 (15.797)
	95% CI		-62.69, -52.45		-66.97, -56.98

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (20.0 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hpmas.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.8.1 Descriptive Statistics of 3-HPMA Urinary Concentration Adjusted for Creatinine (ng/mg creat) by Sex - FAS**

Timepoint	Statistic	THS 2.2			
		Male (N=39)		Female (N=41)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 3	n	39	39	40	40
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	419.86 (26.84%)		457.30 (56.68%)	
	Geometric 95% CI	385.46, 457.34		386.27, 541.39	
	Median	422.33	-47.02	463.58	-52.02
	Q25, Q75	341.38, 503.02	-58.05, -36.66	390.76, 553.52	-59.33, -40.12
	Min, Max	276.9, 719.8	-68.7, -21.3	38.8, 1671.4	-83.4, 83.6
	Mean (SD)	434.53 (117.867)	-47.00 (13.887)	509.50 (247.011)	-42.62 (31.661)
	95% CI		-51.50, -42.49		-52.75, -32.49

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (20.0 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hpmas.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.8.1 Descriptive Statistics of 3-HPMA Urinary Concentration Adjusted for Creatinine (ng/mg creat) by Sex - FAS**

Timepoint	Statistic	THS 2.2			
		Male (N=39)		Female (N=41)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 4	n	39	39	40	40
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	267.59 (38.59%)		294.17 (50.93%)	
	Geometric 95% CI	237.15, 301.94		252.28, 343.00	
	Median	255.45	-66.43	307.03	-68.63
	Q25, Q75	192.58, 344.40	-72.13, -60.15	244.62, 377.21	-74.31, -60.93
	Min, Max	113.4, 597.0	-84.1, -37.5	33.7, 537.7	-83.8, 35.7
	Mean (SD)	286.54 (110.938)	-65.59 (11.460)	320.46 (116.067)	-62.37 (24.450)
	95% CI		-69.31, -61.87		-70.19, -54.54

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (20.0 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hpmas.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.8.1 Descriptive Statistics of 3-HPMA Urinary Concentration Adjusted for Creatinine (ng/mg creat) by Sex - FAS**

Timepoint	Statistic	THS 2.2			
		Male (N=39)		Female (N=41)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 5	n	39	39	40	40
	BLOQ - n (%)	0		1 (2.5%)	
	Geometric Mean (CV%)	386.70 (27.29%)		418.04 (55.84%)	
	Geometric 95% CI	354.52, 421.80		353.88, 493.83	
	Median	381.15	-54.41	428.77	-57.12
	Q25, Q75	298.99, 462.50	-58.85, -44.29	327.40, 531.01	-65.47, -46.13
	Min, Max	239.8, 683.6	-72.8, -23.0	36.8, 1261.8	-73.9, 44.6
	Mean (SD)	400.64 (110.083)	-51.26 (12.615)	462.45 (197.654)	-48.14 (26.394)
	95% CI		-55.35, -47.16		-56.58, -39.69

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (20.0 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hpmas.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.8.1 Descriptive Statistics of 3-HPMA Urinary Concentration Adjusted for Creatinine (ng/mg creat) by Sex - FAS**

Timepoint	Statistic	CC			
		Male (N=21)		Female (N=20)	
		Raw value	%Change (*)	Raw value	%Change (*)
Baseline	n	21		20	
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	746.14 (60.18%)		859.37 (30.45%)	
	Geometric 95% CI	579.32, 960.98		747.58, 987.88	
	Median	646.34		854.91	
	Q25, Q75	546.53, 1029.24		765.59, 1068.75	
	Min, Max	287.9, 2246.7		432.9, 1398.2	
	Mean (SD)	865.86 (505.664)		894.66 (252.731)	
	95% CI				

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (20.0 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hpmas.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.8.1 Descriptive Statistics of 3-HPMA Urinary Concentration Adjusted for Creatinine (ng/mg creat) by Sex - FAS**

Timepoint	Statistic	CC			
		Male (N=21)		Female (N=20)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 1	n	21	21	20	20
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	900.62 (51.87%)		1037.39 (34.65%)	
	Geometric 95% CI	721.20, 1124.67		886.17, 1214.42	
	Median	904.35	15.91	1001.96	16.22
	Q25, Q75	646.70, 1396.45	11.92, 43.62	905.63, 1299.80	4.63, 35.56
	Min, Max	427.8, 2102.3	-26.6, 114.1	469.1, 1863.6	-14.5, 103.3
	Mean (SD)	1006.98 (485.267)	25.00 (35.227)	1093.06 (360.421)	23.70 (29.347)
	95% CI		8.96, 41.04		9.96, 37.44

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (20.0 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hpmas.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.8.1 Descriptive Statistics of 3-HPMA Urinary Concentration Adjusted for Creatinine (ng/mg creat) by Sex - FAS**

Timepoint	Statistic	CC			
		Male (N=21)		Female (N=20)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 2	n	21	21	20	20
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	727.88 (62.62%)		842.71 (34.97%)	
	Geometric 95% CI	560.22, 945.73		718.85, 987.90	
	Median	587.93	0.44	820.18	-5.10
	Q25, Q75	467.10, 1153.85	-13.42, 15.64	648.34, 1026.80	-18.78, 17.57
	Min, Max	191.2, 1617.6	-67.8, 66.2	510.0, 1646.6	-29.7, 48.7
	Mean (SD)	841.90 (440.955)	2.80 (30.891)	891.88 (322.499)	0.50 (23.441)
	95% CI		-11.27, 16.86		-10.47, 11.48

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (20.0 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hpmas.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.8.1 Descriptive Statistics of 3-HPMA Urinary Concentration Adjusted for Creatinine (ng/mg creat) by Sex - FAS**

Timepoint	Statistic	CC			
		Male (N=21)		Female (N=20)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 3	n	21	21	20	20
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	837.94 (47.43%)		1009.47 (35.46%)	
	Geometric 95% CI	682.61, 1028.62		859.31, 1185.86	
	Median	771.68	13.29	1049.73	19.25
	Q25, Q75	583.00, 1107.44	0.09, 27.06	810.13, 1158.45	2.52, 38.65
	Min, Max	493.0, 1906.5	-46.4, 122.3	516.9, 1932.0	-57.9, 132.4
	Mean (SD)	928.43 (459.535)	17.16 (35.894)	1067.76 (374.452)	23.28 (38.155)
	95% CI		0.82, 33.50		5.42, 41.14

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (20.0 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hpmas.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.8.1 Descriptive Statistics of 3-HPMA Urinary Concentration Adjusted for Creatinine (ng/mg creat) by Sex - FAS**

Timepoint	Statistic	CC			
		Male (N=21)		Female (N=20)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 4	n	21	21	20	20
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	764.96 (59.00%)		819.72 (36.48%)	
	Geometric 95% CI	596.48, 981.04		694.76, 967.16	
	Median	642.03	4.97	921.33	-6.76
	Q25, Q75	491.03, 1232.52	-10.79, 21.05	703.62, 978.90	-18.66, 9.46
	Min, Max	318.5, 1806.9	-45.1, 62.3	379.6, 1403.7	-35.5, 50.2
	Mean (SD)	881.47 (482.181)	6.46 (28.928)	865.12 (269.943)	-2.45 (21.907)
	95% CI		-6.71, 19.64		-12.71, 7.81

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (20.0 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hpmas.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.8.1 Descriptive Statistics of 3-HPMA Urinary Concentration Adjusted for Creatinine (ng/mg creat) by Sex - FAS**

Timepoint	Statistic	CC			
		Male (N=21)		Female (N=20)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 5	n	21	21	20	20
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	889.78 (45.24%)		976.36 (33.06%)	
	Geometric 95% CI	731.13, 1082.86		839.74, 1135.20	
	Median	991.56	18.14	908.72	7.70
	Q25, Q75	641.79, 1187.82	2.26, 41.70	800.00, 1195.08	-4.83, 42.81
	Min, Max	451.6, 1803.3	-44.8, 115.1	578.2, 1896.6	-19.3, 69.8
	Mean (SD)	971.76 (419.839)	23.91 (35.202)	1027.50 (353.303)	16.46 (26.854)
	95% CI		7.89, 39.94		3.89, 29.03

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (20.0 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hpmas.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.8.1 Descriptive Statistics of 3-HPMA Urinary Concentration Adjusted for Creatinine (ng/mg creat) by Sex - FAS**

Timepoint	Statistic	SA			
		Male (N=20)		Female (N=19)	
		Raw value	%Change (*)	Raw value	%Change (*)
Baseline	n	20		19	
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	759.58 (45.42%)		984.49 (46.07%)	
	Geometric 95% CI	620.23, 930.23		796.86, 1216.28	
	Median	718.25		1048.39	
	Q25, Q75	537.90, 1040.47		700.82, 1340.50	
	Min, Max	376.6, 1618.3		352.7, 2530.7	
	Mean (SD)	830.57 (366.977)		1077.13 (487.229)	
	95% CI				

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (20.0 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hpmas.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.8.1 Descriptive Statistics of 3-HPMA Urinary Concentration Adjusted for Creatinine (ng/mg creat) by Sex - FAS**

Timepoint	Statistic	SA			
		Male (N=20)		Female (N=19)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 1	n	20	20	19	19
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	336.73 (28.70%)		401.94 (37.00%)	
	Geometric 95% CI	295.19, 384.12		338.21, 477.68	
	Median	346.96	-57.32	403.65	-57.06
	Q25, Q75	267.95, 422.91	-62.40, -48.97	326.57, 478.27	-70.24, -45.67
	Min, Max	212.8, 522.7	-72.3, -28.7	174.5, 794.7	-80.8, -24.2
	Mean (SD)	349.12 (92.981)	-54.15 (12.438)	426.34 (150.685)	-56.48 (15.534)
	95% CI		-59.98, -48.32		-63.97, -48.99

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (20.0 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hpmas.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.8.1 Descriptive Statistics of 3-HPMA Urinary Concentration Adjusted for Creatinine (ng/mg creat) by Sex - FAS**

Timepoint	Statistic	SA			
		Male (N=20)		Female (N=19)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 2	n	20	20	19	19
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	203.54 (20.28%)		202.07 (38.61%)	
	Geometric 95% CI	185.28, 223.59		168.83, 241.84	
	Median	204.17	-72.05	194.14	-77.40
	Q25, Q75	178.08, 232.15	-80.98, -62.18	177.64, 277.19	-83.80, -71.75
	Min, Max	118.6, 269.0	-85.3, -55.3	66.1, 351.1	-91.5, -60.4
	Mean (SD)	207.27 (38.966)	-71.51 (9.882)	214.10 (68.918)	-77.45 (9.220)
	95% CI		-76.14, -66.88		-81.90, -73.00

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (20.0 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hpmas.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.8.1 Descriptive Statistics of 3-HPMA Urinary Concentration Adjusted for Creatinine (ng/mg creat) by Sex - FAS**

Timepoint	Statistic	SA			
		Male (N=20)		Female (N=19)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 3	n	20	20	19	19
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	283.81 (18.72%)		301.00 (55.01%)	
	Geometric 95% CI	260.20, 309.56		234.93, 385.65	
	Median	270.55	-64.15	352.94	-65.62
	Q25, Q75	250.13, 307.74	-70.52, -51.77	253.21, 369.32	-75.23, -58.80
	Min, Max	221.1, 462.5	-79.8, -36.5	74.1, 532.2	-90.4, -47.8
	Mean (SD)	288.78 (58.898)	-60.60 (13.511)	332.52 (127.107)	-67.04 (11.644)
	95% CI		-66.93, -54.27		-72.65, -61.42

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (20.0 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hpmas.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.8.1 Descriptive Statistics of 3-HPMA Urinary Concentration Adjusted for Creatinine (ng/mg creat) by Sex - FAS**

Timepoint	Statistic	SA			
		Male (N=20)		Female (N=19)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 4	n	20	20	19	19
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	149.92 (31.45%)		154.87 (51.15%)	
	Geometric 95% CI	129.84, 173.10		122.76, 195.38	
	Median	156.77	-81.10	149.39	-81.89
	Q25, Q75	113.32, 185.62	-85.08, -72.43	111.50, 242.71	-87.98, -79.59
	Min, Max	89.2, 249.0	-89.3, -65.5	67.2, 354.1	-94.1, -66.2
	Mean (SD)	156.67 (47.271)	-79.14 (6.974)	172.30 (80.640)	-83.03 (6.341)
	95% CI		-82.41, -75.88		-86.09, -79.97

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (20.0 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hpmas.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.8.1 Descriptive Statistics of 3-HPMA Urinary Concentration Adjusted for Creatinine (ng/mg creat) by Sex - FAS**

Timepoint	Statistic	SA			
		Male (N=20)		Female (N=19)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 5	n	20	20	19	19
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	226.78 (21.91%)		267.29 (27.68%)	
	Geometric 95% CI	204.92, 250.97		234.49, 304.69	
	Median	230.20	-73.44	264.55	-72.54
	Q25, Q75	197.95, 258.91	-77.63, -57.49	216.32, 335.32	-78.68, -67.04
	Min, Max	126.9, 357.5	-84.4, -44.9	149.5, 436.5	-85.8, -44.5
	Mean (SD)	231.71 (48.463)	-67.83 (12.954)	276.65 (74.072)	-71.16 (10.434)
	95% CI		-73.89, -61.76		-76.19, -66.13

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (20.0 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hpmas.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.8.2 Descriptive Statistics of 3-HPMA Urinary Concentration Adjusted for Creatinine (ng/mg creat) by Cigarette Consumption - FAS**

Timepoint	Statistic	THS 2.2			
		10-19 cig/day (N=41)		>19 cig/day (N=39)	
		Raw value	%Change (*)	Raw value	%Change (*)
Baseline	n	41		39	
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	705.04 (45.56%)		1014.37 (64.37%)	
	Geometric 95% CI	614.72, 808.62		838.13, 1227.67	
	Median	726.54		1149.17	
	Q25, Q75	574.24, 859.00		764.76, 1495.48	
	Min, Max	254.5, 1654.0		94.6, 2495.7	
	Mean (SD)	770.07 (328.066)		1153.81 (512.917)	
	95% CI				

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (20.0 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hpmac.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.8.2 Descriptive Statistics of 3-HPMA Urinary Concentration Adjusted for Creatinine (ng/mg creat) by Cigarette Consumption - FAS**

Timepoint	Statistic	THS 2.2			
		10-19 cig/day (N=41)		>19 cig/day (N=39)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 1	n	41	41	39	39
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	421.08 (32.02%)		532.40 (43.32%)	
	Geometric 95% CI	381.54, 464.72		465.43, 609.00	
	Median	416.07	-42.83	561.58	-49.42
	Q25, Q75	358.60, 531.25	-51.57, -27.97	434.16, 677.42	-58.69, -40.16
	Min, Max	188.7, 747.8	-65.8, 77.1	82.2, 953.7	-70.1, 28.6
	Mean (SD)	440.80 (132.449)	-36.46 (26.435)	568.45 (180.644)	-43.71 (23.603)
	95% CI		-44.80, -28.11		-51.36, -36.05

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (20.0 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hpmac.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.8.2 Descriptive Statistics of 3-HPMA Urinary Concentration Adjusted for Creatinine (ng/mg creat) by Cigarette Consumption - FAS**

Timepoint	Statistic	THS 2.2			
		10-19 cig/day (N=41)		>19 cig/day (N=39)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 2	n	41	41	39	39
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	292.54 (41.03%)		343.12 (45.90%)	
	Geometric 95% CI	258.29, 331.33		297.78, 395.37	
	Median	292.35	-59.95	369.32	-68.13
	Q25, Q75	237.74, 356.72	-65.18, -51.74	314.56, 416.31	-73.56, -54.39
	Min, Max	124.7, 730.6	-78.7, 13.1	53.7, 639.8	-86.3, -24.6
	Mean (SD)	315.91 (132.321)	-56.11 (16.860)	368.05 (115.343)	-63.73 (13.864)
	95% CI		-61.44, -50.78		-68.23, -59.24

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (20.0 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hpmac.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.8.2 Descriptive Statistics of 3-HPMA Urinary Concentration Adjusted for Creatinine (ng/mg creat) by Cigarette Consumption - FAS**

Timepoint	Statistic	THS 2.2			
		10-19 cig/day (N=41)		>19 cig/day (N=39)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 3	n	40	40	39	39
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	410.87 (30.26%)		468.59 (54.54%)	
	Geometric 95% CI	373.76, 451.66		397.15, 552.88	
	Median	402.82	-46.67	472.51	-55.00
	Q25, Q75	329.05, 505.34	-54.45, -36.98	415.05, 562.17	-65.01, -43.17
	Min, Max	263.2, 845.1	-66.6, 83.6	38.8, 1671.4	-83.4, 20.2
	Mean (SD)	429.67 (137.868)	-39.30 (27.214)	516.41 (236.679)	-50.41 (20.185)
	95% CI		-48.01, -30.59		-56.95, -43.86

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (20.0 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hpmac.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.8.2 Descriptive Statistics of 3-HPMA Urinary Concentration Adjusted for Creatinine (ng/mg creat) by Cigarette Consumption - FAS**

Timepoint	Statistic	THS 2.2			
		10-19 cig/day (N=41)		>19 cig/day (N=39)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 4	n	40	40	39	39
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	262.01 (36.50%)		301.32 (52.29%)	
	Geometric 95% CI	233.99, 293.39		256.94, 353.38	
	Median	260.44	-66.57	319.38	-70.65
	Q25, Q75	202.49, 331.24	-71.83, -58.14	237.06, 405.89	-78.23, -63.16
	Min, Max	113.4, 597.0	-80.2, 35.7	33.7, 537.7	-84.1, -24.2
	Mean (SD)	278.90 (105.304)	-59.97 (23.067)	329.17 (118.478)	-68.04 (13.051)
	95% CI		-67.35, -52.59		-72.28, -63.81

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (20.0 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hpmac.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.8.2 Descriptive Statistics of 3-HPMA Urinary Concentration Adjusted for Creatinine (ng/mg creat) by Cigarette Consumption - FAS**

Timepoint	Statistic	THS 2.2			
		10-19 cig/day (N=41)		>19 cig/day (N=39)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 5	n	40	40	39	39
	BLOQ - n (%)	0		1 (2.6%)	
	Geometric Mean (CV%)	368.49 (26.88%)		440.12 (55.15%)	
	Geometric 95% CI	338.65, 400.97		372.40, 520.14	
	Median	362.60	-50.59	462.50	-57.61
	Q25, Q75	299.66, 424.91	-58.05, -40.50	381.15, 567.30	-68.35, -47.02
	Min, Max	239.8, 683.6	-73.9, 44.6	36.8, 1261.8	-73.7, 16.8
	Mean (SD)	381.73 (107.429)	-45.64 (22.353)	483.43 (192.302)	-53.82 (18.208)
	95% CI		-52.79, -38.49		-59.73, -47.92

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (20.0 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hpmac.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.8.2 Descriptive Statistics of 3-HPMA Urinary Concentration Adjusted for Creatinine (ng/mg creat) by Cigarette Consumption - FAS**

Timepoint	Statistic	CC			
		10-19 cig/day (N=21)		>19 cig/day (N=20)	
		Raw value	%Change (*)	Raw value	%Change (*)
Baseline	n	21		20	
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	720.31 (42.95%)		891.75 (49.86%)	
	Geometric 95% CI	597.29, 868.66		715.29, 1111.75	
	Median	736.66		879.61	
	Q25, Q75	546.53, 1029.24		711.47, 1139.12	
	Min, Max	347.1, 1398.2		287.9, 2246.7	
	Mean (SD)	779.12 (312.010)		985.74 (456.050)	
	95% CI				

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (20.0 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hpmac.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.8.2 Descriptive Statistics of 3-HPMA Urinary Concentration Adjusted for Creatinine (ng/mg creat) by Cigarette Consumption - FAS**

Timepoint	Statistic	CC			
		10-19 cig/day (N=21)		>19 cig/day (N=20)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 1	n	21	21	20	20
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	922.78 (43.56%)		1011.25 (45.41%)	
	Geometric 95% CI	763.31, 1115.55		825.75, 1238.42	
	Median	904.35	22.76	1018.79	12.97
	Q25, Q75	699.45, 1341.47	13.30, 43.62	847.86, 1413.56	1.38, 34.51
	Min, Max	427.8, 1863.6	-21.7, 114.1	437.9, 2102.3	-26.6, 52.1
	Mean (SD)	1000.45 (408.533)	32.82 (37.965)	1099.91 (448.061)	15.49 (22.120)
	95% CI		15.53, 50.10		5.13, 25.85

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (20.0 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hpmac.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.8.2 Descriptive Statistics of 3-HPMA Urinary Concentration Adjusted for Creatinine (ng/mg creat) by Cigarette Consumption - FAS**

Timepoint	Statistic	CC			
		10-19 cig/day (N=21)		>19 cig/day (N=20)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 2	n	21	21	20	20
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	706.75 (52.05%)		869.18 (47.03%)	
	Geometric 95% CI	565.56, 883.20		705.12, 1071.41	
	Median	625.97	-2.52	899.72	-1.72
	Q25, Q75	539.29, 1123.94	-13.83, 15.64	648.34, 1199.47	-18.70, 16.22
	Min, Max	191.2, 1459.3	-67.8, 66.2	393.0, 1646.6	-43.4, 36.5
	Mean (SD)	784.72 (352.281)	3.09 (30.746)	951.93 (405.364)	0.20 (23.606)
	95% CI		-10.91, 17.09		-10.85, 11.25

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (20.0 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hpmac.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.8.2 Descriptive Statistics of 3-HPMA Urinary Concentration Adjusted for Creatinine (ng/mg creat) by Cigarette Consumption - FAS**

Timepoint	Statistic	CC			
		10-19 cig/day (N=21)		>19 cig/day (N=20)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 3	n	21	21	20	20
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	855.66 (41.74%)		987.52 (43.13%)	
	Geometric 95% CI	713.00, 1026.87		813.96, 1198.10	
	Median	782.61	19.69	1057.32	14.50
	Q25, Q75	634.24, 1062.13	3.80, 38.18	672.32, 1204.76	0.49, 27.66
	Min, Max	503.3, 1932.0	-46.4, 132.4	493.0, 1906.5	-57.9, 85.4
	Mean (SD)	928.47 (415.862)	24.45 (40.801)	1067.71 (424.818)	15.62 (32.216)
	95% CI		5.87, 43.03		0.54, 30.71

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (20.0 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hpmac.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.8.2 Descriptive Statistics of 3-HPMA Urinary Concentration Adjusted for Creatinine (ng/mg creat) by Cigarette Consumption - FAS**

Timepoint	Statistic	CC			
		10-19 cig/day (N=21)		>19 cig/day (N=20)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 4	n	21	21	20	20
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	720.07 (48.50%)		873.47 (46.99%)	
	Geometric 95% CI	584.14, 887.61		708.71, 1076.53	
	Median	762.26	-0.67	940.63	-3.52
	Q25, Q75	491.03, 993.17	-11.58, 12.01	631.17, 1157.74	-16.92, 15.84
	Min, Max	318.5, 1492.9	-41.7, 62.3	403.6, 1806.9	-45.1, 50.2
	Mean (SD)	793.91 (352.862)	3.25 (27.145)	957.07 (415.127)	0.92 (25.001)
	95% CI		-9.11, 15.61		-10.78, 12.63

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (20.0 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hpmac.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.8.2 Descriptive Statistics of 3-HPMA Urinary Concentration Adjusted for Creatinine (ng/mg creat) by Cigarette Consumption - FAS**

Timepoint	Statistic	CC			
		10-19 cig/day (N=21)		>19 cig/day (N=20)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 5	n	21	21	20	20
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	850.01 (39.16%)		1024.38 (38.08%)	
	Geometric 95% CI	715.75, 1009.45		862.33, 1216.89	
	Median	746.63	7.99	1063.81	18.85
	Q25, Q75	646.20, 1187.82	-1.19, 47.18	829.72, 1206.24	-1.41, 37.69
	Min, Max	489.1, 1896.6	-19.3, 115.1	451.6, 1803.3	-44.8, 62.4
	Mean (SD)	912.23 (369.474)	22.32 (35.359)	1090.00 (388.982)	18.13 (27.017)
	95% CI		6.22, 38.42		5.48, 30.78

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (20.0 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hpmac.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.8.2 Descriptive Statistics of 3-HPMA Urinary Concentration Adjusted for Creatinine (ng/mg creat) by Cigarette Consumption - FAS**

Timepoint	Statistic	SA			
		10-19 cig/day (N=19)		>19 cig/day (N=20)	
		Raw value	%Change (*)	Raw value	%Change (*)
Baseline	n	19		20	
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	672.34 (39.32%)		1091.22 (39.92%)	
	Geometric 95% CI	560.05, 807.15		911.49, 1306.38	
	Median	674.60		1053.53	
	Q25, Q75	482.72, 910.87		837.32, 1451.39	
	Min, Max	352.7, 1271.9		481.7, 2530.7	
	Mean (SD)	719.50 (275.243)		1170.32 (463.411)	
	95% CI				

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (20.0 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hpmac.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.8.2 Descriptive Statistics of 3-HPMA Urinary Concentration Adjusted for Creatinine (ng/mg creat) by Cigarette Consumption - FAS**

Timepoint	Statistic	SA			
		10-19 cig/day (N=19)		>19 cig/day (N=20)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 1	n	19	19	20	20
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	328.56 (29.51%)		407.80 (34.71%)	
	Geometric 95% CI	285.85, 377.65		348.25, 477.53	
	Median	333.80	-50.52	410.33	-61.59
	Q25, Q75	239.55, 439.19	-59.58, -42.58	346.96, 477.97	-71.35, -53.52
	Min, Max	212.8, 491.6	-64.7, -28.7	174.5, 794.7	-80.8, -24.2
	Mean (SD)	341.31 (93.320)	-50.02 (10.970)	429.90 (144.706)	-60.29 (14.766)
	95% CI		-55.31, -44.73		-67.20, -53.37

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (20.0 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hpmac.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.8.2 Descriptive Statistics of 3-HPMA Urinary Concentration Adjusted for Creatinine (ng/mg creat) by Cigarette Consumption - FAS**

Timepoint	Statistic	SA			
		10-19 cig/day (N=19)		>19 cig/day (N=20)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 2	n	19	19	20	20
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	193.17 (23.63%)		212.43 (35.05%)	
	Geometric 95% CI	172.64, 216.14		181.14, 249.12	
	Median	192.73	-71.31	226.26	-78.90
	Q25, Q75	172.64, 210.23	-77.41, -61.66	186.65, 265.70	-84.39, -73.95
	Min, Max	118.6, 351.1	-83.8, -55.3	66.1, 316.6	-91.5, -59.4
	Mean (SD)	198.40 (49.908)	-70.05 (8.556)	222.19 (58.240)	-78.54 (9.484)
	95% CI		-74.18, -65.92		-82.98, -74.10

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (20.0 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hpmac.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.8.2 Descriptive Statistics of 3-HPMA Urinary Concentration Adjusted for Creatinine (ng/mg creat) by Cigarette Consumption - FAS**

Timepoint	Statistic	SA			
		10-19 cig/day (N=19)		>19 cig/day (N=20)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 3	n	19	19	20	20
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	272.04 (36.91%)		312.44 (41.07%)	
	Geometric 95% CI	228.99, 323.17		259.74, 375.85	
	Median	253.21	-60.99	327.81	-70.45
	Q25, Q75	232.71, 353.77	-69.84, -47.82	276.31, 367.16	-75.64, -62.41
	Min, Max	106.4, 532.2	-73.7, -36.5	74.1, 525.1	-90.4, -36.8
	Mean (SD)	288.28 (100.888)	-58.07 (11.621)	330.80 (95.846)	-69.12 (11.901)
	95% CI		-63.67, -52.46		-74.70, -63.55

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (20.0 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hpmac.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.8.2 Descriptive Statistics of 3-HPMA Urinary Concentration Adjusted for Creatinine (ng/mg creat) by Cigarette Consumption - FAS**

Timepoint	Statistic	SA			
		10-19 cig/day (N=19)		>19 cig/day (N=20)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 4	n	19	19	20	20
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	133.26 (37.02%)		172.93 (41.45%)	
	Geometric 95% CI	112.11, 158.38		143.53, 208.36	
	Median	132.47	-80.77	179.39	-82.37
	Q25, Q75	107.23, 183.90	-83.53, -72.74	132.68, 241.77	-88.25, -79.50
	Min, Max	73.8, 278.6	-90.3, -65.5	67.2, 354.1	-94.1, -66.2
	Mean (SD)	141.73 (52.668)	-79.29 (6.268)	185.71 (70.005)	-82.70 (7.161)
	95% CI		-82.32, -76.26		-86.06, -79.34

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (20.0 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hpmac.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.8.2 Descriptive Statistics of 3-HPMA Urinary Concentration Adjusted for Creatinine (ng/mg creat) by Cigarette Consumption - FAS**

Timepoint	Statistic	SA			
		10-19 cig/day (N=19)		>19 cig/day (N=20)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 5	n	19	19	20	20
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	226.52 (27.06%)		265.40 (22.84%)	
	Geometric 95% CI	199.28, 257.48		238.81, 294.94	
	Median	216.33	-67.05	268.63	-75.04
	Q25, Q75	196.42, 252.37	-74.99, -52.00	232.55, 310.19	-79.88, -71.85
	Min, Max	126.9, 436.5	-84.4, -44.5	149.5, 385.4	-85.8, -44.9
	Mean (SD)	234.58 (68.204)	-64.31 (12.076)	271.67 (58.907)	-74.34 (9.321)
	95% CI		-70.13, -58.48		-78.71, -69.97

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (20.0 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hpmac.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.9 Descriptive Statistics of S-PMA Urinary Concentration Adjusted for Creatinine (pg/mg creat) - FAS**

Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
		Raw value	%Change (*)	Raw value	%Change (*)
Baseline	n	80		41	
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	2394.03 (102.40%)		2765.20 (77.42%)	
	Geometric 95% CI	1982.83, 2890.50		2227.38, 3432.88	
	Median	2582.08		2903.85	
	Q25, Q75	1489.86, 4439.96		2018.73, 3794.75	
	Min, Max	395.9, 11111.1		357.1, 8830.3	
	Mean (SD)	3225.58 (2306.758)		3399.93 (2200.241)	
	95% CI				

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.0250 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_spma.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.9 Descriptive Statistics of S-PMA Urinary Concentration Adjusted for Creatinine (pg/mg creat) - FAS**

Timepoint	Statistic	SA (N=39)	
		Raw value	%Change (*)
Baseline	n	39	
	BLOQ - n (%)	0	
	Geometric Mean (CV%)	2153.82 (116.41%)	
	Geometric 95% CI	1595.57, 2907.38	
	Median	2251.48	
	Q25, Q75	939.80, 4473.12	
	Min, Max	336.7, 9539.9	
	Mean (SD)	3064.10 (2368.257)	
	95% CI		

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.0250 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_spma.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.9 Descriptive Statistics of S-PMA Urinary Concentration Adjusted for Creatinine (pg/mg creat) - FAS**

Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 1	n	80	80	41	41
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	1024.34 (114.28%)		2874.73 (75.34%)	
	Geometric 95% CI	835.80, 1255.41		2326.43, 3552.26	
	Median	1252.76	-57.86	2934.58	7.13
	Q25, Q75	619.47, 1943.10	-64.42, -46.99	2208.06, 3861.39	-6.97, 12.61
	Min, Max	130.5, 4554.9	-81.0, -14.6	390.7, 10305.2	-53.4, 123.8
	Mean (SD)	1429.50 (1048.096)	-55.62 (11.952)	3511.57 (2277.853)	6.93 (27.319)
	95% CI		-58.28, -52.96		-1.70, 15.56

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.0250 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_spma.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.9 Descriptive Statistics of S-PMA Urinary Concentration Adjusted for Creatinine (pg/mg creat) - FAS**

Timepoint	Statistic	SA (N=39)	
		Raw value	%Change (*)
Day 1	n	39	39
	BLOQ - n (%)	0	
	Geometric Mean (CV%)	953.37 (132.83%)	
	Geometric 95% CI	687.54, 1321.98	
	Median	1367.43	-53.77
	Q25, Q75	439.70, 1983.25	-64.81, -45.70
	Min, Max	161.0, 5357.1	-79.3, -26.6
	Mean (SD)	1438.98 (1225.088)	-53.92 (12.897)
	95% CI		-58.10, -49.73

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.0250 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_spma.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.3.9 Descriptive Statistics of S-PMA Urinary Concentration Adjusted for Creatinine (pg/mg creat) - FAS**

Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 2	n	80	80	41	41
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	518.55 (93.26%)		2667.74 (83.51%)	
	Geometric 95% CI	434.84, 618.37		2120.50, 3356.20	
	Median	597.58	-79.31	2701.30	-4.85
	Q25, Q75	303.30, 859.05	-84.48, -73.20	2076.50, 3924.53	-16.22, 20.17
	Min, Max	96.5, 3355.4	-90.5, -1.9	299.8, 11642.4	-78.0, 88.6
	Mean (SD)	690.50 (552.190)	-75.73 (14.618)	3343.30 (2287.854)	2.25 (32.267)
	95% CI		-78.98, -72.47		-7.94, 12.44

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.0250 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_spma.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.9 Descriptive Statistics of S-PMA Urinary Concentration Adjusted for Creatinine (pg/mg creat) - FAS**

Timepoint	Statistic	SA (N=39)	
		Raw value	%Change (*)
Day 2	n	39	39
	BLOQ - n (%)	0	
	Geometric Mean (CV%)	450.61 (82.26%)	
	Geometric 95% CI	356.94, 568.86	
	Median	443.62	-79.87
	Q25, Q75	267.35, 681.07	-86.59, -70.61
	Min, Max	104.7, 1629.8	-91.5, -42.1
	Mean (SD)	573.49 (409.796)	-76.43 (12.574)
	95% CI		-80.51, -72.35

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.0250 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_spma.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.9 Descriptive Statistics of S-PMA Urinary Concentration Adjusted for Creatinine (pg/mg creat) - FAS**

Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 3	n	79	79	41	41
	BLOQ - n (%)	1 (1.3%)		0	
	Geometric Mean (CV%)	276.88 (103.50%)		2691.34 (73.96%)	
	Geometric 95% CI	228.70, 335.20		2184.88, 3315.21	
	Median	291.67	-89.71	2877.25	4.63
	Q25, Q75	159.43, 431.21	-91.15, -86.08	2085.76, 3590.82	-14.64, 24.19
	Min, Max	43.1, 10732.9	-95.5, 25.7	285.3, 9225.0	-89.2, 139.1
	Mean (SD)	474.16 (1198.169)	-86.71 (13.679)	3227.45 (1926.536)	5.51 (37.198)
	95% CI		-89.78, -83.64		-6.23, 17.26

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.0250 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_spma.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.9 Descriptive Statistics of S-PMA Urinary Concentration Adjusted for Creatinine (pg/mg creat) - FAS**

Timepoint	Statistic	SA (N=39)	
		Raw value	%Change (*)
Day 3	n	39	39
	BLOQ - n (%)	0	
	Geometric Mean (CV%)	224.24 (83.83%)	
	Geometric 95% CI	177.01, 284.08	
	Median	210.08	-89.61
	Q25, Q75	124.40, 372.52	-92.81, -84.76
	Min, Max	52.0, 1149.4	-96.2, -66.5
	Mean (SD)	292.23 (234.786)	-88.10 (6.553)
	95% CI		-90.23, -85.97

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.0250 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_spma.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.9 Descriptive Statistics of S-PMA Urinary Concentration Adjusted for Creatinine (pg/mg creat) - FAS**

Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 4	n	79	79	41	41
	BLOQ - n (%)	1 (1.3%)		0	
	Geometric Mean (CV%)	188.72 (74.72%)		2897.36 (73.70%)	
	Geometric 95% CI	162.56, 219.09		2353.53, 3566.87	
	Median	203.43	-92.47	2892.38	5.90
	Q25, Q75	116.85, 282.26	-94.51, -90.35	1959.33, 4189.61	-11.33, 22.77
	Min, Max	36.3, 1244.9	-97.3, -76.2	340.5, 9258.1	-67.5, 126.2
	Mean (SD)	235.17 (180.582)	-91.26 (4.733)	3507.14 (2190.484)	9.62 (33.161)
	95% CI		-92.32, -90.19		-0.85, 20.09

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.0250 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_spma.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.9 Descriptive Statistics of S-PMA Urinary Concentration Adjusted for Creatinine (pg/mg creat) - FAS**

Timepoint	Statistic	SA (N=39)	
		Raw value	%Change (*)
Day 4	n	39	39
	BLOQ - n (%)	0	
	Geometric Mean (CV%)	158.09 (75.09%)	
	Geometric 95% CI	127.27, 196.35	
	Median	142.35	-92.22
	Q25, Q75	96.50, 269.34	-95.33, -90.33
	Min, Max	54.2, 690.5	-97.7, -70.5
	Mean (SD)	198.15 (146.040)	-91.39 (5.390)
	95% CI		-93.14, -89.63

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.0250 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_spma.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.9 Descriptive Statistics of S-PMA Urinary Concentration Adjusted for Creatinine (pg/mg creat) - FAS**

Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 5	n	79	79	41	41
	BLOQ - n (%)	3 (3.8%)		0	
	Geometric Mean (CV%)	164.45 (63.10%)		2922.81 (75.82%)	
	Geometric 95% CI	144.45, 187.22		2362.80, 3615.54	
	Median	159.33	-93.30	2981.82	4.24
	Q25, Q75	113.29, 237.12	-95.28, -91.56	2294.64, 4431.95	-8.99, 21.22
	Min, Max	43.9, 873.8	-97.7, -75.3	352.0, 9345.5	-46.4, 127.5
	Mean (SD)	194.16 (126.312)	-92.03 (5.222)	3548.31 (2162.396)	9.48 (31.561)
	95% CI		-93.20, -90.85		-0.48, 19.45

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.0250 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_spma.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.9 Descriptive Statistics of S-PMA Urinary Concentration Adjusted for Creatinine (pg/mg creat) - FAS**

Timepoint	Statistic	SA (N=39)	
		Raw value	%Change (*)
Day 5	n	39	39
	BLOQ - n (%)	0	
	Geometric Mean (CV%)	143.70 (53.43%)	
	Geometric 95% CI	122.15, 169.04	
	Median	142.29	-93.82
	Q25, Q75	95.23, 171.04	-96.31, -88.82
	Min, Max	67.2, 509.0	-97.9, -70.9
	Mean (SD)	164.41 (98.838)	-91.52 (6.754)
	95% CI		-93.72, -89.33

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.0250 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_spma.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.9.1 Descriptive Statistics of S-PMA Urinary Concentration Adjusted for Creatinine (pg/mg creat) by Sex - FAS**

Timepoint	Statistic	THS 2.2			
		Male (N=39)		Female (N=41)	
		Raw value	%Change (*)	Raw value	%Change (*)
Baseline	n	39		41	
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	1980.64 (101.19%)		2867.08 (98.51%)	
	Geometric 95% CI	1508.69, 2600.22		2210.81, 3718.16	
	Median	2293.68		3409.09	
	Q25, Q75	1159.93, 3403.98		1824.10, 5510.89	
	Min, Max	406.2, 7671.6		395.9, 11111.1	
	Mean (SD)	2652.55 (1915.308)		3770.65 (2529.736)	
	95% CI				

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.0250 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_spmas.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.9.1 Descriptive Statistics of S-PMA Urinary Concentration Adjusted for Creatinine (pg/mg creat) by Sex - FAS**

Timepoint	Statistic	THS 2.2			
		Male (N=39)		Female (N=41)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 1	n	39	39	41	41
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	769.62 (111.81%)		1344.49 (103.10%)	
	Geometric 95% CI	574.77, 1030.52		1027.79, 1758.77	
	Median	908.35	-60.08	1669.15	-52.67
	Q25, Q75	345.22, 1444.91	-66.24, -52.91	1072.82, 2323.95	-62.29, -43.52
	Min, Max	130.5, 3612.1	-81.0, -40.8	149.0, 4554.9	-75.8, -14.6
	Mean (SD)	1078.87 (845.385)	-60.04 (9.153)	1763.02 (1121.035)	-51.42 (12.860)
	95% CI		-63.01, -57.07		-55.48, -47.35

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.0250 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_spmas.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.9.1 Descriptive Statistics of S-PMA Urinary Concentration Adjusted for Creatinine (pg/mg creat) by Sex - FAS**

Timepoint	Statistic	THS 2.2			
		Male (N=39)		Female (N=41)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 2	n	39	39	41	41
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	414.32 (94.54%)		641.94 (83.96%)	
	Geometric 95% CI	319.76, 536.83		509.76, 808.39	
	Median	421.78	-79.32	698.15	-78.96
	Q25, Q75	239.92, 711.68	-85.30, -74.05	445.42, 1051.43	-82.92, -72.59
	Min, Max	96.5, 3355.4	-90.5, -1.9	105.8, 2245.7	-88.8, -27.3
	Mean (SD)	576.85 (592.311)	-75.80 (17.190)	798.61 (494.307)	-75.65 (11.887)
	95% CI		-81.38, -70.23		-79.41, -71.89

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.0250 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_spmas.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.9.1 Descriptive Statistics of S-PMA Urinary Concentration Adjusted for Creatinine (pg/mg creat) by Sex - FAS**

Timepoint	Statistic	THS 2.2			
		Male (N=39)		Female (N=41)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 3	n	39	39	40	40
	BLOQ - n (%)	1 (2.6%)		0	
	Geometric Mean (CV%)	193.73 (72.29%)		392.19 (110.17%)	
	Geometric 95% CI	157.00, 239.05		294.90, 521.58	
	Median	206.32	-90.42	401.23	-87.70
	Q25, Q75	121.57, 326.81	-91.70, -88.17	253.47, 583.89	-90.12, -84.66
	Min, Max	43.1, 555.8	-95.5, -77.7	63.6, 10732.9	-93.0, 25.7
	Mean (SD)	232.76 (133.672)	-89.59 (3.900)	709.52 (1654.906)	-83.90 (18.521)
	95% CI		-90.86, -88.33		-89.83, -77.98

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.0250 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_spmas.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.9.1 Descriptive Statistics of S-PMA Urinary Concentration Adjusted for Creatinine (pg/mg creat) by Sex - FAS**

Timepoint	Statistic	THS 2.2			
		Male (N=39)		Female (N=41)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 4	n	39	39	40	40
	BLOQ - n (%)	1 (2.6%)		0	
	Geometric Mean (CV%)	137.74 (61.17%)		256.54 (68.17%)	
	Geometric 95% CI	114.73, 165.37		210.54, 312.58	
	Median	148.77	-93.01	258.97	-91.73
	Q25, Q75	95.74, 215.52	-95.45, -91.52	185.27, 380.89	-93.80, -89.50
	Min, Max	36.3, 353.7	-97.3, -77.9	67.6, 1244.9	-95.5, -76.2
	Mean (SD)	158.54 (80.289)	-92.11 (4.515)	309.88 (217.583)	-90.43 (4.848)
	95% CI		-93.58, -90.64		-91.98, -88.87

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.0250 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_spmas.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.9.1 Descriptive Statistics of S-PMA Urinary Concentration Adjusted for Creatinine (pg/mg creat) by Sex - FAS**

Timepoint	Statistic	THS 2.2			
		Male (N=39)		Female (N=41)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 5	n	39	39	40	40
	BLOQ - n (%)	1 (2.6%)		2 (5.0%)	
	Geometric Mean (CV%)	125.81 (47.57%)		213.52 (62.51%)	
	Geometric 95% CI	108.67, 145.65		177.69, 256.57	
	Median	137.31	-93.81	227.32	-93.05
	Q25, Q75	94.82, 172.72	-95.86, -92.05	153.14, 294.18	-95.13, -90.66
	Min, Max	43.9, 305.2	-97.7, -75.6	46.0, 873.8	-97.7, -75.3
	Mean (SD)	138.13 (58.826)	-92.54 (4.893)	248.78 (149.459)	-91.53 (5.539)
	95% CI		-94.13, -90.95		-93.30, -89.75

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.0250 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_spmas.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.9.1 Descriptive Statistics of S-PMA Urinary Concentration Adjusted for Creatinine (pg/mg creat) by Sex - FAS**

Timepoint	Statistic	CC			
		Male (N=21)		Female (N=20)	
		Raw value	%Change (*)	Raw value	%Change (*)
Baseline	n	21		20	
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	2593.23 (75.24%)		2958.06 (81.27%)	
	Geometric 95% CI	1911.82, 3517.49		2119.69, 4128.03	
	Median	2681.74		3066.62	
	Q25, Q75	1340.57, 3727.17		2225.91, 5062.50	
	Min, Max	663.4, 8830.3		357.1, 8760.5	
	Mean (SD)	3206.63 (2260.616)		3602.88 (2174.327)	
	95% CI				

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.0250 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_spmas.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.9.1 Descriptive Statistics of S-PMA Urinary Concentration Adjusted for Creatinine (pg/mg creat) by Sex - FAS**

Timepoint	Statistic	CC			
		Male (N=21)		Female (N=20)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 1	n	21	21	20	20
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	2729.83 (66.16%)		3035.17 (86.46%)	
	Geometric 95% CI	2075.03, 3591.25		2139.58, 4305.64	
	Median	2996.43	8.29	2825.67	5.12
	Q25, Q75	1782.60, 3861.39	-2.20, 12.65	2220.10, 4997.24	-8.44, 11.59
	Min, Max	735.3, 8636.4	-53.4, 123.8	390.7, 10305.2	-23.6, 35.0
	Mean (SD)	3215.77 (1925.272)	10.08 (35.504)	3822.16 (2612.161)	3.62 (14.896)
	95% CI		-6.08, 26.25		-3.35, 10.60

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.0250 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_spmas.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.9.1 Descriptive Statistics of S-PMA Urinary Concentration Adjusted for Creatinine (pg/mg creat) by Sex - FAS**

Timepoint	Statistic	CC			
		Male (N=21)		Female (N=20)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 2	n	21	21	20	20
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	2413.89 (76.22%)		2963.05 (91.58%)	
	Geometric 95% CI	1773.90, 3284.79		2056.48, 4269.28	
	Median	2500.00	-7.41	2941.78	-0.15
	Q25, Q75	1610.90, 3768.12	-21.12, 21.24	2308.88, 4760.00	-14.32, 16.94
	Min, Max	498.7, 8175.9	-78.0, 88.6	299.8, 11642.4	-46.1, 51.7
	Mean (SD)	2941.05 (1863.003)	1.60 (39.069)	3765.66 (2645.650)	2.93 (24.171)
	95% CI		-16.19, 19.39		-8.38, 14.25

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.0250 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_spmas.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.9.1 Descriptive Statistics of S-PMA Urinary Concentration Adjusted for Creatinine (pg/mg creat) by Sex - FAS**

Timepoint	Statistic	CC			
		Male (N=21)		Female (N=20)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 3	n	21	21	20	20
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	2608.53 (61.62%)		2781.12 (88.41%)	
	Geometric 95% CI	2014.90, 3377.05		1948.76, 3969.00	
	Median	2865.25	6.42	2975.15	4.22
	Q25, Q75	1902.65, 3476.96	-15.05, 24.46	2197.70, 4035.98	-5.74, 18.24
	Min, Max	706.0, 9225.0	-59.0, 139.1	285.3, 7984.5	-89.2, 58.2
	Mean (SD)	3021.34 (1783.612)	7.56 (41.584)	3443.86 (2090.170)	3.36 (32.912)
	95% CI		-11.37, 26.50		-12.05, 18.77

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.0250 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_spmas.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.9.1 Descriptive Statistics of S-PMA Urinary Concentration Adjusted for Creatinine (pg/mg creat) by Sex - FAS**

Timepoint	Statistic	CC			
		Male (N=21)		Female (N=20)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 4	n	21	21	20	20
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	2822.02 (62.16%)		2978.64 (87.43%)	
	Geometric 95% CI	2175.54, 3660.61		2093.46, 4238.09	
	Median	2798.35	11.12	3273.20	5.79
	Q25, Q75	1853.51, 4141.74	-9.94, 33.59	1983.38, 4986.05	-16.49, 12.54
	Min, Max	952.6, 9148.7	-67.5, 126.2	340.5, 9258.1	-27.0, 57.0
	Mean (SD)	3306.44 (2041.116)	16.34 (41.240)	3717.88 (2371.670)	2.57 (20.557)
	95% CI		-2.44, 35.12		-7.06, 12.20

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.0250 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_spmas.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.9.1 Descriptive Statistics of S-PMA Urinary Concentration Adjusted for Creatinine (pg/mg creat) by Sex - FAS**

Timepoint	Statistic	CC			
		Male (N=21)		Female (N=20)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 5	n	21	21	20	20
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	2858.87 (68.79%)		2991.48 (85.34%)	
	Geometric 95% CI	2153.56, 3795.17		2116.12, 4228.94	
	Median	2981.82	5.83	3021.90	3.70
	Q25, Q75	1950.43, 4431.95	-3.56, 31.45	2415.87, 5347.44	-11.12, 13.69
	Min, Max	844.9, 9345.5	-46.4, 127.5	352.0, 9050.4	-27.2, 25.0
	Mean (SD)	3414.59 (2126.739)	16.33 (40.737)	3688.71 (2245.558)	2.29 (15.618)
	95% CI		-2.21, 34.88		-5.02, 9.61

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.0250 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_spmas.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.9.1 Descriptive Statistics of S-PMA Urinary Concentration Adjusted for Creatinine (pg/mg creat) by Sex - FAS**

Timepoint	Statistic	SA			
		Male (N=20)		Female (N=19)	
		Raw value	%Change (*)	Raw value	%Change (*)
Baseline	n	20		19	
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	1673.13 (118.23%)		2809.70 (104.66%)	
	Geometric 95% CI	1080.08, 2591.81		1856.22, 4252.94	
	Median	1963.70		3236.32	
	Q25, Q75	723.50, 4131.49		1920.12, 5806.46	
	Min, Max	336.7, 7988.3		394.5, 9539.9	
	Mean (SD)	2424.35 (2031.926)		3737.53 (2559.083)	
	95% CI				

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.0250 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_spmas.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.9.1 Descriptive Statistics of S-PMA Urinary Concentration Adjusted for Creatinine (pg/mg creat) by Sex - FAS**

Timepoint	Statistic	SA			
		Male (N=20)		Female (N=19)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 1	n	20	20	19	19
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	687.41 (115.96%)		1345.20 (131.42%)	
	Geometric 95% CI	446.26, 1058.86		830.10, 2179.93	
	Median	923.29	-55.13	1458.12	-51.81
	Q25, Q75	310.14, 1445.56	-65.27, -51.00	752.73, 3206.90	-59.81, -37.96
	Min, Max	161.0, 2431.8	-79.3, -30.3	184.3, 5357.1	-74.2, -26.6
	Mean (SD)	963.53 (702.607)	-57.48 (11.129)	1939.45 (1460.071)	-50.17 (13.837)
	95% CI		-62.69, -52.26		-56.84, -43.49

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.0250 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_spmas.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.9.1 Descriptive Statistics of S-PMA Urinary Concentration Adjusted for Creatinine (pg/mg creat) by Sex - FAS**

Timepoint	Statistic	SA			
		Male (N=20)		Female (N=19)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 2	n	20	20	19	19
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	310.98 (61.70%)		665.81 (74.00%)	
	Geometric 95% CI	238.39, 405.68		484.19, 915.53	
	Median	320.14	-83.52	642.86	-77.88
	Q25, Q75	209.91, 507.25	-87.25, -72.85	426.35, 1280.00	-83.59, -70.61
	Min, Max	104.7, 681.1	-91.5, -44.7	161.4, 1629.8	-87.0, -42.1
	Mean (SD)	357.77 (181.237)	-78.87 (11.996)	800.57 (462.296)	-73.85 (12.970)
	95% CI		-84.49, -73.25		-80.11, -67.60

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.0250 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_spmas.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.9.1 Descriptive Statistics of S-PMA Urinary Concentration Adjusted for Creatinine (pg/mg creat) by Sex - FAS**

Timepoint	Statistic	SA			
		Male (N=20)		Female (N=19)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 3	n	20	20	19	19
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	151.94 (59.42%)		337.80 (76.32%)	
	Geometric 95% CI	117.46, 196.54		243.69, 468.24	
	Median	150.20	-91.35	318.39	-88.47
	Q25, Q75	99.67, 237.39	-93.85, -86.34	208.02, 603.69	-91.74, -83.64
	Min, Max	52.0, 383.8	-96.2, -74.9	98.8, 1149.4	-93.6, -66.5
	Mean (SD)	175.14 (97.148)	-89.29 (6.660)	415.48 (274.126)	-86.85 (6.374)
	95% CI		-92.41, -86.16		-89.93, -83.77

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.0250 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_spmas.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.9.1 Descriptive Statistics of S-PMA Urinary Concentration Adjusted for Creatinine (pg/mg creat) by Sex - FAS**

Timepoint	Statistic	SA			
		Male (N=20)		Female (N=19)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 4	n	20	20	19	19
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	108.78 (55.49%)		234.31 (63.84%)	
	Geometric 95% CI	85.35, 138.63		176.77, 310.59	
	Median	98.35	-92.92	211.03	-91.87
	Q25, Q75	70.40, 150.62	-96.04, -90.91	129.74, 380.26	-94.38, -88.06
	Min, Max	54.2, 283.2	-97.7, -80.5	100.6, 690.5	-95.7, -70.5
	Mean (SD)	124.52 (71.046)	-92.21 (4.935)	275.65 (165.330)	-90.52 (5.837)
	95% CI		-94.53, -89.90		-93.33, -87.70

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.0250 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_spmas.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.9.1 Descriptive Statistics of S-PMA Urinary Concentration Adjusted for Creatinine (pg/mg creat) by Sex - FAS**

Timepoint	Statistic	SA			
		Male (N=20)		Female (N=19)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 5	n	20	20	19	19
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	106.79 (31.80%)		196.41 (50.54%)	
	Geometric 95% CI	92.35, 123.48		156.07, 247.16	
	Median	106.59	-93.67	169.33	-94.22
	Q25, Q75	80.63, 142.94	-96.61, -90.09	142.29, 274.83	-94.83, -88.73
	Min, Max	67.2, 171.0	-97.9, -74.0	95.2, 509.0	-96.6, -70.9
	Mean (SD)	111.76 (34.493)	-91.46 (7.303)	219.83 (114.217)	-91.59 (6.325)
	95% CI		-94.89, -88.04		-94.64, -88.53

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.0250 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_spmas.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.9.2 Descriptive Statistics of S-PMA Urinary Concentration Adjusted for Creatinine (pg/mg creat) by Cigarette Consumption - FAS**

Timepoint	Statistic	THS 2.2			
		10-19 cig/day (N=41)		>19 cig/day (N=39)	
		Raw value	%Change (*)	Raw value	%Change (*)
Baseline	n	41		39	
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	1835.08 (98.66%)		3166.11 (92.57%)	
	Geometric 95% CI	1414.62, 2380.52		2453.44, 4085.79	
	Median	2138.64		3182.32	
	Q25, Q75	935.51, 3409.10		2051.28, 6227.96	
	Min, Max	395.9, 7036.4		419.0, 11111.1	
	Mean (SD)	2447.42 (1728.546)		4043.64 (2564.578)	
	95% CI				

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.0250 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_spmac.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.9.2 Descriptive Statistics of S-PMA Urinary Concentration Adjusted for Creatinine (pg/mg creat) by Cigarette Consumption - FAS**

Timepoint	Statistic	THS 2.2			
		10-19 cig/day (N=41)		>19 cig/day (N=39)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 1	n	41	41	39	39
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	785.05 (109.57%)		1354.93 (105.70%)	
	Geometric 95% CI	593.14, 1039.06		1023.27, 1794.08	
	Median	1048.08	-57.57	1669.15	-58.15
	Q25, Q75	377.83, 1569.98	-64.62, -45.90	872.93, 2539.69	-63.32, -49.66
	Min, Max	139.7, 3321.1	-81.0, -31.9	130.5, 4554.9	-75.8, -14.6
	Mean (SD)	1085.80 (790.426)	-55.41 (12.425)	1790.81 (1168.179)	-55.84 (11.592)
	95% CI		-59.34, -51.48		-59.60, -52.08

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.0250 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_spmac.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.9.2 Descriptive Statistics of S-PMA Urinary Concentration Adjusted for Creatinine (pg/mg creat) by Cigarette Consumption - FAS**

Timepoint	Statistic	THS 2.2			
		10-19 cig/day (N=41)		>19 cig/day (N=39)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 2	n	41	41	39	39
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	437.04 (100.26%)		620.69 (80.95%)	
	Geometric 95% CI	335.87, 568.67		493.10, 781.29	
	Median	517.24	-77.42	687.02	-80.98
	Q25, Q75	239.92, 731.67	-82.34, -71.39	392.04, 1051.43	-85.48, -74.62
	Min, Max	96.5, 3355.4	-90.5, -1.9	140.9, 2245.7	-87.8, -46.9
	Mean (SD)	610.34 (586.537)	-72.42 (18.386)	774.78 (507.489)	-79.20 (8.016)
	95% CI		-78.23, -66.62		-81.80, -76.59

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.0250 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_spmac.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.9.2 Descriptive Statistics of S-PMA Urinary Concentration Adjusted for Creatinine (pg/mg creat) by Cigarette Consumption - FAS**

Timepoint	Statistic	THS 2.2			
		10-19 cig/day (N=41)		>19 cig/day (N=39)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 3	n	40	40	39	39
	BLOQ - n (%)	1 (2.5%)		0	
	Geometric Mean (CV%)	220.68 (84.54%)		349.42 (113.86%)	
	Geometric 95% CI	174.48, 279.09		260.00, 469.58	
	Median	256.36	-88.67	349.53	-90.23
	Q25, Q75	134.23, 377.74	-91.13, -85.51	188.80, 534.40	-91.15, -86.70
	Min, Max	43.1, 902.1	-95.3, -69.5	79.4, 10732.9	-95.5, 25.7
	Mean (SD)	278.45 (183.041)	-87.40 (5.193)	674.88 (1682.475)	-86.01 (18.852)
	95% CI		-89.06, -85.73		-92.13, -79.90

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.0250 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_spmac.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.9.2 Descriptive Statistics of S-PMA Urinary Concentration Adjusted for Creatinine (pg/mg creat) by Cigarette Consumption - FAS**

Timepoint	Statistic	THS 2.2			
		10-19 cig/day (N=41)		>19 cig/day (N=39)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 4	n	40	40	39	39
	BLOQ - n (%)	1 (2.5%)		0	
	Geometric Mean (CV%)	159.59 (67.90%)		224.13 (76.57%)	
	Geometric 95% CI	131.06, 194.33		179.83, 279.34	
	Median	189.28	-91.87	238.32	-92.73
	Q25, Q75	105.74, 237.40	-93.77, -89.55	140.56, 331.96	-95.35, -91.16
	Min, Max	36.3, 454.1	-97.3, -76.2	50.7, 1244.9	-96.1, -77.9
	Mean (SD)	188.72 (105.967)	-90.42 (5.096)	282.81 (225.414)	-92.12 (4.221)
	95% CI		-92.05, -88.79		-93.49, -90.75

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.0250 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_spmac.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.9.2 Descriptive Statistics of S-PMA Urinary Concentration Adjusted for Creatinine (pg/mg creat) by Cigarette Consumption - FAS**

Timepoint	Statistic	THS 2.2			
		10-19 cig/day (N=41)		>19 cig/day (N=39)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 5	n	40	40	39	39
	BLOQ - n (%)	1 (2.5%)		2 (5.1%)	
	Geometric Mean (CV%)	146.32 (56.60%)		185.39 (67.30%)	
	Geometric 95% CI	123.61, 173.19		152.06, 226.01	
	Median	146.61	-92.87	182.85	-94.51
	Q25, Q75	108.27, 208.03	-94.19, -89.00	143.65, 258.39	-95.86, -92.35
	Min, Max	43.9, 395.0	-97.7, -75.3	46.0, 873.8	-97.7, -75.6
	Mean (SD)	166.08 (82.799)	-90.89 (5.577)	222.95 (155.025)	-93.19 (4.615)
	95% CI		-92.68, -89.11		-94.69, -91.69

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.0250 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_spmac.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.9.2 Descriptive Statistics of S-PMA Urinary Concentration Adjusted for Creatinine (pg/mg creat) by Cigarette Consumption - FAS**

Timepoint	Statistic	CC			
		10-19 cig/day (N=21)		>19 cig/day (N=20)	
		Raw value	%Change (*)	Raw value	%Change (*)
Baseline	n	21		20	
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	2985.05 (57.20%)		2551.76 (98.42%)	
	Geometric 95% CI	2343.02, 3803.01		1736.06, 3750.74	
	Median	3333.33		2435.68	
	Q25, Q75	2271.42, 3794.75		1679.65, 4086.66	
	Min, Max	1166.1, 7956.1		357.1, 8830.3	
	Mean (SD)	3401.54 (1791.826)		3398.24 (2609.996)	
	95% CI				

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.0250 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_spmac.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.9.2 Descriptive Statistics of S-PMA Urinary Concentration Adjusted for Creatinine (pg/mg creat) by Cigarette Consumption - FAS**

Timepoint	Statistic	CC			
		10-19 cig/day (N=21)		>19 cig/day (N=20)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 1	n	21	21	20	20
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	3208.47 (59.89%)		2561.61 (90.31%)	
	Geometric 95% CI	2493.76, 4128.02		1784.61, 3676.90	
	Median	3082.10	6.09	2665.03	8.50
	Q25, Q75	2550.09, 3751.69	-7.54, 12.65	1660.68, 3869.70	-4.52, 11.72
	Min, Max	1032.6, 8281.5	-30.1, 123.8	390.7, 10305.2	-53.4, 23.3
	Mean (SD)	3691.14 (2029.709)	11.36 (34.074)	3323.03 (2552.215)	2.28 (17.453)
	95% CI		-4.15, 26.88		-5.90, 10.45

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.0250 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_spmac.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.9.2 Descriptive Statistics of S-PMA Urinary Concentration Adjusted for Creatinine (pg/mg creat) by Cigarette Consumption - FAS**

Timepoint	Statistic	CC			
		10-19 cig/day (N=21)		>19 cig/day (N=20)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 2	n	21	21	20	20
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	2778.38 (75.56%)		2556.30 (94.14%)	
	Geometric 95% CI	2046.14, 3772.66		1760.75, 3711.32	
	Median	2813.99	-7.03	2441.38	7.57
	Q25, Q75	2395.14, 3924.53	-18.65, 15.02	1742.68, 3903.48	-13.41, 25.59
	Min, Max	498.7, 7938.5	-78.0, 88.6	299.8, 11642.4	-53.0, 51.7
	Mean (SD)	3329.89 (1892.339)	0.27 (36.314)	3357.38 (2692.527)	4.33 (28.194)
	95% CI		-16.26, 16.81		-8.87, 17.53

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.0250 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_spmac.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.9.2 Descriptive Statistics of S-PMA Urinary Concentration Adjusted for Creatinine (pg/mg creat) by Cigarette Consumption - FAS**

Timepoint	Statistic	CC			
		10-19 cig/day (N=21)		>19 cig/day (N=20)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 3	n	21	21	20	20
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	3104.24 (55.46%)		2316.77 (89.67%)	
	Geometric 95% CI	2452.34, 3929.42		1617.18, 3319.00	
	Median	3181.82	3.81	2372.73	5.52
	Q25, Q75	2691.58, 3590.82	-14.05, 24.18	1774.99, 3670.41	-15.39, 25.10
	Min, Max	1070.0, 7984.5	-48.1, 139.1	285.3, 9225.0	-89.2, 58.2
	Mean (SD)	3511.70 (1836.048)	9.13 (38.077)	2928.98 (2020.447)	1.71 (36.839)
	95% CI		-8.20, 26.47		-15.54, 18.96

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.0250 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_spmac.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.9.2 Descriptive Statistics of S-PMA Urinary Concentration Adjusted for Creatinine (pg/mg creat) by Cigarette Consumption - FAS**

Timepoint	Statistic	CC			
		10-19 cig/day (N=21)		>19 cig/day (N=20)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 4	n	21	21	20	20
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	3062.84 (59.21%)		2733.22 (89.66%)	
	Geometric 95% CI	2386.49, 3930.88		1907.92, 3915.53	
	Median	3011.58	4.30	2672.69	10.70
	Q25, Q75	2400.00, 4189.61	-16.79, 12.57	1906.42, 4196.52	-8.00, 33.16
	Min, Max	1229.8, 8539.6	-34.3, 126.2	340.5, 9258.1	-67.5, 82.4
	Mean (SD)	3520.38 (1925.023)	6.47 (33.608)	3493.25 (2490.053)	12.93 (33.221)
	95% CI		-8.83, 21.77		-2.62, 28.48

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.0250 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_spmac.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.9.2 Descriptive Statistics of S-PMA Urinary Concentration Adjusted for Creatinine (pg/mg creat) by Cigarette Consumption - FAS**

Timepoint	Statistic	CC			
		10-19 cig/day (N=21)		>19 cig/day (N=20)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 5	n	21	21	20	20
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	3101.60 (61.00%)		2746.16 (92.19%)	
	Geometric 95% CI	2401.10, 4006.45		1902.46, 3964.00	
	Median	3119.27	1.02	2656.63	8.19
	Q25, Q75	2471.65, 4418.72	-9.18, 13.56	1867.33, 4522.68	0.35, 25.37
	Min, Max	966.7, 6918.1	-33.4, 127.5	352.0, 9345.5	-46.4, 83.8
	Mean (SD)	3557.17 (1821.361)	7.98 (35.239)	3539.00 (2520.314)	11.07 (28.014)
	95% CI		-8.07, 24.02		-2.05, 24.18

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.0250 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_spmac.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.9.2 Descriptive Statistics of S-PMA Urinary Concentration Adjusted for Creatinine (pg/mg creat) by Cigarette Consumption - FAS**

Timepoint	Statistic	SA			
		10-19 cig/day (N=19)		>19 cig/day (N=20)	
		Raw value	%Change (*)	Raw value	%Change (*)
Baseline	n	19		20	
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	2118.16 (84.98%)		2188.24 (152.06%)	
	Geometric 95% CI	1484.68, 3021.93		1311.18, 3651.97	
	Median	2261.21		2079.43	
	Q25, Q75	1552.63, 4060.85		871.74, 5984.95	
	Min, Max	394.5, 5153.8		336.7, 9539.9	
	Mean (SD)	2603.27 (1445.388)		3501.89 (2970.519)	
	95% CI				

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.0250 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_spmac.sas

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**Table 15.2.3.9.2 Descriptive Statistics of S-PMA Urinary Concentration Adjusted for Creatinine (pg/mg creat) by Cigarette Consumption - FAS**

Timepoint	Statistic	SA			
		10-19 cig/day (N=19)		>19 cig/day (N=20)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 1	n	19	19	20	20
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	909.04 (106.96%)		997.49 (164.13%)	
	Geometric 95% CI	596.72, 1384.82		584.21, 1703.13	
	Median	1367.71	-54.84	1031.73	-53.49
	Q25, Q75	533.33, 1464.91	-65.65, -47.67	401.57, 2493.87	-60.66, -45.28
	Min, Max	163.6, 3206.9	-79.3, -26.6	161.0, 5357.1	-74.2, -30.3
	Mean (SD)	1195.42 (745.907)	-55.02 (13.907)	1670.35 (1535.877)	-52.86 (12.125)
	95% CI		-61.73, -48.32		-58.54, -47.18

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.0250 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_spmac.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.9.2 Descriptive Statistics of S-PMA Urinary Concentration Adjusted for Creatinine (pg/mg creat) by Cigarette Consumption - FAS**

Timepoint	Statistic	SA			
		10-19 cig/day (N=19)		>19 cig/day (N=20)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 2	n	19	19	20	20
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	414.04 (73.99%)		488.34 (91.27%)	
	Geometric 95% CI	301.12, 569.32		339.23, 702.98	
	Median	496.82	-82.78	435.97	-77.64
	Q25, Q75	228.33, 661.61	-87.17, -70.61	292.42, 885.49	-84.63, -69.14
	Min, Max	104.7, 1373.3	-89.5, -42.1	126.1, 1629.8	-91.5, -44.7
	Mean (SD)	501.35 (313.143)	-77.90 (12.596)	642.03 (482.514)	-75.03 (12.716)
	95% CI		-83.97, -71.82		-80.99, -69.08

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.0250 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_spmac.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.9.2 Descriptive Statistics of S-PMA Urinary Concentration Adjusted for Creatinine (pg/mg creat) by Cigarette Consumption - FAS**

Timepoint	Statistic	SA			
		10-19 cig/day (N=19)		>19 cig/day (N=20)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 3	n	19	19	20	20
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	213.73 (71.84%)		234.71 (97.27%)	
	Geometric 95% CI	156.61, 291.67		160.20, 343.86	
	Median	222.91	-89.95	201.08	-89.09
	Q25, Q75	124.40, 372.52	-93.43, -86.32	123.97, 433.65	-92.65, -83.27
	Min, Max	52.0, 603.7	-95.7, -66.5	81.3, 1149.4	-96.2, -74.9
	Mean (SD)	255.35 (146.987)	-88.59 (6.606)	327.27 (295.174)	-87.63 (6.638)
	95% CI		-91.78, -85.40		-90.74, -84.52

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.0250 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_spmac.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.9.2 Descriptive Statistics of S-PMA Urinary Concentration Adjusted for Creatinine (pg/mg creat) by Cigarette Consumption - FAS**

Timepoint	Statistic	SA			
		10-19 cig/day (N=19)		>19 cig/day (N=20)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 4	n	19	19	20	20
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	150.95 (60.48%)		165.18 (90.15%)	
	Geometric 95% CI	115.33, 197.56		115.13, 236.98	
	Median	142.35	-92.28	147.69	-91.89
	Q25, Q75	100.61, 256.64	-95.33, -90.95	79.63, 324.80	-95.08, -87.79
	Min, Max	54.2, 379.9	-97.7, -70.5	59.9, 690.5	-97.6, -80.5
	Mean (SD)	173.69 (92.284)	-91.60 (5.863)	221.39 (182.814)	-91.18 (5.044)
	95% CI		-94.43, -88.77		-93.54, -88.81

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.0250 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_spmac.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.3.9.2 Descriptive Statistics of S-PMA Urinary Concentration Adjusted for Creatinine (pg/mg creat) by Cigarette Consumption - FAS**

Timepoint	Statistic	SA			
		10-19 cig/day (N=19)		>19 cig/day (N=20)	
		Raw value	%Change (*)	Raw value	%Change (*)
Day 5	n	19	19	20	20
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	132.53 (41.58%)		155.17 (63.46%)	
	Geometric 95% CI	109.32, 160.66		118.19, 203.73	
	Median	127.25	-94.18	149.90	-93.67
	Q25, Q75	95.23, 162.89	-96.49, -91.49	92.68, 209.46	-96.24, -88.46
	Min, Max	67.2, 274.8	-97.0, -70.9	71.8, 509.0	-97.9, -74.0
	Mean (SD)	142.94 (58.228)	-92.42 (6.023)	184.81 (124.204)	-90.67 (7.436)
	95% CI		-95.33, -89.51		-94.16, -87.19

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.0250 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_spmac.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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15.2.4 Secondary Endpoints Summary Tables

**Table 15.2.4.1 Primary Biomarkers of Exposure versus SA on Day 5 - FAS**

Variable	Statistic	THS 2.2 (N=80)	SA (N=39)	THS : SA Ratio (%)
Evening COHb (%)	n	79	39	
	Geometric LS Mean (CV%)	1.06	0.98	108.16 (16.84)
	95% CI	1.02, 1.11	0.93, 1.04	101.37, 115.41
Urinary MHBMA (pg/mg creat)	n	79	39	
	Geometric LS Mean (CV%)	194.05	168.01	115.51 (54.57)
	95% CI	173.23, 217.38	142.88, 197.55	94.78, 140.76
Urinary 3-HPMA (ng/mg creat)	n	79	39	
	Geometric LS Mean (CV%)	398.87	242.56	164.44 (26.10)
	95% CI	376.74, 422.30	223.64, 263.09	148.90, 181.60
Urinary S-PMA (pg/mg creat)	n	79	39	
	Geometric LS Mean (CV%)	164.51	153.66	107.06 (37.30)
	95% CI	151.82, 178.25	137.03, 172.31	93.08, 123.13

Note: SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Adjusted geometric least squares (LS) means and confidence intervals (CIs) from an ANCOVA model conducted on log-transformed Day 5 values with log-transformed baseline value, study arm, sex and CC consumption reported at screening as fixed effect factors. Geometrical CV% of the ratio is estimated from the residual mean squares.

Baseline is defined as the last assessment prior to Day 1 product use for THS 2.2 and CC subjects and prior to 06:29 AM on Day 1 for SA subjects.

Note: The urinary parameters are concentrations adjusted for creatinine.

Appendix 15.4.4.1 and 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/tl_anlprimbiosafas.sas

Program Run: 27OCT14 cvn_ahedge Program Status: FINAL

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**Table 15.2.4.2 Urinary Quantity Excreted of MHBMA, 3-HPMA and S-PMA over 24 hours on Day 5 - FAS**

Variable	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	THS : CC Ratio (%)	THS : SA Ratio (%)
Urinary Quantity Excreted of MHBMA over 24 hours (ng)	n	79	41	39		
	Geometric LS Mean (SE) ¹	279.67	3424.15	263.97	8.17 (0.100)	105.95 (0.102)
	95% CI	249.33, 313.71	2916.83, 4019.70	224.04, 311.02	6.70, 9.96	86.73, 129.42
Urinary Quantity Excreted of 3-HPMA over 24 hours (µg)	n	79	41	39		
	Geometric LS Mean (SE) ¹	574.08	1417.81	382.96	40.49 (0.062)	149.91 (0.063)
	95% CI	534.66, 616.41	1284.46, 1565.01	346.09, 423.76	35.84, 45.74	132.45, 169.66
Urinary Quantity Excreted of S-PMA over 24 hours (ng)	n	79	41	39		
	Geometric LS Mean (SE) ¹	237.17	4039.58	243.42	5.87 (0.078)	97.43 (0.080)
	95% CI	216.86, 259.38	3565.57, 4576.59	214.19, 276.64	5.03, 6.85	83.34, 113.89

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Adjusted geometric least squares (LS) means and confidence intervals (CIs) from an ANCOVA model conducted on log-transformed Day 5 values with log-transformed baseline value, study arm, sex and CC consumption reported at screening as fixed effect factors. Geometrical CV% of the ratio is estimated from the residual mean squares.

Baseline is defined as the last assessment prior to Day 1 product use for THS 2.2 and CC subjects and prior to 06:29 AM on Day 1 for SA subjects.

Note: 1 = Standard errors presented from analysis performed on the log_e scale.

Note: Collection over 24 hours starting on the Day stated in the title.

Appendix 15.4.4.2 and 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/tl_anluexcbiosafas.sas

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Program Run: 27OCT14 cvn_ahedge Program Status: FINAL

**Table 15.2.4.3 Descriptive Statistics of Urinary Quantity Excreted of MHBMA over 24 hours (ng) - FAS**

Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
		Raw value	% Change (*)	Raw value	% Change (*)
Baseline	n	80		41	
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	3027.75 (108.31%)		3814.21 (93.34%)	
	Geometric 95% CI	2488.72, 3683.51		2970.95, 4896.82	
	Median	3974.13		3760.00	
	Q25, Q75	1742.45, 5840.25		2838.00, 5880.00	
	Min, Max	352.0, 13158.0		340.8, 15529.5	
	Mean (SD)	4096.56 (2741.754)		4953.33 (3599.456)	
	95% CI				

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.100 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_uquant.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.3 Descriptive Statistics of Urinary Quantity Excreted of MHBMA over 24 hours (ng) - FAS**

Timepoint	Statistic	SA (N=39)	
		Raw value	% Change (*)
Baseline	n	39	
	BLOQ - n (%)	0	
	Geometric Mean (CV%)	2685.41 (151.00%)	
	Geometric 95% CI	1886.15, 3823.35	
	Median	3078.00	
	Q25, Q75	939.90, 6030.00	
	Min, Max	262.6, 13517.8	
	Mean (SD)	4248.37 (3597.708)	
	95% CI		

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.100 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_uquant.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.3 Descriptive Statistics of Urinary Quantity Excreted of MHBMA over 24 hours (ng) - FAS**

Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
		Raw value	% Change (*)	Raw value	% Change (*)
Day 1	n	80	80	41	41
	BLOQ - n (%)	4 (5.0%)		0	
	Geometric Mean (CV%)	843.01 (83.58%)		3586.22 (102.47%)	
	Geometric 95% CI	716.94, 991.24		2744.70, 4685.74	
	Median	957.56	-74.60	3961.50	-5.72
	Q25, Q75	614.25, 1282.50	-80.00, -66.43	2369.15, 5644.00	-22.38, 7.61
	Min, Max	77.5, 8176.0	-86.1, -6.9	348.3, 15060.3	-46.3, 121.3
	Mean (SD)	1075.06 (961.876)	-69.42 (15.032)	4851.96 (3798.522)	-1.86 (31.508)
	95% CI		-72.77, -66.07		-11.81, 8.09

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.100 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_uquant.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.3 Descriptive Statistics of Urinary Quantity Excreted of MHBMA over 24 hours (ng) - FAS**

Timepoint	Statistic	SA (N=39)	
		Raw value	% Change (*)
Day 1	n	39	39
	BLOQ - n (%)	1 (2.6%)	
	Geometric Mean (CV%)	837.60 (79.03%)	
	Geometric 95% CI	668.30, 1049.79	
	Median	846.80	-73.00
	Q25, Q75	582.79, 1358.50	-79.88, -56.64
	Min, Max	142.5, 2727.0	-85.6, 23.6
	Mean (SD)	1033.73 (657.093)	-63.60 (23.419)
	95% CI		-71.19, -56.00

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.100 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_uquant.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.3 Descriptive Statistics of Urinary Quantity Excreted of MHBMA over 24 hours (ng) - FAS**

Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
		Raw value	% Change (*)	Raw value	% Change (*)
Day 2	n	80	80	41	41
	BLOQ - n (%)	7 (8.8%)		0	
	Geometric Mean (CV%)	431.84 (59.44%)		3424.95 (109.68%)	
	Geometric 95% CI	382.08, 488.07		2587.24, 4533.91	
	Median	447.53	-88.79	3680.00	-10.01
	Q25, Q75	347.10, 576.18	-91.62, -74.26	2600.75, 5612.00	-28.06, 19.12
	Min, Max	105.0, 8008.0	-96.4, 37.9	322.0, 18124.0	-92.8, 179.8
	Mean (SD)	546.67 (860.641)	-79.47 (21.842)	4674.78 (3663.729)	-0.04 (44.404)
	95% CI		-84.34, -74.61		-14.06, 13.98

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.100 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_uquant.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.3 Descriptive Statistics of Urinary Quantity Excreted of MHBMA over 24 hours (ng) - FAS**

Timepoint	Statistic	SA (N=39)	
		Raw value	% Change (*)
Day 2	n	39	39
	BLOQ - n (%)	3 (7.7%)	
	Geometric Mean (CV%)	428.89 (44.78%)	
	Geometric 95% CI	373.38, 492.64	
	Median	457.30	-86.84
	Q25, Q75	363.20, 537.60	-92.59, -54.33
	Min, Max	147.5, 977.6	-97.9, 31.3
	Mean (SD)	464.47 (177.714)	-71.70 (33.342)
	95% CI		-82.52, -60.89

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.100 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_uquant.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.3 Descriptive Statistics of Urinary Quantity Excreted of MHBMA over 24 hours (ng) - FAS**

Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
		Raw value	% Change (*)	Raw value	% Change (*)
Day 3	n	79	79	41	41
	BLOQ - n (%)	9 (11.4%)		1 (2.4%)	
	Geometric Mean (CV%)	356.09 (62.09%)		3324.73 (116.09%)	
	Geometric 95% CI	313.34, 404.68		2483.83, 4450.31	
	Median	373.33	-90.53	3630.00	-2.87
	Q25, Q75	303.60, 463.20	-92.94, -80.93	2491.65, 5544.00	-23.95, 13.80
	Min, Max	80.0, 6930.0	-98.6, 28.4	132.5, 14832.0	-97.7, 57.7
	Mean (SD)	457.67 (750.993)	-81.80 (20.999)	4580.78 (3541.524)	-3.06 (31.705)
	95% CI		-86.51, -77.10		-13.07, 6.95

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.100 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_uquant.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.3 Descriptive Statistics of Urinary Quantity Excreted of MHBMA over 24 hours (ng) - FAS**

Timepoint	Statistic	SA (N=39)	
		Raw value	% Change (*)
Day 3	n	39	39
	BLOQ - n (%)	9 (23.1%)	
	Geometric Mean (CV%)	331.38 (54.55%)	
	Geometric 95% CI	280.85, 391.00	
	Median	364.00	-90.70
	Q25, Q75	249.60, 469.80	-94.08, -61.55
	Min, Max	116.3, 712.4	-98.4, 2.2
	Mean (SD)	371.53 (166.484)	-76.22 (28.587)
	95% CI		-85.49, -66.95

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.100 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_uquant.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.3 Descriptive Statistics of Urinary Quantity Excreted of MHBMA over 24 hours (ng) - FAS**

Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
		Raw value	% Change (*)	Raw value	% Change (*)
Day 4	n	79	79	41	41
	BLOQ - n (%)	16 (20.3%)		0	
	Geometric Mean (CV%)	315.48 (48.52%)		3640.03 (94.63%)	
	Geometric 95% CI	284.60, 349.70		2828.05, 4685.14	
	Median	351.53	-91.71	3726.00	-4.80
	Q25, Q75	245.00, 439.45	-93.84, -81.09	2508.00, 5486.25	-25.26, 16.41
	Min, Max	90.0, 690.2	-98.9, -36.5	443.7, 17821.5	-43.0, 175.2
	Mean (SD)	346.12 (137.865)	-84.62 (15.317)	4813.61 (3814.045)	0.26 (36.860)
	95% CI		-88.06, -81.19		-11.38, 11.90

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.100 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_uquant.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.3 Descriptive Statistics of Urinary Quantity Excreted of MHBMA over 24 hours (ng) - FAS**

Timepoint	Statistic	SA (N=39)	
		Raw value	% Change (*)
Day 4	n	39	39
	BLOQ - n (%)	13 (33.3%)	
	Geometric Mean (CV%)	279.30 (54.17%)	
	Geometric 95% CI	236.95, 329.21	
	Median	305.10	-93.33
	Q25, Q75	196.87, 401.48	-95.80, -62.52
	Min, Max	100.0, 777.8	-98.2, 7.0
	Mean (SD)	315.08 (157.825)	-76.73 (31.183)
	95% CI		-86.85, -66.62

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.100 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_uquant.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.4.3 Descriptive Statistics of Urinary Quantity Excreted of MHBMA over 24 hours (ng) - FAS**

Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
		Raw value	% Change (*)	Raw value	% Change (*)
Day 5	n	79	79	41	41
	BLOQ - n (%)	34 (43.0%)		1 (2.4%)	
	Geometric Mean (CV%)	277.00 (44.49%)		3603.43 (112.80%)	
	Geometric 95% CI	251.85, 304.67		2707.24, 4796.30	
	Median	301.30	-92.38	3687.45	-2.70
	Q25, Q75	215.00, 387.20	-95.03, -86.19	2489.80, 5582.50	-21.95, 7.97
	Min, Max	112.5, 647.5	-97.7, -39.8	140.0, 16351.0	-58.9, 126.0
	Mean (SD)	301.28 (119.751)	-86.42 (14.000)	4948.11 (3917.928)	-0.39 (34.033)
	95% CI		-89.56, -83.28		-11.13, 10.36

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.100 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_uquant.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.3 Descriptive Statistics of Urinary Quantity Excreted of MHBMA over 24 hours (ng) - FAS**

Timepoint	Statistic	SA (N=39)	
		Raw value	% Change (*)
Day 5	n	39	39
	BLOQ - n (%)	16 (41.0%)	
	Geometric Mean (CV%)	255.97 (50.13%)	
	Geometric 95% CI	219.54, 298.43	
	Median	292.60	-93.64
	Q25, Q75	156.25, 361.20	-96.07, -67.51
	Min, Max	105.0, 573.2	-98.2, -16.0
	Mean (SD)	284.29 (128.347)	-79.91 (25.476)
	95% CI		-88.18, -71.65

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.100 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_uquant.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.4.4 Descriptive Statistics of Urinary Quantity Excreted of 3-HPMA over 24 hours (µg) - FAS**

Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
		Raw value	% Change (*)	Raw value	% Change (*)
Baseline	n	80		41	
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	1349.85 (56.09%)		1317.25 (48.59%)	
	Geometric 95% CI	1201.54, 1516.46		1139.09, 1523.27	
	Median	1481.33		1309.38	
	Q25, Q75	1085.90, 1943.80		968.00, 1644.30	
	Min, Max	144, 3680		545, 3977	
	Mean (SD)	1505.74 (629.335)		1472.29 (783.334)	
	95% CI				

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (20.0 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_uquant2.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.4 Descriptive Statistics of Urinary Quantity Excreted of 3-HPMA over 24 hours (μg) - FAS**

Timepoint	Statistic	SA (N=39)	
		Raw value	% Change (*)
Baseline	n	39	
	BLOQ - n (%)	0	
	Geometric Mean (CV%)	1361.96 (47.69%)	
	Geometric 95% CI	1176.09, 1577.21	
	Median	1305.50	
	Q25, Q75	871.70, 1824.00	
	Min, Max	641, 3788	
	Mean (SD)	1512.55 (762.444)	
	95% CI		

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (20.0 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_uquant2.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.4 Descriptive Statistics of Urinary Quantity Excreted of 3-HPMA over 24 hours (µg) - FAS**

Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
		Raw value	% Change (*)	Raw value	% Change (*)
Day 1	n	80	80	41	41
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	679.06 (41.55%)		1406.91 (53.32%)	
	Geometric 95% CI	621.35, 742.12		1201.44, 1647.52	
	Median	685.90	-50.66	1236.90	3.49
	Q25, Q75	539.70, 852.67	-61.43, -38.54	1062.59, 1921.45	-3.54, 25.13
	Min, Max	102, 1606	-75, 33	464, 4292	-44, 96
	Mean (SD)	728.74 (268.480)	-46.13 (21.376)	1596.75 (871.097)	10.66 (30.534)
	95% CI		-50.90, -41.37		1.01, 20.30

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (20.0 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_uquant2.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.4 Descriptive Statistics of Urinary Quantity Excreted of 3-HPMA over 24 hours (µg) - FAS**

Timepoint	Statistic	SA (N=39)	
		Raw value	% Change (*)
Day 1	n	39	39
	BLOQ - n (%)	0	
	Geometric Mean (CV%)	545.30 (37.68%)	
	Geometric 95% CI	484.56, 613.66	
	Median	574.75	-60.35
	Q25, Q75	427.20, 684.00	-69.94, -49.75
	Min, Max	253, 1159	-80, -8
	Mean (SD)	580.99 (210.144)	-57.81 (14.357)
	95% CI		-62.47, -53.15

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (20.0 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_uquant2.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.4 Descriptive Statistics of Urinary Quantity Excreted of 3-HPMA over 24 hours (µg) - FAS**

Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
		Raw value	% Change (*)	Raw value	% Change (*)
Day 2	n	80	80	41	41
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	472.89 (54.26%)		1246.88 (60.70%)	
	Geometric 95% CI	422.34, 529.49		1044.85, 1487.98	
	Median	479.45	-64.17	1086.40	-6.48
	Q25, Q75	381.82, 616.96	-73.80, -55.73	878.70, 1986.00	-22.27, 25.87
	Min, Max	74, 2475	-89, 72	296, 3718	-82, 124
	Mean (SD)	534.71 (302.941)	-61.62 (20.343)	1455.48 (858.209)	1.88 (38.960)
	95% CI		-66.15, -57.09		-10.43, 14.18

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (20.0 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_uquant2.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.4 Descriptive Statistics of Urinary Quantity Excreted of 3-HPMA over 24 hours (µg) - FAS**

Timepoint	Statistic	SA (N=39)	
		Raw value	% Change (*)
Day 2	n	39	39
	BLOQ - n (%)	0	
	Geometric Mean (CV%)	312.02 (41.30%)	
	Geometric 95% CI	274.35, 354.85	
	Median	318.60	-75.96
	Q25, Q75	261.80, 432.53	-82.93, -67.56
	Min, Max	82, 596	-94, -54
	Mean (SD)	333.97 (115.199)	-74.80 (10.380)
	95% CI		-78.17, -71.43

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (20.0 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_uquant2.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.4 Descriptive Statistics of Urinary Quantity Excreted of 3-HPMA over 24 hours (µg) - FAS**

Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
		Raw value	% Change (*)	Raw value	% Change (*)
Day 3	n	79	79	41	41
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	648.21 (46.89%)		1420.92 (50.78%)	
	Geometric 95% CI	586.61, 716.28		1221.58, 1652.79	
	Median	679.40	-52.48	1275.00	8.76
	Q25, Q75	534.00, 816.00	-62.80, -44.07	1070.00, 1802.00	-9.39, 23.45
	Min, Max	46, 2121	-84, 64	485, 3875	-65, 111
	Mean (SD)	701.08 (272.039)	-49.24 (21.426)	1596.38 (836.800)	12.81 (33.815)
	95% CI		-54.04, -44.43		2.13, 23.49

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (20.0 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_uquant2.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.4 Descriptive Statistics of Urinary Quantity Excreted of 3-HPMA over 24 hours (µg) - FAS**

Timepoint	Statistic	SA (N=39)	
		Raw value	% Change (*)
Day 3	n	39	39
	BLOQ - n (%)	0	
	Geometric Mean (CV%)	448.05 (45.32%)	
	Geometric 95% CI	389.47, 515.44	
	Median	491.40	-64.48
	Q25, Q75	389.55, 594.15	-77.03, -57.13
	Min, Max	111, 1163	-91, -26
	Mean (SD)	485.21 (185.047)	-64.29 (14.146)
	95% CI		-68.88, -59.70

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (20.0 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_uquant2.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.4 Descriptive Statistics of Urinary Quantity Excreted of 3-HPMA over 24 hours (µg) - FAS**

Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
		Raw value	% Change (*)	Raw value	% Change (*)
Day 4	n	79	79	41	41
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	410.09 (48.74%)		1206.43 (57.92%)	
	Geometric 95% CI	369.80, 454.77		1018.06, 1429.65	
	Median	389.20	-70.38	1073.50	-10.39
	Q25, Q75	331.20, 552.21	-76.27, -63.53	867.50, 1545.00	-19.62, 6.33
	Min, Max	43, 974	-88, 15	490, 3692	-48, 122
	Mean (SD)	448.88 (182.047)	-67.07 (17.144)	1406.10 (874.889)	-4.65 (29.949)
	95% CI		-70.91, -63.22		-14.11, 4.80

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (20.0 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_uquant2.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.4 Descriptive Statistics of Urinary Quantity Excreted of 3-HPMA over 24 hours (μg) - FAS**

Timepoint	Statistic	SA (N=39)	
		Raw value	% Change (*)
Day 4	n	39	39
	BLOQ - n (%)	0	
	Geometric Mean (CV%)	224.12 (46.87%)	
	Geometric 95% CI	193.97, 258.95	
	Median	246.05	-82.67
	Q25, Q75	147.75, 299.61	-87.50, -78.12
	Min, Max	97, 564	-93, -61
	Mean (SD)	246.41 (109.940)	-82.07 (7.610)
	95% CI		-84.54, -79.60

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (20.0 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_uquant2.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.4 Descriptive Statistics of Urinary Quantity Excreted of 3-HPMA over 24 hours (µg) - FAS**

Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
		Raw value	% Change (*)	Raw value	% Change (*)
Day 5	n	79	79	41	41
	BLOQ - n (%)	1 (1.3%)		0	
	Geometric Mean (CV%)	577.54 (58.00%)		1398.20 (56.32%)	
	Geometric 95% CI	511.92, 651.58		1184.73, 1650.12	
	Median	636.00	-55.99	1271.90	0.53
	Q25, Q75	484.40, 781.00	-66.17, -46.36	954.80, 2079.18	-10.53, 29.59
	Min, Max	45, 1264	-86, 21	548, 3963	-34, 117
	Mean (SD)	642.84 (252.625)	-54.63 (17.490)	1604.83 (888.940)	10.22 (32.754)
	95% CI		-58.55, -50.71		-0.12, 20.57

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (20.0 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_uquant2.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.4 Descriptive Statistics of Urinary Quantity Excreted of 3-HPMA over 24 hours (µg) - FAS**

Timepoint	Statistic	SA (N=39)	
		Raw value	% Change (*)
Day 5	n	39	39
	BLOQ - n (%)	0	
	Geometric Mean (CV%)	385.41 (31.04%)	
	Geometric 95% CI	349.32, 425.23	
	Median	399.30	-72.37
	Q25, Q75	330.40, 459.90	-77.95, -57.66
	Min, Max	209, 680	-87, -47
	Mean (SD)	402.32 (116.356)	-69.49 (11.844)
	95% CI		-73.34, -65.65

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (20.0 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_uquant2.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.5 Descriptive Statistics of Urinary Quantity Excreted of S-PMA over 24 hours (ng) - FAS**

Timepoint	Statistic	THS 2.2		CC	
		Raw value	% Change (*)	Raw value	% Change (*)
Baseline	n	80		41	
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	3838.72 (94.24%)		4556.63 (83.71%)	
	Geometric 95% CI	3214.68, 4583.90		3620.35, 5735.04	
	Median	4346.25		5049.00	
	Q25, Q75	2333.60, 6957.00		3120.00, 7236.00	
	Min, Max	552.2, 14792.0		390.5, 20438.3	
	Mean (SD)	4984.31 (3237.856)		5714.28 (3967.905)	
	95% CI				

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.250 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_uquant3.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.5 Descriptive Statistics of Urinary Quantity Excreted of S-PMA over 24 hours (ng) - FAS**

Timepoint	Statistic	SA	
		Raw value	% Change (*)
Baseline	n	39	
	BLOQ - n (%)	0	
	Geometric Mean (CV%)	3403.50 (115.28%)	
	Geometric 95% CI	2526.30, 4585.29	
	Median	4104.00	
	Q25, Q75	1562.30, 7530.00	
	Min, Max	467.1, 18904.0	
	Mean (SD)	4873.45 (3967.404)	
	95% CI		

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.250 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_uquant3.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.5 Descriptive Statistics of Urinary Quantity Excreted of S-PMA over 24 hours (ng) - FAS**

Timepoint	Statistic	THS 2.2		CC	
		Raw value	% Change (*)	Raw value	% Change (*)
Day 1	n	80	80	41	41
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	1473.42 (108.42%)		4191.52 (84.93%)	
	Geometric 95% CI	1210.94, 1792.79		3321.69, 5289.12	
	Median	1639.65	-62.03	3892.00	-8.65
	Q25, Q75	829.32, 2531.13	-68.18, -53.46	2652.00, 7900.50	-22.72, 4.54
	Min, Max	140.1, 6160.0	-82.9, -22.4	415.0, 17632.0	-44.0, 79.3
	Mean (SD)	2003.98 (1405.532)	-59.86 (12.335)	5287.55 (3586.346)	-4.60 (27.756)
	95% CI		-62.61, -57.11		-13.37, 4.16

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.250 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_uquant3.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.5 Descriptive Statistics of Urinary Quantity Excreted of S-PMA over 24 hours (ng) - FAS**

Timepoint	Statistic	SA	
		Raw value	% Change (*)
Day 1	n	39	39
	BLOQ - n (%)	0	
	Geometric Mean (CV%)	1416.33 (123.80%)	
	Geometric 95% CI	1036.23, 1935.85	
	Median	1717.00	-59.44
	Q25, Q75	656.00, 3024.00	-65.37, -46.66
	Min, Max	191.2, 5355.0	-80.6, -24.1
	Mean (SD)	2037.99 (1512.243)	-56.15 (14.131)
	95% CI		-60.73, -51.56

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.250 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_uquant3.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.5 Descriptive Statistics of Urinary Quantity Excreted of S-PMA over 24 hours (ng) - FAS**

Timepoint	Statistic	THS 2.2		CC	
		Raw value	% Change (*)	Raw value	% Change (*)
Day 2	n	80	80	41	41
	BLOQ - n (%)	0		0	
	Geometric Mean (CV%)	775.53 (91.82%)		4254.75 (93.92%)	
	Geometric 95% CI	651.67, 922.93		3310.31, 5468.65	
	Median	868.05	-80.14	4462.50	-7.92
	Q25, Q75	488.48, 1336.32	-85.82, -75.07	3094.00, 7296.00	-21.15, 19.26
	Min, Max	144.7, 11368.0	-92.3, 15.1	309.0, 17755.5	-87.7, 186.1
	Mean (SD)	1072.43 (1318.596)	-76.46 (17.422)	5528.98 (4021.701)	3.52 (47.697)
	95% CI		-80.34, -72.58		-11.54, 18.58

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.250 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_uquant3.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.5 Descriptive Statistics of Urinary Quantity Excreted of S-PMA over 24 hours (ng) - FAS**

Timepoint	Statistic	SA	
		Raw value	% Change (*)
Day 2	n	39	39
	BLOQ - n (%)	0	
	Geometric Mean (CV%)	693.22 (68.19%)	
	Geometric 95% CI	567.38, 846.97	
	Median	718.40	-80.22
	Q25, Q75	461.55, 1196.35	-86.33, -70.45
	Min, Max	217.3, 2018.8	-95.5, -42.9
	Mean (SD)	823.01 (467.005)	-76.65 (12.685)
	95% CI		-80.76, -72.53

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.250 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_uquant3.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.5 Descriptive Statistics of Urinary Quantity Excreted of S-PMA over 24 hours (ng) - FAS**

Timepoint	Statistic	THS 2.2		CC	
		Raw value	% Change (*)	Raw value	% Change (*)
Day 3	n	79	79	41	41
	BLOQ - n (%)	1 (1.3%)		0	
	Geometric Mean (CV%)	409.37 (94.75%)		4167.47 (86.44%)	
	Geometric 95% CI	342.18, 489.76		3292.13, 5275.57	
	Median	439.20	-90.13	4320.00	-4.92
	Q25, Q75	243.75, 648.60	-92.13, -87.10	2714.82, 7095.00	-18.55, 15.70
	Min, Max	65.0, 13620.0	-96.0, 42.4	445.2, 18105.8	-91.1, 112.4
	Mean (SD)	655.46 (1517.822)	-87.41 (15.418)	5294.82 (3727.833)	-0.41 (36.701)
	95% CI		-90.87, -83.95		-12.00, 11.18

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.250 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_uquant3.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.5 Descriptive Statistics of Urinary Quantity Excreted of S-PMA over 24 hours (ng) - FAS**

Timepoint	Statistic	SA	
		Raw value	% Change (*)
Day 3	n	39	39
	BLOQ - n (%)	0	
	Geometric Mean (CV%)	344.02 (66.61%)	
	Geometric 95% CI	282.66, 418.69	
	Median	298.80	-89.41
	Q25, Q75	230.40, 600.60	-93.33, -84.63
	Min, Max	103.1, 1060.8	-96.4, -68.8
	Mean (SD)	409.45 (243.966)	-88.47 (6.128)
	95% CI		-90.46, -86.48

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.250 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_uquant3.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.5 Descriptive Statistics of Urinary Quantity Excreted of S-PMA over 24 hours (ng) - FAS**

Timepoint	Statistic	THS 2.2		CC	
		Raw value	% Change (*)	Raw value	% Change (*)
Day 4	n	79	79	41	41
	BLOQ - n (%)	1 (1.3%)		0	
	Geometric Mean (CV%)	275.68 (66.17%)		4417.91 (82.43%)	
	Geometric 95% CI	240.87, 315.52		3519.77, 5545.21	
	Median	294.10	-93.26	4521.00	-7.10
	Q25, Q75	189.00, 413.10	-94.93, -91.41	2805.00, 7020.00	-17.33, 10.06
	Min, Max	60.0, 1197.0	-96.9, -75.9	442.2, 19838.0	-50.0, 224.9
	Mean (SD)	326.86 (198.255)	-92.03 (4.354)	5567.80 (3996.553)	3.20 (46.086)
	95% CI		-93.01, -91.05		-11.35, 17.75

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.250 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_uquant3.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.5 Descriptive Statistics of Urinary Quantity Excreted of S-PMA over 24 hours (ng) - FAS**

Timepoint	Statistic	SA	
		Raw value	% Change (*)
Day 4	n	39	39
	BLOQ - n (%)	0	
	Geometric Mean (CV%)	232.61 (61.24%)	
	Geometric 95% CI	193.72, 279.31	
	Median	195.80	-92.92
	Q25, Q75	141.80, 393.90	-95.74, -90.22
	Min, Max	113.2, 690.3	-97.9, -71.7
	Mean (SD)	274.17 (169.492)	-91.89 (5.132)
	95% CI		-93.56, -90.22

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.250 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_uquant3.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.5 Descriptive Statistics of Urinary Quantity Excreted of S-PMA over 24 hours (ng) - FAS**

Timepoint	Statistic	THS 2.2		CC	
		Raw value	% Change (*)	Raw value	% Change (*)
Day 5	n	79	79	41	41
	BLOQ - n (%)	3 (3.8%)		0	
	Geometric Mean (CV%)	236.11 (60.35%)		4389.49 (92.62%)	
	Geometric 95% CI	208.40, 267.50		3424.00, 5627.22	
	Median	249.75	-94.19	4216.00	-5.14
	Q25, Q75	172.00, 346.50	-95.94, -91.62	2983.50, 7850.00	-18.43, 9.67
	Min, Max	35.6, 1085.4	-98.2, -77.2	316.4, 21331.0	-41.7, 237.0
	Mean (SD)	271.66 (150.385)	-92.84 (4.598)	5733.63 (4340.338)	1.59 (43.292)
	95% CI		-93.88, -91.81		-12.08, 15.26

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.250 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_uquant3.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.5 Descriptive Statistics of Urinary Quantity Excreted of S-PMA over 24 hours (ng) - FAS**

Timepoint	Statistic	SA	
		Raw value	% Change (*)
Day 5	n	39	39
	BLOQ - n (%)	0	
	Geometric Mean (CV%)	225.42 (43.24%)	
	Geometric 95% CI	197.10, 257.79	
	Median	220.63	-93.71
	Q25, Q75	154.10, 319.50	-96.16, -89.44
	Min, Max	119.5, 507.5	-98.6, -72.3
	Mean (SD)	245.26 (104.255)	-91.65 (6.187)
	95% CI		-93.66, -89.64

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.250 ng/mL).

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_uquant3.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.6 Analysis of Exhaled CO (ppm) on Day 5 - FAS**

Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	THS : CC Ratio (%)	THS : SA Ratio (%)
Missing	6	0	7		
n	73	41	32		
Geometric LS Mean (SE) ¹	2.03	12.33	2.11	16.48 (0.085)	96.38 (0.093)
95% CI	1.83, 2.25	10.79, 14.09	1.81, 2.46	13.95, 19.47	80.34, 115.61

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Adjusted geometric least squares (LS) means and confidence intervals (CIs) from an ANCOVA model conducted on log-transformed Day 5 values with log-transformed baseline value, study arm, sex and CC consumption reported at screening as fixed effect factors.

Baseline is defined as the last assessment prior to Day 1 product use for THS 2.2 and CC subjects and prior to 06:29 AM on Day 1 for SA subjects.

Note: 1 = Standard errors presented from analysis performed on the log_e scale.

Note: The evening value on day 5 is analyzed.

Note: Missing numbers show number of observations missing at a timepoint where the subject was still on the study due to 0 values recorded, which cannot be log-transformed.

Appendix 15.4.4.6 and 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/tl_anlsecondbio.sas

Program Run: 27OCT14 cvn_ahedge Program Status: FINAL

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**Table 15.2.4.7 Descriptive Statistics of Exhaled CO (ppm) - FAS**

Timepoint	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)
Day -2	n	80	41	39
	BLOQ - n (%)	0	0	0
	Geometric Mean (CV%)	15.9 (46.3%)	14.6 (66.9%)	17.6 (49.2%)
	95% CI	14.4, 17.6	12.0, 17.8	15.1, 20.5
	Median	17.5	15.0	19.0
	Q25, Q75	12.0, 22.0	8.0, 23.0	14.0, 23.0
	Min, Max	4, 37	4, 44	3, 33
Day -1, 15 min < T ₀	n	80	41	39
	BLOQ - n (%)	0	0	0
	Geometric Mean (CV%)	10.0 (50.6%)	9.8 (55.2%)	10.9 (39.0%)
	95% CI	8.9, 11.1	8.2, 11.5	9.6, 12.4
	Median	10.0	10.0	11.0
	Q25, Q75	7.0, 15.0	7.0, 13.0	9.0, 14.0
	Min, Max	2, 23	3, 22	4, 22

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: T₀ = Time of first product use on study day.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_co.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.7 Descriptive Statistics of Exhaled CO (ppm) - FAS**

Timepoint	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)
Day -1, 12:00-02:00 PM	n	80	41	39
	BLOQ - n (%)	0	0	0
	Geometric Mean (CV%)	14.2 (58.9%)	12.6 (64.6%)	14.0 (40.0%)
	95% CI	12.6, 16.1	10.4, 15.2	12.3, 15.9
	Median	16.0	14.0	14.0
	Q25, Q75	10.0, 21.0	9.0, 19.0	11.0, 18.0
	Min, Max	2, 45	2, 28	4, 26
Day -1, 04:00-06:00 PM	n	80	41	39
	BLOQ - n (%)	0	0	0
	Geometric Mean (CV%)	15.9 (41.9%)	15.2 (41.8%)	15.5 (35.9%)
	95% CI	14.5, 17.5	13.3, 17.3	13.8, 17.4
	Median	16.0	16.0	16.0
	Q25, Q75	12.0, 21.5	11.0, 20.0	12.0, 19.0
	Min, Max	5, 38	6, 35	7, 34

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: T₀ = Time of first product use on study day.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_co.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.7 Descriptive Statistics of Exhaled CO (ppm) - FAS**

Timepoint	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)
Day -1, 08:00-10:00 PM	n	80	41	39
	BLOQ - n (%)	0	0	0
	Geometric Mean (CV%)	14.4 (59.4%)	13.5 (44.9%)	15.1 (35.7%)
	95% CI	12.7, 16.3	11.7, 15.5	13.5, 16.9
	Median	16.0	15.0	14.0
	Q25, Q75	11.0, 21.0	11.0, 18.0	11.0, 21.0
	Min, Max	3, 41	4, 28	7, 27
Day 0, 15 min < T ₀	n	80	41	39
	BLOQ - n (%)	0	0	0
	Geometric Mean (CV%)	7.9 (47.0%)	7.6 (55.4%)	8.6 (37.8%)
	95% CI	7.1, 8.8	6.4, 9.0	7.6, 9.7
	Median	8.0	9.0	8.0
	Q25, Q75	6.0, 11.0	5.0, 11.0	7.0, 12.0
	Min, Max	2, 19	3, 18	4, 16

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: T₀ = Time of first product use on study day.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_co.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.7 Descriptive Statistics of Exhaled CO (ppm) - FAS**

Timepoint	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)
Day 0, 12:00-02:00 PM	n	80	41	39
	BLOQ - n (%)	0	0	0
	Geometric Mean (CV%)	16.0 (54.0%)	15.1 (52.1%)	15.9 (49.9%)
	95% CI	14.2, 18.0	12.9, 17.7	13.6, 18.6
	Median	16.5	16.0	15.0
	Q25, Q75	12.5, 22.0	12.0, 21.0	12.0, 23.0
	Min, Max	2, 51	4, 50	5, 36
Day 0, 04:00-06:00 PM	n	80	41	39
	BLOQ - n (%)	0	0	0
	Geometric Mean (CV%)	14.6 (48.6%)	15.1 (47.1%)	15.6 (38.3%)
	95% CI	13.1, 16.2	13.0, 17.4	13.8, 17.7
	Median	14.0	15.0	16.0
	Q25, Q75	10.5, 20.5	12.0, 20.0	12.0, 21.0
	Min, Max	6, 41	5, 48	7, 33

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: T₀ = Time of first product use on study day.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_co.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.7 Descriptive Statistics of Exhaled CO (ppm) - FAS**

Timepoint	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)
Day 0, 08:00-10:00 PM	n	80	41	39
	BLOQ - n (%)	0	0	0
	Geometric Mean (CV%)	15.4 (54.8%)	15.6 (36.5%)	16.9 (42.1%)
	95% CI	13.7, 17.3	13.9, 17.5	14.8, 19.3
	Median	16.0	14.0	18.0
	Q25, Q75	11.5, 20.0	13.0, 18.0	13.0, 23.0
	Min, Max	1, 42	7, 43	5, 34
Day 1, 15 min < T ₀	n	80	41	0
	BLOQ - n (%)	0	0	
	Geometric Mean (CV%)	8.5 (40.1%)	8.2 (46.0%)	
	95% CI	7.8, 9.3	7.1, 9.5	
	Median	8.0	8.0	
	Q25, Q75	7.0, 12.0	6.0, 12.0	
	Min, Max	2, 18	3, 17	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: T₀ = Time of first product use on study day.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_co.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.7 Descriptive Statistics of Exhaled CO (ppm) - FAS**

Timepoint	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)
Day 1, 08:00-10:00 AM	n	0	0	39
	BLOQ - n (%)			0
	Geometric Mean (CV%)			7.7 (37.3%)
	95% CI			6.8, 8.7
	Median			8.0
	Q25, Q75			6.0, 10.0
	Min, Max			4, 15
Day 1, 12:00-02:00 PM	n	80	41	39
	BLOQ - n (%)	0	0	0
	Geometric Mean (CV%)	5.3 (49.2%)	13.4 (39.1%)	5.5 (52.7%)
	95% CI	4.8, 6.0	11.9, 15.2	4.6, 6.5
	Median	6.0	14.0	6.0
	Q25, Q75	4.0, 7.0	10.0, 17.0	4.0, 7.0
	Min, Max	1, 12	7, 34	0, 12

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: T₀ = Time of first product use on study day.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_co.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.7 Descriptive Statistics of Exhaled CO (ppm) - FAS**

Timepoint	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)
Day 1, 04:00-06:00 PM	n	80	41	39
	BLOQ - n (%)	0	0	0
	Geometric Mean (CV%)	3.7 (59.5%)	13.5 (43.4%)	4.1 (48.6%)
	95% CI	3.3, 4.3	11.8, 15.4	3.5, 4.9
	Median	4.0	14.0	4.0
	Q25, Q75	3.0, 6.0	10.0, 17.0	3.0, 5.0
	Min, Max	0, 10	5, 32	1, 9
Day 1, 08:00-10:00 PM	n	80	41	39
	BLOQ - n (%)	0	0	0
	Geometric Mean (CV%)	3.2 (57.7%)	11.9 (47.8%)	3.1 (53.5%)
	95% CI	2.8, 3.6	10.2, 13.7	2.6, 3.7
	Median	3.0	11.0	3.0
	Q25, Q75	2.0, 4.5	9.0, 16.0	2.0, 5.0
	Min, Max	1, 34	5, 42	0, 8

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: T₀ = Time of first product use on study day.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_co.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.7 Descriptive Statistics of Exhaled CO (ppm) - FAS**

Timepoint	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)
Day 2, 15 min < T ₀	n	80	41	0
	BLOQ - n (%)	0	0	
	Geometric Mean (CV%)	3.1 (40.1%)	8.1 (44.8%)	
	95% CI	2.8, 3.4	7.1, 9.4	
	Median	3.0	8.0	
	Q25, Q75	2.0, 4.0	6.0, 11.0	
	Min, Max	1, 6	3, 21	
Day 2, 08:00-10:00 AM	n	0	0	39
	BLOQ - n (%)			0
	Geometric Mean (CV%)			2.9 (42.7%)
	95% CI			2.5, 3.3
	Median			3.0
	Q25, Q75			2.0, 4.0
	Min, Max			1, 6

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: T₀ = Time of first product use on study day.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_co.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.7 Descriptive Statistics of Exhaled CO (ppm) - FAS**

Timepoint	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)
Day 2, 12:00-02:00 PM	n	80	41	39
	BLOQ - n (%)	0	0	0
	Geometric Mean (CV%)	2.7 (49.2%)	11.6 (56.1%)	2.5 (47.6%)
	95% CI	2.4, 3.1	9.8, 13.7	2.1, 3.0
	Median	3.0	12.0	3.0
	Q25, Q75	2.0, 4.0	9.0, 15.0	2.0, 4.0
	Min, Max	0, 7	2, 34	0, 6
Day 2, 04:00-06:00 PM	n	80	41	39
	BLOQ - n (%)	0	0	0
	Geometric Mean (CV%)	2.7 (53.5%)	13.0 (45.5%)	2.4 (50.8%)
	95% CI	2.4, 3.1	11.3, 14.9	2.0, 2.8
	Median	3.0	14.0	2.0
	Q25, Q75	2.0, 4.0	10.0, 18.0	2.0, 3.0
	Min, Max	1, 9	5, 37	0, 6

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: T₀ = Time of first product use on study day.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_co.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.7 Descriptive Statistics of Exhaled CO (ppm) - FAS**

Timepoint	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)
Day 2, 08:00-10:00 PM	n	80	41	39
	BLOQ - n (%)	0	0	0
	Geometric Mean (CV%)	2.3 (57.4%)	13.2 (44.7%)	2.5 (60.0%)
	95% CI	2.0, 2.7	11.5, 15.2	2.0, 3.1
	Median	2.0	13.0	2.0
	Q25, Q75	1.5, 3.0	10.0, 18.0	1.0, 4.0
	Min, Max	0, 8	6, 34	0, 8
Day 3, 15 min < T ₀	n	80	41	0
	BLOQ - n (%)	0	0	
	Geometric Mean (CV%)	2.9 (49.6%)	7.6 (45.5%)	
	95% CI	2.5, 3.2	6.6, 8.7	
	Median	3.0	7.0	
	Q25, Q75	2.0, 4.0	6.0, 9.0	
	Min, Max	1, 7	3, 18	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: T₀ = Time of first product use on study day.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_co.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.7 Descriptive Statistics of Exhaled CO (ppm) - FAS**

Timepoint	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)
Day 3, 08:00-10:00 AM	n	0	0	39
	BLOQ - n (%)			0
	Geometric Mean (CV%)			2.8 (46.5%)
	95% CI			2.4, 3.3
	Median			3.0
	Q25, Q75			2.0, 4.0
	Min, Max			1, 5
Day 3, 12:00-02:00 PM	n	80	41	39
	BLOQ - n (%)	0	0	0
	Geometric Mean (CV%)	2.9 (50.2%)	12.1 (44.3%)	2.6 (51.7%)
	95% CI	2.5, 3.2	10.6, 13.9	2.2, 3.1
	Median	3.0	12.0	3.0
	Q25, Q75	2.0, 4.0	8.0, 16.0	2.0, 4.0
	Min, Max	0, 6	6, 31	0, 7

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: T₀ = Time of first product use on study day.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_co.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.7 Descriptive Statistics of Exhaled CO (ppm) - FAS**

Timepoint	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)
Day 3, 04:00-06:00 PM	n	80	41	39
	BLOQ - n (%)	0	0	0
	Geometric Mean (CV%)	2.9 (48.8%)	14.8 (37.1%)	2.5 (51.4%)
	95% CI	2.6, 3.3	13.2, 16.7	2.1, 3.0
	Median	3.0	14.0	2.0
	Q25, Q75	2.0, 4.0	12.0, 18.0	2.0, 4.0
	Min, Max	0, 8	8, 42	0, 7
Day 3, 08:00-10:00 PM	n	79	41	39
	BLOQ - n (%)	0	0	0
	Geometric Mean (CV%)	2.6 (52.9%)	13.8 (40.0%)	2.0 (55.8%)
	95% CI	2.2, 2.9	12.2, 15.7	1.7, 2.5
	Median	3.0	13.0	2.0
	Q25, Q75	2.0, 4.0	11.0, 17.0	1.0, 3.0
	Min, Max	0, 6	6, 35	0, 5

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: T₀ = Time of first product use on study day.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_co.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.7 Descriptive Statistics of Exhaled CO (ppm) - FAS**

Timepoint	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)
Day 4, 15 min < T ₀	n	79	41	0
	BLOQ - n (%)	0	0	
	Geometric Mean (CV%)	2.9 (42.2%)	8.6 (35.8%)	
	95% CI	2.6, 3.2	7.7, 9.7	
	Median	3.0	9.0	
	Q25, Q75	2.0, 4.0	7.0, 11.0	
	Min, Max	1, 6	4, 17	
Day 4, 08:00-10:00 AM	n	0	0	39
	BLOQ - n (%)			0
	Geometric Mean (CV%)			2.7 (36.5%)
	95% CI			2.4, 3.1
	Median			3.0
	Q25, Q75			2.0, 4.0
	Min, Max			0, 5

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: T₀ = Time of first product use on study day.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_co.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.7 Descriptive Statistics of Exhaled CO (ppm) - FAS**

Timepoint	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)
Day 4, 12:00-02:00 PM	n	79	41	39
	BLOQ - n (%)	0	0	0
	Geometric Mean (CV%)	2.9 (43.7%)	12.2 (45.7%)	2.3 (48.3%)
	95% CI	2.6, 3.2	10.6, 14.1	1.9, 2.8
	Median	3.0	13.0	2.0
	Q25, Q75	2.0, 4.0	9.0, 16.0	2.0, 3.0
	Min, Max	1, 8	5, 25	0, 6
Day 4, 04:00-06:00 PM	n	79	41	39
	BLOQ - n (%)	0	0	0
	Geometric Mean (CV%)	2.8 (50.4%)	13.3 (51.5%)	2.7 (41.5%)
	95% CI	2.5, 3.2	11.4, 15.6	2.3, 3.1
	Median	3.0	13.0	2.0
	Q25, Q75	2.0, 4.0	9.0, 19.0	2.0, 4.0
	Min, Max	0, 7	3, 33	0, 5

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: T₀ = Time of first product use on study day.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_co.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.7 Descriptive Statistics of Exhaled CO (ppm) - FAS**

Timepoint	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)
Day 4, 08:00-10:00 PM	n	79	41	39
	BLOQ - n (%)	0	0	0
	Geometric Mean (CV%)	2.6 (44.6%)	14.2 (39.1%)	2.5 (46.2%)
	95% CI	2.3, 2.9	12.6, 16.1	2.1, 2.9
	Median	2.0	14.0	2.0
	Q25, Q75	2.0, 3.0	10.0, 18.0	2.0, 3.0
	Min, Max	0, 6	7, 32	0, 5
Day 5, 15 min < T ₀	n	79	41	0
	BLOQ - n (%)	0	0	
	Geometric Mean (CV%)	2.3 (48.3%)	7.3 (42.8%)	
	95% CI	2.0, 2.6	6.4, 8.4	
	Median	2.0	7.0	
	Q25, Q75	2.0, 3.0	6.0, 9.0	
	Min, Max	0, 5	3, 18	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: T₀ = Time of first product use on study day.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_co.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.7 Descriptive Statistics of Exhaled CO (ppm) - FAS**

Timepoint	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)
Day 5, 08:00-10:00 AM	n	0	0	39
	BLOQ - n (%)			0
	Geometric Mean (CV%)			2.0 (41.2%)
	95% CI			1.7, 2.3
	Median			2.0
	Q25, Q75			2.0, 3.0
	Min, Max			1, 4
Day 5, 12:00-02:00 PM	n	79	41	39
	BLOQ - n (%)	0	0	0
	Geometric Mean (CV%)	2.1 (43.4%)	11.2 (48.3%)	2.1 (52.0%)
	95% CI	1.9, 2.4	9.7, 13.0	1.7, 2.6
	Median	2.0	10.0	2.0
	Q25, Q75	1.0, 3.0	8.0, 15.0	1.0, 3.0
	Min, Max	0, 4	4, 26	0, 7

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: T₀ = Time of first product use on study day.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_co.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.7 Descriptive Statistics of Exhaled CO (ppm) - FAS**

Timepoint	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)
Day 5, 04:00-06:00 PM	n	79	41	39
	BLOQ - n (%)	0	0	0
	Geometric Mean (CV%)	2.2 (51.0%)	11.8 (53.7%)	2.0 (45.9%)
	95% CI	1.9, 2.5	10.1, 13.9	1.7, 2.4
	Median	2.0	11.0	2.0
	Q25, Q75	1.0, 3.0	9.0, 17.0	1.0, 3.0
	Min, Max	0, 4	2, 45	0, 5
Day 5, 08:00-10:00 PM	n	79	41	39
	BLOQ - n (%)	0	0	0
	Geometric Mean (CV%)	2.0 (46.5%)	12.3 (52.4%)	2.2 (51.5%)
	95% CI	1.8, 2.3	10.5, 14.4	1.8, 2.6
	Median	2.0	11.0	2.0
	Q25, Q75	1.0, 3.0	9.0, 16.0	1.0, 3.0
	Min, Max	0, 5	5, 39	0, 6

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: T₀ = Time of first product use on study day.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_co.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.7 Descriptive Statistics of Exhaled CO (ppm) - FAS**

Timepoint	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)
Day 6/Discharge	n	79	41	39
	BLOQ - n (%)	0	0	0
	Geometric Mean (CV%)	2.3 (38.4%)	6.5 (41.8%)	2.1 (45.8%)
	95% CI	2.1, 2.6	5.6, 7.4	1.8, 2.4
	Median	2.0	7.0	2.0
	Q25, Q75	2.0, 3.0	5.0, 8.0	2.0, 3.0
	Min, Max	0, 4	3, 19	1, 4

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: T₀ = Time of first product use on study day.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_co.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.8 Descriptive Statistics of Time Matched Changes (%) in Exhaled CO - FAS**

Timepoint	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)
Day 1, 15 min < T ₀	n	80	41	0
	Mean (SD)	12.74 (36.973)	22.74 (80.317)	
	95% CI	4.50, 20.97	-2.62, 48.10	
	Median	14.29	0.00	
	Q25, Q75	-12.50, 33.34	-18.19, 33.34	
	Min, Max	-46.2, 160.0	-50.0, 433.3	
Day 1, 12:00-02:00 PM	n	80	41	39
	Mean (SD)	-62.94 (22.968)	-2.89 (49.999)	-61.94 (19.505)
	95% CI	-68.06, -57.82	-18.68, 12.89	-68.27, -55.62
	Median	-66.67	-14.29	-60.00
	Q25, Q75	-75.00, -57.73	-30.00, 5.00	-77.78, -52.63
	Min, Max	-88.9, 100.0	-53.3, 225.0	-100.0, 0.0
Day 1, 04:00-06:00 PM	n	80	41	39
	Mean (SD)	-72.28 (11.988)	-2.72 (43.835)	-71.35 (11.616)
	95% CI	-74.95, -69.60	-16.56, 11.12	-75.12, -67.58
	Median	-72.47	-9.09	-73.33
	Q25, Q75	-81.54, -64.70	-30.44, 12.50	-80.00, -66.66
	Min, Max	-100.0, -40.0	-64.3, 180.0	-90.9, -37.5

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: T₀ = Time of first product use on study day.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_cotmc.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.8 Descriptive Statistics of Time Matched Changes (%) in Exhaled CO - FAS**

Timepoint	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)
Day 1, 08:00-10:00 PM	n	80	41	39
	Mean (SD)	-71.83 (45.359)	-19.99 (25.081)	-79.62 (13.786)
	95% CI	-81.93, -61.73	-27.91, -12.07	-84.10, -75.15
	Median	-79.47	-16.67	-82.14
	Q25, Q75	-85.72, -71.82	-36.67, -2.32	-86.96, -73.68
	Min, Max	-97.1, 277.8	-65.2, 33.3	-100.0, -20.0
Day 2, 15 min < T ₀	n	80	41	0
	Mean (SD)	-57.62 (17.580)	17.12 (53.821)	
	95% CI	-61.53, -53.70	0.13, 34.11	
	Median	-60.00	0.00	
	Q25, Q75	-69.62, -50.00	-18.19, 33.34	
	Min, Max	-83.3, 0.0	-50.0, 233.3	
Day 2, 12:00-02:00 PM	n	80	41	39
	Mean (SD)	-80.30 (13.410)	-12.10 (52.665)	-82.15 (11.387)
	95% CI	-83.29, -77.31	-28.73, 4.53	-85.85, -78.46
	Median	-81.82	-22.73	-84.62
	Q25, Q75	-87.87, -76.47	-41.67, 0.00	-90.48, -75.00
	Min, Max	-100.0, 0.0	-75.0, 160.0	-100.0, -40.0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: T₀ = Time of first product use on study day.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_cotmc.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.8 Descriptive Statistics of Time Matched Changes (%) in Exhaled CO - FAS**

Timepoint	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)
Day 2, 04:00-06:00 PM	n	80	41	39
	Mean (SD)	-78.08 (12.408)	-4.50 (46.724)	-83.92 (10.429)
	95% CI	-80.85, -75.32	-19.25, 10.25	-87.31, -80.53
	Median	-81.53	-7.69	-86.67
	Q25, Q75	-86.96, -70.58	-40.00, 14.29	-90.48, -78.57
	Min, Max	-95.7, -43.8	-63.2, 180.0	-100.0, -50.0
Day 2, 08:00-10:00 PM	n	80	41	39
	Mean (SD)	-82.66 (14.273)	-10.89 (29.592)	-83.86 (13.888)
	95% CI	-85.84, -79.48	-20.23, -1.54	-88.37, -79.36
	Median	-85.71	-16.67	-86.96
	Q25, Q75	-90.24, -78.57	-30.00, 8.00	-91.67, -76.92
	Min, Max	-100.0, 0.0	-50.0, 90.0	-100.0, -20.0
Day 3, 15 min < T ₀	n	80	41	0
	Mean (SD)	-59.21 (21.611)	15.10 (80.395)	
	95% CI	-64.03, -54.40	-10.28, 40.48	
	Median	-66.67	-9.09	
	Q25, Q75	-72.73, -50.00	-27.28, 33.34	
	Min, Max	-90.9, 20.0	-62.5, 433.3	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: T₀ = Time of first product use on study day.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_cotmc.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.8 Descriptive Statistics of Time Matched Changes (%) in Exhaled CO - FAS**

Timepoint	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)
Day 3, 12:00-02:00 PM	n	80	41	39
	Mean (SD)	-80.12 (17.284)	-12.24 (39.778)	-83.21 (9.332)
	95% CI	-83.98, -76.27	-24.80, 0.32	-86.24, -80.18
	Median	-82.84	-20.00	-84.21
	Q25, Q75	-87.75, -75.73	-42.86, 0.00	-88.58, -78.57
	Min, Max	-100.0, 50.0	-64.3, 111.1	-100.0, -63.2
Day 3, 04:00-06:00 PM	n	80	41	39
	Mean (SD)	-77.26 (12.718)	2.90 (33.914)	-83.00 (10.971)
	95% CI	-80.10, -74.43	-7.81, 13.61	-86.56, -79.44
	Median	-78.42	0.00	-83.33
	Q25, Q75	-86.24, -70.29	-15.00, 18.75	-90.33, -77.41
	Min, Max	-100.0, -16.7	-47.4, 160.0	-100.0, -50.0
Day 3, 08:00-10:00 PM	n	79	41	39
	Mean (SD)	-80.00 (23.323)	-8.07 (24.844)	-87.77 (8.173)
	95% CI	-85.23, -74.77	-15.92, -0.22	-90.43, -85.12
	Median	-85.00	-14.29	-88.24
	Q25, Q75	-90.00, -75.00	-25.00, 12.50	-94.74, -84.00
	Min, Max	-100.0, 100.0	-56.5, 43.8	-100.0, -69.2

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: T₀ = Time of first product use on study day.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_cotmc.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.8 Descriptive Statistics of Time Matched Changes (%) in Exhaled CO - FAS**

Timepoint	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)
Day 4, 15 min < T ₀	n	79	41	0
	Mean (SD)	-58.80 (21.605)	27.63 (73.645)	
	95% CI	-63.64, -53.95	4.38, 50.88	
	Median	-66.67	0.00	
	Q25, Q75	-73.34, -50.00	-16.67, 60.00	
	Min, Max	-91.7, 0.0	-44.4, 366.7	
Day 4, 12:00-02:00 PM	n	79	41	39
	Mean (SD)	-78.36 (18.225)	-11.11 (42.541)	-82.93 (18.487)
	95% CI	-82.45, -74.27	-24.54, 2.33	-88.93, -76.93
	Median	-81.25	-25.00	-84.00
	Q25, Q75	-87.50, -75.00	-37.50, 5.56	-91.31, -80.00
	Min, Max	-93.8, 50.0	-65.2, 125.0	-100.0, 20.0
Day 4, 04:00-06:00 PM	n	79	41	39
	Mean (SD)	-78.41 (13.375)	-3.93 (40.277)	-83.15 (10.930)
	95% CI	-81.41, -75.40	-16.65, 8.79	-86.70, -79.60
	Median	-81.82	-7.69	-84.21
	Q25, Q75	-87.50, -72.22	-30.44, 14.29	-90.33, -76.47
	Min, Max	-100.0, -28.6	-78.6, 140.0	-100.0, -50.0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: T₀ = Time of first product use on study day.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_cotmc.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.8 Descriptive Statistics of Time Matched Changes (%) in Exhaled CO - FAS**

Timepoint	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)
Day 4, 08:00-10:00 PM	n	79	41	39
	Mean (SD)	-81.92 (13.461)	-4.71 (29.010)	-85.45 (6.789)
	95% CI	-84.94, -78.90	-13.87, 4.45	-87.66, -83.25
	Median	-84.62	-8.33	-85.71
	Q25, Q75	-89.29, -77.77	-23.08, 8.00	-90.00, -81.25
	Min, Max	-100.0, 0.0	-50.0, 75.0	-100.0, -71.4
Day 5, 15 min < T ₀	n	79	41	0
	Mean (SD)	-67.14 (19.978)	10.97 (58.524)	
	95% CI	-71.62, -62.66	-7.51, 29.45	
	Median	-71.43	-12.50	
	Q25, Q75	-80.00, -57.14	-27.28, 44.45	
	Min, Max	-100.0, 0.0	-72.7, 166.7	
Day 5, 12:00-02:00 PM	n	79	41	39
	Mean (SD)	-86.26 (12.416)	-16.84 (43.850)	-85.91 (10.427)
	95% CI	-89.04, -83.47	-30.69, -3.00	-89.30, -82.53
	Median	-87.50	-30.77	-88.00
	Q25, Q75	-92.60, -82.35	-45.00, 0.00	-92.31, -80.00
	Min, Max	-100.0, 0.0	-71.4, 125.0	-100.0, -60.0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: T₀ = Time of first product use on study day.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_cotmc.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.8 Descriptive Statistics of Time Matched Changes (%) in Exhaled CO - FAS**

Timepoint	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)
Day 5, 04:00-06:00 PM	n	79	41	39
	Mean (SD)	-83.50 (10.281)	-15.16 (33.046)	-86.96 (7.731)
	95% CI	-85.81, -81.19	-25.60, -4.73	-89.47, -84.45
	Median	-85.71	-21.43	-87.50
	Q25, Q75	-90.91, -76.92	-35.72, -6.66	-91.31, -83.33
	Min, Max	-100.0, -50.0	-71.4, 80.0	-100.0, -62.5
Day 5, 08:00-10:00 PM	n	79	41	39
	Mean (SD)	-85.48 (12.643)	-16.59 (27.444)	-88.22 (8.738)
	95% CI	-88.32, -82.64	-25.25, -7.92	-91.06, -85.38
	Median	-87.50	-21.43	-88.89
	Q25, Q75	-92.31, -80.00	-31.82, 4.17	-94.45, -84.00
	Min, Max	-100.0, 0.0	-69.6, 33.3	-100.0, -68.4

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: T₀ = Time of first product use on study day.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_cotmc.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.9 Analysis of Urinary 1-OHP on Day 5 - FAS**

Variable	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	THS : CC Ratio (%)	THS : SA Ratio (%)
Urinary Quantity over n 24 hours (ng)		79	41	39		
	Geometric LS Mean (SE) ¹	117.52	270.63	133.23	43.43 (0.072)	88.21 (0.074)
	95% CI	108.15, 127.71	241.14, 303.71	118.37, 149.96	37.67, 50.06	76.33, 101.93
Concentration Adjusted for Creatinine (pg/mg creat)	n	79	41	39		
	Geometric LS Mean (SE) ¹	81.48	183.70	84.32	44.35 (0.054)	96.63 (0.055)
	95% CI	76.57, 86.70	168.53, 200.23	77.19, 92.12	39.88, 49.33	86.73, 107.65

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Adjusted geometric least squares (LS) means and confidence intervals (CIs) from an ANCOVA model conducted on log-transformed Day 5 values with log-transformed baseline value, study arm, sex and CC consumption reported at screening as fixed effect factors.

Baseline is defined as the last assessment prior to Day 1 product use for THS 2.2 and CC subjects and prior to 06:29 AM on Day 1 for SA subjects.

Note: 1 = Standard errors presented from analysis performed on the log_e scale.

Note: Collection over 24 hours starting on the Day stated in the title.

Appendix 15.4.4.9 and 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/tl_anlsecondbio.sas

Program Run: 27OCT14 cvn_ahedge Program Status: FINAL

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**Table 15.2.4.10 Descriptive Statistics of Urinary 1-OHP - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Quantity Excreted over 24 hours (ng)	Baseline	n	80		41	
		BLOQ - n (%)	0		0	
		Geometric Mean (CV%)	349.06 (44.26%)		359.90 (37.84%)	
		Geometric 95% CI	317.70, 383.51		320.65, 403.96	
		Median	373.30		312.80	
		Q25, Q75	263.10, 471.75		273.07, 488.05	
		Min, Max	119.5, 786.3		210.1, 731.0	
		Mean (SD)	379.26 (150.891)		385.19 (150.156)	
		95% CI				

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (10.0 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_1ohp.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.10 Descriptive Statistics of Urinary 1-OHP - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Quantity Excreted over 24 hours (ng)	Day 1	n	80	80	41	41
		BLOQ - n (%)	0		0	
		Geometric Mean (CV%)	206.99 (41.39%)		310.19 (38.27%)	
		Geometric 95% CI	189.46, 226.14		276.02, 348.59	
		Median	207.88	-42.69	313.20	-13.72
		Q25, Q75	162.70, 270.48	-47.81, -34.69	215.00, 429.40	-22.22, -5.36
		Min, Max	83.7, 481.8	-64.0, 20.3	166.8, 666.4	-45.3, 47.4
		Mean (SD)	223.11 (85.985)	-39.00 (15.772)	331.44 (122.614)	-12.32 (17.317)
		95% CI		-42.51, -35.49		-17.79, -6.85

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (10.0 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_1ohp.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.10 Descriptive Statistics of Urinary 1-OHP - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Quantity Excreted over 24 hours (ng)	Day 2	n	80	80	41	41
		BLOQ - n (%)	0		0	
		Geometric Mean (CV%)	194.76 (43.17%)		333.53 (37.89%)	
		Geometric 95% CI	177.63, 213.53		297.12, 374.41	
		Median	194.04	-43.62	340.43	-9.38
		Q25, Q75	153.92, 252.12	-53.71, -35.93	253.50, 448.40	-21.93, 1.37
		Min, Max	69.6, 621.6	-72.6, 31.1	161.0, 725.0	-45.6, 112.9
		Mean (SD)	211.94 (92.835)	-41.24 (20.317)	355.88 (130.657)	-3.62 (30.509)
		95% CI		-45.77, -36.72		-13.25, 6.02

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (10.0 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_1ohp.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.10 Descriptive Statistics of Urinary 1-OHP - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Quantity Excreted over 24 hours (ng)	Day 3	n	79	79	41	41
		BLOQ - n (%)	0		0	
		Geometric Mean (CV%)	157.96 (44.88%)		295.73 (41.49%)	
		Geometric 95% CI	143.51, 173.87		260.77, 335.37	
		Median	151.34	-57.58	282.00	-20.40
		Q25, Q75	120.25, 214.40	-63.90, -44.08	215.83, 387.50	-27.07, -4.83
		Min, Max	58.5, 683.1	-74.8, 21.5	156.3, 853.2	-67.5, 70.2
		Mean (SD)	174.96 (97.942)	-52.07 (18.380)	320.83 (142.246)	-14.35 (25.769)
		95% CI		-56.19, -47.95		-22.48, -6.21

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (10.0 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_1ohp.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.10 Descriptive Statistics of Urinary 1-OHP - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Quantity Excreted over 24 hours (ng)	Day 4	n	79	79	41	41
		BLOQ - n (%)	0		0	
		Geometric Mean (CV%)	126.93 (40.69%)		270.74 (39.00%)	
		Geometric 95% CI	116.27, 138.56		240.42, 304.88	
		Median	127.76	-64.86	276.21	-24.41
		Q25, Q75	89.96, 154.72	-70.48, -55.00	195.75, 359.55	-37.50, -12.23
		Min, Max	61.4, 522.0	-79.0, -0.8	137.3, 502.3	-49.7, 69.9
		Mean (SD)	137.84 (65.800)	-61.52 (14.258)	289.46 (104.904)	-22.32 (21.568)
		95% CI		-64.72, -58.32		-29.13, -15.51

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (10.0 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_1ohp.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.10 Descriptive Statistics of Urinary 1-OHP - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Quantity Excreted over 24 hours (ng)	Day 5	n	79	79	41	41
		BLOQ - n (%)	1 (1.3%)		0	
		Geometric Mean (CV%)	116.61 (50.79%)		274.61 (46.82%)	
		Geometric 95% CI	104.74, 129.82		238.61, 316.04	
		Median	115.50	-66.63	279.96	-25.27
		Q25, Q75	89.10, 158.65	-73.04, -59.09	185.40, 400.73	-40.99, -8.10
		Min, Max	20.0, 473.0	-93.6, -10.4	135.0, 723.8	-63.6, 216.1
		Mean (SD)	129.85 (64.016)	-64.00 (14.401)	301.88 (133.126)	-17.26 (44.017)
		95% CI		-67.23, -60.77		-31.16, -3.36

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (10.0 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_1ohp.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.10 Descriptive Statistics of Urinary 1-OHP - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Baseline	n	80		41	
		BLOQ - n (%)	0		0	
		Geometric Mean (CV%)	217.69 (44.91%)		218.41 (34.83%)	
		Geometric 95% CI	197.88, 239.48		196.28, 243.03	
		Median	215.25		216.88	
		Q25, Q75	163.86, 304.95		180.02, 261.88	
		Min, Max	81.2, 524.1		106.3, 476.8	
		Mean (SD)	237.95 (101.712)		230.84 (79.290)	
		95% CI				

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (10.0 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_1ohp.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.4.10 Descriptive Statistics of Urinary 1-OHP - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Day 1	n	80	80	41	41
		BLOQ - n (%)	0		0	
		Geometric Mean (CV%)	143.90 (36.15%)		212.74 (36.83%)	
		Geometric 95% CI	133.10, 155.58		190.09, 238.09	
		Median	145.42	-34.70	218.13	-2.59
		Q25, Q75	110.65, 195.30	-41.73, -28.00	174.78, 259.22	-9.33, 7.98
		Min, Max	66.9, 280.3	-64.0, 38.4	84.4, 489.2	-35.3, 32.9
		Mean (SD)	152.71 (52.540)	-32.52 (14.677)	226.09 (81.208)	-1.56 (14.179)
		95% CI		-35.79, -29.25		-6.04, 2.92

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (10.0 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_1ohp.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.10 Descriptive Statistics of Urinary 1-OHP - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Day 2	n	80	80	41	41
		BLOQ - n (%)	0		0	
		Geometric Mean (CV%)	130.22 (37.21%)		209.13 (36.46%)	
		Geometric 95% CI	120.19, 141.09		187.06, 233.79	
		Median	123.64	-38.26	218.80	-3.86
		Q25, Q75	104.02, 156.97	-50.30, -28.58	173.07, 251.56	-16.92, 7.73
		Min, Max	64.8, 461.5	-69.2, 27.9	88.3, 440.5	-42.4, 62.2
		Mean (SD)	139.54 (59.331)	-37.48 (19.600)	221.72 (75.875)	-2.25 (20.448)
		95% CI		-41.85, -33.11		-8.71, 4.21

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (10.0 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_1ohp.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.4.10 Descriptive Statistics of Urinary 1-OHP - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Day 3	n	79	79	41	41
		BLOQ - n (%)	0		0	
		Geometric Mean (CV%)	106.84 (39.07%)		190.98 (38.40%)	
		Geometric 95% CI	98.19, 116.26		169.88, 214.70	
		Median	106.19	-53.19	197.73	-10.66
		Q25, Q75	81.27, 127.69	-61.53, -41.22	160.76, 241.13	-22.55, 3.64
		Min, Max	50.6, 470.3	-71.9, 75.1	78.8, 415.0	-58.0, 41.0
		Mean (SD)	115.72 (57.235)	-47.82 (22.087)	203.54 (71.894)	-10.24 (20.305)
		95% CI		-52.77, -42.87		-16.65, -3.82

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (10.0 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_1ohp.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.10 Descriptive Statistics of Urinary 1-OHP - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Day 4	n	79	79	41	41
		BLOQ - n (%)	0		0	
		Geometric Mean (CV%)	86.89 (37.42%)		177.56 (40.69%)	
		Geometric 95% CI	80.12, 94.23		156.92, 200.91	
		Median	86.71	-62.01	188.39	-19.52
		Q25, Q75	65.98, 108.53	-68.55, -53.01	156.57, 230.12	-31.71, -4.62
		Min, Max	35.4, 228.1	-81.6, 16.6	59.5, 356.1	-44.0, 14.0
		Mean (SD)	92.95 (37.004)	-57.64 (16.988)	190.04 (66.895)	-17.06 (16.665)
		95% CI		-61.45, -53.83		-22.33, -11.80

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (10.0 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_1ohp.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.10 Descriptive Statistics of Urinary 1-OHP - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Day 5	n	79	79	41	41
		BLOQ - n (%)	1 (1.3%)		0	
		Geometric Mean (CV%)	81.22 (37.86%)		182.85 (41.49%)	
		Geometric 95% CI	74.82, 88.16		161.24, 207.37	
		Median	78.76	-65.70	181.51	-19.88
		Q25, Q75	63.46, 100.36	-70.07, -54.40	155.32, 230.77	-26.97, -3.81
		Min, Max	38.5, 239.9	-84.4, 0.2	79.2, 400.0	-50.9, 72.3
		Mean (SD)	87.10 (35.878)	-60.17 (15.704)	197.22 (78.220)	-13.52 (24.042)
		95% CI			-63.70, -56.65	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (10.0 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_1ohp.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.10 Descriptive Statistics of Urinary 1-OHP - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Quantity Excreted over 24 hours (ng)	Baseline	n	39	
		BLOQ - n (%)	0	
		Geometric Mean (CV%)	353.89 (44.99%)	
		Geometric 95% CI	307.91, 406.73	
		Median	368.15	
		Q25, Q75	272.81, 474.00	
		Min, Max	120.4, 939.8	
		Mean (SD)	385.56 (161.914)	
		95% CI		

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (10.0 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_1ohp.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.10 Descriptive Statistics of Urinary 1-OHP - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Quantity Excreted over 24 hours (ng)	Day 1	n	39	39
		BLOQ - n (%)	0	
		Geometric Mean (CV%)	221.12 (50.32%)	
		Geometric 95% CI	189.56, 257.94	
		Median	215.05	-40.04
		Q25, Q75	163.20, 283.92	-46.71, -33.83
		Min, Max	75.5, 721.5	-56.4, 65.1
		Mean (SD)	248.69 (138.326)	-34.73 (23.128)
		95% CI		-42.23, -27.23

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (10.0 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_1ohp.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.10 Descriptive Statistics of Urinary 1-OHP - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Quantity Excreted over 24 hours (ng)	Day 2	n	39	39
		BLOQ - n (%)	0	
		Geometric Mean (CV%)	212.92 (49.20%)	
		Geometric 95% CI	183.09, 247.61	
		Median	205.68	-43.11
		Q25, Q75	155.04, 305.20	-53.01, -17.55
		Min, Max	88.0, 691.2	-80.4, 76.8
		Mean (SD)	237.61 (122.515)	-35.18 (27.182)
		95% CI		-44.00, -26.37

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (10.0 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_1ohp.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.10 Descriptive Statistics of Urinary 1-OHP - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Quantity Excreted over 24 hours (ng)	Day 3	n	39	39
		BLOQ - n (%)	0	
		Geometric Mean (CV%)	167.59 (48.18%)	
		Geometric 95% CI	144.52, 194.35	
		Median	157.08	-52.77
		Q25, Q75	131.01, 227.34	-64.20, -41.66
		Min, Max	77.2, 514.8	-73.1, 13.2
		Mean (SD)	186.52 (95.086)	-49.75 (19.026)
		95% CI		-55.93, -43.58

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (10.0 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_1ohp.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.10 Descriptive Statistics of Urinary 1-OHP - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Quantity Excreted over 24 hours (ng)	Day 4	n	39	39
		BLOQ - n (%)	0	
		Geometric Mean (CV%)	127.43 (49.22%)	
		Geometric 95% CI	109.57, 148.20	
		Median	116.55	-64.84
		Q25, Q75	95.11, 184.16	-70.52, -57.62
		Min, Max	58.8, 578.9	-85.0, -8.8
		Mean (SD)	144.04 (90.301)	-61.68 (14.725)
		95% CI		-66.46, -56.90

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (10.0 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_1ohp.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.10 Descriptive Statistics of Urinary 1-OHP - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Quantity Excreted over 24 hours (ng)	Day 5	n	39	39
		BLOQ - n (%)	0	
		Geometric Mean (CV%)	133.54 (49.89%)	
		Geometric 95% CI	114.61, 155.59	
		Median	124.26	-64.68
		Q25, Q75	93.60, 172.55	-72.15, -48.66
		Min, Max	69.7, 594.8	-83.0, -6.3
		Mean (SD)	152.08 (100.035)	-58.69 (19.238)
		95% CI		-64.93, -52.45

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (10.0 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_1ohp.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.10 Descriptive Statistics of Urinary 1-OHP - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Baseline	n	39	
		BLOQ - n (%)	0	
		Geometric Mean (CV%)	223.95 (37.87%)	
		Geometric 95% CI	198.89, 252.17	
		Median	230.54	
		Q25, Q75	181.27, 290.27	
		Min, Max	101.7, 473.8	
		Mean (SD)	238.60 (85.689)	
		95% CI		

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (10.0 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_1ohp.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.10 Descriptive Statistics of Urinary 1-OHP - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Day 1	n	39	39
		BLOQ - n (%)	0	
		Geometric Mean (CV%)	148.84 (37.85%)	
		Geometric 95% CI	132.19, 167.59	
		Median	153.47	-33.01
		Q25, Q75	111.66, 190.64	-40.67, -24.99
		Min, Max	65.9, 350.0	-69.4, 14.8
		Mean (SD)	158.93 (60.642)	-31.82 (15.412)
		95% CI		-36.82, -26.82

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (10.0 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_1ohp.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.10 Descriptive Statistics of Urinary 1-OHP - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Day 2	n	39	39
		BLOQ - n (%)	0	
		Geometric Mean (CV%)	138.40 (33.09%)	
		Geometric 95% CI	124.67, 153.65	
		Median	136.42	-36.79
		Q25, Q75	114.72, 182.80	-49.21, -17.50
		Min, Max	70.9, 304.2	-71.0, 5.1
		Mean (SD)	145.66 (48.681)	-35.12 (19.815)
		95% CI		-41.55, -28.70

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (10.0 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_1ohp.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.10 Descriptive Statistics of Urinary 1-OHP - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Day 3	n	39	39
		BLOQ - n (%)	0	
		Geometric Mean (CV%)	109.24 (36.66%)	
		Geometric 95% CI	97.36, 122.57	
		Median	117.99	-48.78
		Q25, Q75	87.42, 134.57	-60.85, -40.93
		Min, Max	53.3, 322.6	-74.0, -9.3
		Mean (SD)	116.44 (46.442)	-49.18 (14.383)
		95% CI		-53.85, -44.51

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (10.0 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_1ohp.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.10 Descriptive Statistics of Urinary 1-OHP - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Day 4	n	39	39
		BLOQ - n (%)	0	
		Geometric Mean (CV%)	86.60 (37.82%)	
		Geometric 95% CI	76.92, 97.50	
		Median	94.38	-60.24
		Q25, Q75	63.05, 104.59	-67.16, -52.34
		Min, Max	49.5, 297.1	-81.0, -37.3
		Mean (SD)	93.09 (42.424)	-59.82 (10.860)
		95% CI		-63.35, -56.30

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (10.0 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_1ohp.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.10 Descriptive Statistics of Urinary 1-OHP - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Day 5	n	39	39
		BLOQ - n (%)	0	
		Geometric Mean (CV%)	85.13 (38.91%)	
		Geometric 95% CI	75.37, 96.15	
		Median	87.27	-61.08
		Q25, Q75	62.99, 100.00	-68.12, -52.88
		Min, Max	41.2, 293.3	-82.4, -20.3
		Mean (SD)	91.90 (43.439)	-59.94 (13.315)
		95% CI		-64.26, -55.62

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (10.0 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_1ohp.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.11 Analysis of Urinary Total NNN on Day 5 - FAS**

Variable	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	THS : CC Ratio (%)	THS : SA Ratio (%)
Urinary Quantity over n 24 hours (ng)		79	41	39		
	Geometric LS Mean (SE) ¹	2.21	9.44	0.25	23.45 (0.160)	896.44 (0.162)
	95% CI	1.84, 2.66	7.31, 12.19	0.19, 0.33	17.12, 32.12	651.32, 1233.82
Concentration Adjusted for Creatinine (pg/mg creat)	n	79	41	39		
	Geometric LS Mean (SE) ¹	1.54	6.37	0.16	24.12 (0.155)	980.18 (0.158)
	95% CI	1.28, 1.84	4.97, 8.17	0.12, 0.21	17.76, 32.75	718.45, 1337.25

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Adjusted geometric least squares (LS) means and confidence intervals (CIs) from an ANCOVA model conducted on log-transformed Day 5 values with log-transformed baseline value, study arm, sex and CC consumption reported at screening as fixed effect factors.

Baseline is defined as the last assessment prior to Day 1 product use for THS 2.2 and CC subjects and prior to 06:29 AM on Day 1 for SA subjects.

Note: 1 = Standard errors presented from analysis performed on the log_e scale.

Note: Collection over 24 hours starting on the Day stated in the title.

Appendix 15.4.4.11 and 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/tl_anlsecondbio.sas

Program Run: 27OCT14 cvn_ahedge Program Status: FINAL

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**Table 15.2.4.12 Descriptive Statistics of Urinary Total NNN - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Quantity Excreted over 24 hours (ng)	Baseline	n	80		41	
		BLOQ - n (%)	0		0	
		Geometric Mean (CV%)	7.71 (100.95%)		7.15 (61.15%)	
		Geometric 95% CI	6.39, 9.29		5.98, 8.54	
		Median	6.94		6.24	
		Q25, Q75	4.85, 11.95		4.85, 11.16	
		Min, Max	0.7, 73.3		3.1, 25.0	
		Mean (SD)	10.99 (11.267)		8.41 (5.267)	
		95% CI				

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.200 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_tnnn.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.12 Descriptive Statistics of Urinary Total NNN - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Quantity Excreted over 24 hours (ng)	Day 1	n	80	80	41	41
		BLOQ - n (%)	3 (3.8%)		0	
		Geometric Mean (CV%)	2.20 (198.59%)		6.92 (57.46%)	
		Geometric 95% CI	1.65, 2.92		5.84, 8.19	
		Median	1.96	-73.13	6.41	0.22
		Q25, Q75	1.23, 3.06	-80.27, -63.00	4.48, 9.55	-20.09, 20.42
		Min, Max	0.2, 1788.8	-96.8, 2339.6	2.9, 21.1	-46.3, 104.5
		Mean (SD)	30.74 (205.753)	-27.16 (291.489)	7.99 (4.600)	1.31 (32.121)
		95% CI		-92.03, 37.71		-8.83, 11.45

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.200 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_tnnn.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.12 Descriptive Statistics of Urinary Total NNN - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Quantity Excreted over 24 hours (ng)	Day 2	n	80	80	41	41
		BLOQ - n (%)	8 (10.0%)		0	
		Geometric Mean (CV%)	1.60 (217.74%)		7.30 (76.25%)	
		Geometric 95% CI	1.18, 2.15		5.89, 9.04	
		Median	1.55	-80.69	7.54	12.22
		Q25, Q75	0.97, 2.32	-87.49, -67.74	4.96, 12.05	-24.81, 41.23
		Min, Max	0.2, 2929.4	-96.9, 3895.3	0.8, 23.3	-84.5, 94.3
		Mean (SD)	41.04 (327.821)	-21.45 (446.463)	8.92 (5.581)	11.46 (42.036)
		95% CI		-120.81, 77.91		-1.82, 24.73

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.200 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_tnnn.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.12 Descriptive Statistics of Urinary Total NNN - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Quantity Excreted over 24 hours (ng)	Day 3	n	79	79	41	41
		BLOQ - n (%)	5 (6.3%)		0	
		Geometric Mean (CV%)	1.72 (198.50%)		6.81 (72.19%)	
		Geometric 95% CI	1.29, 2.29		5.54, 8.36	
		Median	1.66	-77.53	6.40	-2.02
		Q25, Q75	1.06, 2.57	-85.66, -69.11	4.75, 11.05	-19.58, 18.13
		Min, Max	0.2, 1565.6	-96.5, 2035.3	1.4, 24.2	-72.7, 89.7
		Mean (SD)	26.58 (180.671)	-39.01 (254.840)	8.30 (5.401)	2.03 (36.661)
		95% CI		-96.10, 18.08		-9.55, 13.60

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.200 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_tnnn.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.12 Descriptive Statistics of Urinary Total NNN - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Quantity Excreted over 24 hours (ng)	Day 4	n	79	79	41	41
		BLOQ - n (%)	8 (10.1%)		0	
		Geometric Mean (CV%)	1.40 (217.80%)		7.62 (126.40%)	
		Geometric 95% CI	1.03, 1.89		5.59, 10.37	
		Median	1.23	-84.22	6.37	0.07
		Q25, Q75	0.78, 2.03	-89.18, -74.75	5.27, 9.87	-23.60, 14.61
		Min, Max	0.2, 1188.1	-96.5, 2540.2	2.3, 1260.0	-63.0, 25839.3
		Mean (SD)	26.01 (156.342)	-34.65 (314.132)	38.25 (195.619)	630.75 (4035.540)
		95% CI		-105.01, 35.72		-643.03, 1904.53

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.200 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_tnnn.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.12 Descriptive Statistics of Urinary Total NNN - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Quantity Excreted over 24 hours (ng)	Day 5	n	79	79	41	41
		BLOQ - n (%)	6 (7.6%)		0	
		Geometric Mean (CV%)	2.23 (196.96%)		9.00 (70.03%)	
		Geometric 95% CI	1.68, 2.96		7.36, 10.99	
		Median	2.16	-71.63	8.19	20.31
		Q25, Q75	1.34, 3.28	-84.46, -54.53	5.68, 14.65	-7.41, 63.35
		Min, Max	0.2, 5152.0	-96.2, 6926.7	2.4, 28.1	-45.8, 292.8
		Mean (SD)	68.90 (579.324)	25.27 (787.189)	10.88 (6.875)	40.05 (70.772)
		95% CI		-151.05, 201.60		17.71, 62.39

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.200 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_tnnn.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.12 Descriptive Statistics of Urinary Total NNN - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Baseline	n	80		41	
		BLOQ - n (%)	0		0	
		Geometric Mean (CV%)	4.81 (99.22%)		4.34 (68.37%)	
		Geometric 95% CI	3.99, 5.78		3.56, 5.28	
		Median	4.69		4.50	
		Q25, Q75	2.85, 7.66		2.59, 6.72	
		Min, Max	0.5, 49.9		1.4, 14.8	
		Mean (SD)	6.93 (7.869)		5.19 (3.166)	
		95% CI				

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.200 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_tnnn.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.12 Descriptive Statistics of Urinary Total NNN - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Day 1	n	80	80	41	41
		BLOQ - n (%)	3 (3.8%)		0	
		Geometric Mean (CV%)	1.53 (194.35%)		4.74 (59.39%)	
		Geometric 95% CI	1.15, 2.02		3.98, 5.65	
		Median	1.44	-69.77	4.78	9.62
		Q25, Q75	0.90, 2.13	-77.23, -60.23	3.00, 7.06	-7.88, 34.28
		Min, Max	0.2, 1402.1	-95.8, 2709.2	1.7, 19.8	-39.6, 130.9
		Mean (SD)	25.16 (164.804)	-15.76 (344.327)	5.53 (3.393)	13.87 (33.507)
		95% CI			-92.39, 60.87	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.200 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_tnnn.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.12 Descriptive Statistics of Urinary Total NNN - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Day 2	n	80	80	41	41
		BLOQ - n (%)	8 (10.0%)		0	
		Geometric Mean (CV%)	1.07 (212.16%)		4.58 (74.92%)	
		Geometric 95% CI	0.79, 1.43		3.70, 5.65	
		Median	1.06	-80.11	4.67	15.56
		Q25, Q75	0.62, 1.66	-85.05, -66.40	2.81, 6.84	-14.96, 34.97
		Min, Max	0.1, 2277.5	-96.7, 4463.1	0.5, 18.1	-72.4, 82.7
		Mean (SD)	31.79 (255.002)	-12.86 (509.851)	5.56 (3.488)	12.07 (36.287)
		95% CI		-126.33, 100.61		0.61, 23.53

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.200 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_tnnn.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.12 Descriptive Statistics of Urinary Total NNN - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Day 3	n	79	79	41	41
		BLOQ - n (%)	5 (6.3%)		0	
		Geometric Mean (CV%)	1.16 (198.32%)		4.40 (72.04%)	
		Geometric 95% CI	0.87, 1.55		3.58, 5.40	
		Median	1.03	-75.45	4.40	3.44
		Q25, Q75	0.63, 1.80	-85.55, -64.63	2.67, 7.57	-19.69, 36.28
		Min, Max	0.1, 1339.2	-96.3, 2583.2	1.3, 18.4	-66.8, 109.4
		Mean (SD)	21.69 (153.307)	-30.39 (310.798)	5.41 (3.741)	9.38 (41.420)
		95% CI		-100.01, 39.23		-3.70, 22.46

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.200 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_tnnn.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.12 Descriptive Statistics of Urinary Total NNN - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Day 4	n	79	79	41	41
		BLOQ - n (%)	8 (10.1%)		0	
		Geometric Mean (CV%)	0.96 (215.30%)		4.99 (140.80%)	
		Geometric 95% CI	0.71, 1.29		3.59, 6.95	
		Median	0.86	-83.24	5.02	12.18
		Q25, Q75	0.58, 1.40	-87.53, -72.61	2.48, 7.19	-16.15, 31.10
		Min, Max	0.1, 895.3	-96.0, 2496.3	1.6, 1122.2	-75.9, 29696.2
		Mean (SD)	19.81 (119.618)	-31.19 (316.359)	32.39 (174.479)	731.40 (4636.765)
		95% CI		-102.06, 39.67		-732.15, 2194.95

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.200 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_tnnn.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.12 Descriptive Statistics of Urinary Total NNN - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Day 5	n	79	79	41	41
		BLOQ - n (%)	6 (7.6%)		0	
		Geometric Mean (CV%)	1.55 (190.06%)		5.99 (66.67%)	
		Geometric 95% CI	1.17, 2.05		4.94, 7.26	
		Median	1.42	-69.75	5.77	29.93
		Q25, Q75	0.94, 2.29	-82.75, -53.26	3.59, 9.43	12.84, 74.96
		Min, Max	0.1, 5125.3	-96.2, 10168.9	1.9, 26.8	-32.6, 437.8
		Mean (SD)	67.36 (576.416)	69.60 (1151.373)	7.21 (4.867)	53.83 (87.523)
		95% CI	-188.30, 327.50		26.20, 81.46	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.200 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_tnnn.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.12 Descriptive Statistics of Urinary Total NNN - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Quantity Excreted over 24 hours (ng)	Baseline	n	39	
		BLOQ - n (%)	0	
		Geometric Mean (CV%)	8.11 (93.23%)	
		Geometric 95% CI	6.27, 10.49	
		Median	8.70	
		Q25, Q75	4.81, 14.14	
		Min, Max	1.2, 55.5	
		Mean (SD)	10.84 (9.537)	
		95% CI		

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.200 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_tnnn.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.12 Descriptive Statistics of Urinary Total NNN - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Quantity Excreted over 24 hours (ng)	Day 1	n	39	39
		BLOQ - n (%)	9 (23.1%)	
		Geometric Mean (CV%)	0.98 (136.10%)	
		Geometric 95% CI	0.70, 1.37	
		Median	1.30	-86.78
		Q25, Q75	0.57, 1.87	-90.43, -82.34
		Min, Max	0.1, 13.4	-96.4, -75.9
		Mean (SD)	1.59 (2.139)	-86.80 (5.202)
		95% CI		-88.49, -85.11

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.200 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_tnnn.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.12 Descriptive Statistics of Urinary Total NNN - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Quantity Excreted over 24 hours (ng)	Day 2	n	39	39
		BLOQ - n (%)	37 (94.9%)	
		Geometric Mean (CV%)	0.25 (33.28%)	
		Geometric 95% CI	0.22, 0.29	
		Median	0.26	-97.19
		Q25, Q75	0.20, 0.30	-98.26, -94.60
		Min, Max	0.2, 0.8	-99.2, -83.4
		Mean (SD)	0.27 (0.108)	-95.81 (3.733)
		95% CI		-97.03, -94.60

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.200 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_tnnn.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.12 Descriptive Statistics of Urinary Total NNN - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Quantity Excreted over 24 hours (ng)	Day 3	n	39	39
		BLOQ - n (%)	39 (100%)	
		Geometric Mean (CV%)	0.24 (30.91%)	
		Geometric 95% CI	0.21, 0.27	
		Median	0.24	-97.32
		Q25, Q75	0.21, 0.30	-98.41, -95.01
		Min, Max	0.1, 0.4	-99.5, -80.5
		Mean (SD)	0.25 (0.072)	-95.92 (3.804)
		95% CI		-97.16, -94.69

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.200 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_tnnn.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.12 Descriptive Statistics of Urinary Total NNN - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Quantity Excreted over 24 hours (ng)	Day 4	n	39	39
		BLOQ - n (%)	39 (100%)	
		Geometric Mean (CV%)	0.26 (26.33%)	
		Geometric 95% CI	0.24, 0.29	
		Median	0.26	-97.41
		Q25, Q75	0.21, 0.30	-98.12, -94.68
		Min, Max	0.2, 0.5	-99.6, -77.9
		Mean (SD)	0.27 (0.077)	-95.41 (4.848)
		95% CI		-96.99, -93.84

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.200 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_tnnn.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.12 Descriptive Statistics of Urinary Total NNN - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Quantity Excreted over 24 hours (ng)	Day 5	n	39	39
		BLOQ - n (%)	39 (100%)	
		Geometric Mean (CV%)	0.26 (27.59%)	
		Geometric 95% CI	0.23, 0.28	
		Median	0.27	-97.22
		Q25, Q75	0.20, 0.31	-98.46, -94.48
		Min, Max	0.2, 0.4	-99.5, -74.1
		Mean (SD)	0.26 (0.070)	-95.28 (5.171)
		95% CI		-96.96, -93.60

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.200 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_tnnn.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.12 Descriptive Statistics of Urinary Total NNN - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Baseline	n	39	
		BLOQ - n (%)	0	
		Geometric Mean (CV%)	5.14 (97.93%)	
		Geometric 95% CI	3.93, 6.70	
		Median	5.79	
		Q25, Q75	2.89, 9.79	
		Min, Max	0.9, 25.6	
		Mean (SD)	6.84 (5.125)	
		95% CI		

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.200 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_tnnn.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.12 Descriptive Statistics of Urinary Total NNN - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Day 1	n	39	39
		BLOQ - n (%)	9 (23.1%)	
		Geometric Mean (CV%)	0.66 (131.08%)	
		Geometric 95% CI	0.47, 0.91	
		Median	0.80	-86.56
		Q25, Q75	0.32, 1.33	-90.84, -81.48
		Min, Max	0.1, 6.5	-95.9, -74.7
		Mean (SD)	1.02 (1.114)	-86.07 (5.293)
		95% CI		-87.79, -84.35

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.200 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_tnnn.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.12 Descriptive Statistics of Urinary Total NNN - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Day 2	n	39	39
		BLOQ - n (%)	37 (94.9%)	
		Geometric Mean (CV%)	0.17 (37.28%)	
		Geometric 95% CI	0.14, 0.19	
		Median	0.16	-96.86
		Q25, Q75	0.13, 0.21	-98.54, -94.19
		Min, Max	0.1, 0.3	-99.1, -79.4
		Mean (SD)	0.18 (0.066)	-95.50 (4.243)
		95% CI		-96.88, -94.12

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.200 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_tnnn.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.12 Descriptive Statistics of Urinary Total NNN - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Day 3	n	39	39
		BLOQ - n (%)	39 (100%)	
		Geometric Mean (CV%)	0.16 (40.22%)	
		Geometric 95% CI	0.13, 0.18	
		Median	0.16	-97.19
		Q25, Q75	0.11, 0.22	-98.31, -94.47
		Min, Max	0.1, 0.4	-99.4, -76.5
		Mean (SD)	0.17 (0.067)	-95.62 (4.362)
		95% CI		-97.04, -94.20

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.200 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_tnnn.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.12 Descriptive Statistics of Urinary Total NNN - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Day 4	n	39	39
		BLOQ - n (%)	39 (100%)	
		Geometric Mean (CV%)	0.18 (35.36%)	
		Geometric 95% CI	0.15, 0.20	
		Median	0.18	-96.78
		Q25, Q75	0.13, 0.23	-98.30, -94.29
		Min, Max	0.1, 0.4	-99.5, -65.2
		Mean (SD)	0.19 (0.069)	-94.69 (6.596)
		95% CI		-96.83, -92.55

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.200 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_tnnn.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.12 Descriptive Statistics of Urinary Total NNN - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Day 5	n	39	39
		BLOQ - n (%)	39 (100%)	
		Geometric Mean (CV%)	0.16 (39.28%)	
		Geometric 95% CI	0.14, 0.19	
		Median	0.16	-97.04
		Q25, Q75	0.13, 0.22	-98.46, -94.06
		Min, Max	0.1, 0.4	-99.4, -81.5
		Mean (SD)	0.18 (0.070)	-95.31 (4.606)
		95% CI		-96.81, -93.81

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.200 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_tnnn.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.13 Analysis of Urinary 4-ABP on Day 5 - FAS**

Variable	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	THS : CC Ratio (%)	THS : SA Ratio (%)
Urinary Quantity over n 24 hours (ng)		79	41	39		
	Geometric LS Mean (SE) ¹	2.73	18.77	2.52	14.55 (0.077)	108.54 (0.078)
	95% CI	2.50, 2.99	16.61, 21.22	2.22, 2.86	12.51, 16.92	93.11, 126.53
Concentration Adjusted for Creatinine (pg/mg creat)	n	79	41	39		
	Geometric LS Mean (SE) ¹	1.90	12.69	1.60	14.94 (0.075)	118.51 (0.077)
	95% CI	1.73, 2.07	11.25, 14.31	1.41, 1.81	12.89, 17.33	101.96, 137.75

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Adjusted geometric least squares (LS) means and confidence intervals (CIs) from an ANCOVA model conducted on log-transformed Day 5 values with log-transformed baseline value, study arm, sex and CC consumption reported at screening as fixed effect factors.

Baseline is defined as the last assessment prior to Day 1 product use for THS 2.2 and CC subjects and prior to 06:29 AM on Day 1 for SA subjects.

Note: 1 = Standard errors presented from analysis performed on the log_e scale.

Note: Collection over 24 hours starting on the Day stated in the title.

Appendix 15.4.4.13 and 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/tl_anlsecondbio.sas

Program Run: 27OCT14 cvn_ahedge Program Status: FINAL

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**Table 15.2.4.14 Descriptive Statistics of Urinary 4-ABP - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Quantity Excreted over 24 hours (ng)	Baseline	n	80		41	
		BLOQ - n (%)	0		0	
		Geometric Mean (CV%)	21.24 (53.14%)		21.60 (49.79%)	
		Geometric 95% CI	19.01, 23.74		18.61, 25.06	
		Median	23.24		20.77	
		Q25, Q75	17.17, 28.91		14.87, 31.07	
		Min, Max	3.1, 66.0		9.7, 80.3	
		Mean (SD)	23.57 (9.998)		24.22 (13.116)	
		95% CI				

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (1.50 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_4abp.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.14 Descriptive Statistics of Urinary 4-ABP - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Quantity Excreted over 24 hours (ng)	Day 1	n	80	80	41	41
		BLOQ - n (%)	3 (3.8%)		0	
		Geometric Mean (CV%)	8.17 (56.16%)		18.01 (52.62%)	
		Geometric 95% CI	7.26, 9.18		15.41, 21.06	
		Median	8.89	-62.12	17.36	-12.38
		Q25, Q75	6.22, 11.04	-68.57, -53.59	13.03, 26.03	-28.52, -3.28
		Min, Max	1.5, 26.8	-79.4, -5.2	6.3, 71.9	-47.7, 21.9
		Mean (SD)	9.22 (4.527)	-59.77 (13.046)	20.41 (11.619)	-14.83 (17.267)
		95% CI		-62.68, -56.87		-20.29, -9.38

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (1.50 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_4abp.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.14 Descriptive Statistics of Urinary 4-ABP - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Quantity Excreted over 24 hours (ng)	Day 2	n	80	80	41	41
		BLOQ - n (%)	16 (20.0%)		0	
		Geometric Mean (CV%)	4.82 (60.99%)		20.25 (60.79%)	
		Geometric 95% CI	4.24, 5.46		16.96, 24.18	
		Median	5.31	-77.51	18.05	-5.48
		Q25, Q75	3.54, 6.21	-82.30, -72.20	13.60, 31.66	-14.69, 9.22
		Min, Max	1.5, 30.2	-91.8, 31.2	4.8, 73.8	-84.1, 47.8
		Mean (SD)	5.67 (4.034)	-74.79 (15.245)	23.57 (13.801)	-2.01 (24.418)
		95% CI		-78.19, -71.39		-9.72, 5.70

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (1.50 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_4abp.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.14 Descriptive Statistics of Urinary 4-ABP - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Quantity Excreted over 24 hours (ng)	Day 3	n	79	79	41	41
		BLOQ - n (%)	43 (54.4%)		0	
		Geometric Mean (CV%)	2.88 (68.95%)		19.41 (59.87%)	
		Geometric 95% CI	2.50, 3.31		16.29, 23.12	
		Median	2.25	-86.55	19.60	-6.62
		Q25, Q75	1.72, 4.49	-91.55, -80.63	13.05, 30.07	-20.81, 2.76
		Min, Max	1.1, 25.2	-95.0, -23.0	5.0, 74.9	-83.5, 50.0
		Mean (SD)	3.68 (3.832)	-84.03 (11.604)	22.45 (12.854)	-5.96 (24.339)
		95% CI		-86.64, -81.43		-13.65, 1.73

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (1.50 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_4abp.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.14 Descriptive Statistics of Urinary 4-ABP - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Quantity Excreted over 24 hours (ng)	Day 4	n	79	79	41	41
		BLOQ - n (%)	51 (64.6%)		0	
		Geometric Mean (CV%)	2.62 (53.72%)		20.50 (54.08%)	
		Geometric 95% CI	2.34, 2.94		17.47, 24.06	
		Median	2.25	-88.29	20.38	-0.41
		Q25, Q75	1.87, 3.60	-91.77, -83.38	13.60, 28.29	-19.45, 16.01
		Min, Max	1.2, 28.2	-95.2, -20.5	6.9, 82.2	-69.7, 53.5
		Mean (SD)	3.13 (3.171)	-85.61 (11.092)	23.38 (13.647)	-1.88 (23.400)
		95% CI		-88.10, -83.12		-9.27, 5.52

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (1.50 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_4abp.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.14 Descriptive Statistics of Urinary 4-ABP - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Quantity Excreted over 24 hours (ng)	Day 5	n	79	79	41	41
		BLOQ - n (%)	62 (78.5%)		0	
		Geometric Mean (CV%)	2.73 (48.45%)		18.89 (57.97%)	
		Geometric 95% CI	2.46, 3.03		15.93, 22.39	
		Median	2.31	-88.06	20.13	-12.24
		Q25, Q75	1.91, 3.78	-91.24, -83.13	12.43, 28.02	-23.43, 0.63
		Min, Max	1.2, 21.2	-97.2, -28.3	5.7, 78.0	-49.4, 51.1
		Mean (SD)	3.12 (2.401)	-84.80 (11.442)	21.78 (12.834)	-10.47 (19.793)
		95% CI		-87.37, -82.23		-16.72, -4.22

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (1.50 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_4abp.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.14 Descriptive Statistics of Urinary 4-ABP - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Baseline	n	80		41	
		BLOQ - n (%)	0		0	
		Geometric Mean (CV%)	13.25 (53.32%)		13.11 (48.46%)	
		Geometric 95% CI	11.85, 14.81		11.33, 15.16	
		Median	13.54		12.34	
		Q25, Q75	10.35, 18.40		9.68, 16.99	
		Min, Max	2.8, 39.9		5.8, 34.7	
		Mean (SD)	14.87 (7.167)		14.57 (7.145)	
		95% CI				

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (1.50 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_4abp.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.4.14 Descriptive Statistics of Urinary 4-ABP - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Day 1	n	80	80	41	41
		BLOQ - n (%)	3 (3.8%)		0	
		Geometric Mean (CV%)	5.68 (52.17%)		12.35 (46.02%)	
		Geometric 95% CI	5.08, 6.34		10.75, 14.19	
		Median	6.01	-57.01	11.66	-4.58
		Q25, Q75	4.09, 7.74	-62.90, -49.29	9.91, 15.34	-15.17, 7.46
		Min, Max	1.5, 14.3	-78.0, -25.0	5.2, 35.2	-56.4, 35.8
		Mean (SD)	6.34 (2.920)	-55.75 (11.419)	13.62 (6.614)	-3.85 (18.605)
		95% CI	-58.29, -53.20		-9.73, 2.02	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (1.50 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_4abp.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.14 Descriptive Statistics of Urinary 4-ABP - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Day 2	n	80	80	41	41
		BLOQ - n (%)	16 (20.0%)		0	
		Geometric Mean (CV%)	3.22 (62.74%)		12.70 (54.59%)	
		Geometric 95% CI	2.83, 3.67		10.80, 14.92	
		Median	3.06	-75.03	11.96	2.13
		Q25, Q75	2.30, 4.81	-80.86, -67.97	9.63, 18.29	-8.66, 12.27
		Min, Max	0.8, 12.4	-91.6, -13.8	3.1, 35.9	-71.7, 36.6
		Mean (SD)	3.77 (2.186)	-73.49 (12.328)	14.38 (7.467)	-0.12 (21.578)
		95% CI		-76.24, -70.74		-6.94, 6.69

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (1.50 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_4abp.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.4.14 Descriptive Statistics of Urinary 4-ABP - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Day 3	n	79	79	41	41
		BLOQ - n (%)	43 (54.4%)		0	
		Geometric Mean (CV%)	1.95 (67.85%)		12.53 (54.02%)	
		Geometric 95% CI	1.69, 2.24		10.68, 14.71	
		Median	1.95	-85.54	12.60	2.06
		Q25, Q75	1.20, 2.97	-90.85, -79.42	8.52, 16.35	-15.09, 16.55
		Min, Max	0.6, 19.8	-94.4, -32.1	5.3, 42.4	-79.9, 103.0
		Mean (SD)	2.45 (2.468)	-82.93 (11.305)	14.29 (8.157)	0.74 (29.723)
		95% CI		-85.46, -80.39		-8.65, 10.13

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (1.50 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_4abp.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.14 Descriptive Statistics of Urinary 4-ABP - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Day 4	n	79	79	41	41
		BLOQ - n (%)	51 (64.6%)		0	
		Geometric Mean (CV%)	1.80 (50.82%)		13.44 (52.25%)	
		Geometric 95% CI	1.61, 2.01		11.51, 15.70	
		Median	1.73	-87.96	13.47	6.47
		Q25, Q75	1.26, 2.42	-90.63, -81.44	9.87, 18.90	-9.82, 27.66
		Min, Max	0.8, 12.8	-93.9, -20.3	4.9, 42.2	-80.3, 63.0
		Mean (SD)	2.06 (1.503)	-84.39 (11.260)	15.16 (8.058)	7.46 (28.351)
		95% CI		-86.92, -81.86		-1.50, 16.41

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (1.50 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_4abp.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.14 Descriptive Statistics of Urinary 4-ABP - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Day 5	n	79	79	41	41
		BLOQ - n (%)	62 (78.5%)		0	
		Geometric Mean (CV%)	1.90 (51.46%)		12.58 (43.29%)	
		Geometric 95% CI	1.70, 2.12		11.03, 14.34	
		Median	1.69	-86.07	11.61	-1.51
		Q25, Q75	1.30, 2.68	-91.35, -81.02	9.66, 17.37	-16.26, 5.80
		Min, Max	0.8, 9.2	-95.5, -15.0	5.3, 34.2	-58.4, 69.3
		Mean (SD)	2.16 (1.287)	-82.12 (15.483)	13.72 (6.246)	-1.66 (21.931)
		95% CI			-85.59, -78.64	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (1.50 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_4abp.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.14 Descriptive Statistics of Urinary 4-ABP - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Quantity Excreted over 24 hours (ng)	Baseline	n	39	
		BLOQ - n (%)	0	
		Geometric Mean (CV%)	21.38 (43.83%)	
		Geometric 95% CI	18.66, 24.50	
		Median	22.05	
		Q25, Q75	15.04, 30.15	
		Min, Max	8.3, 59.3	
		Mean (SD)	23.25 (9.881)	
		95% CI		

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (1.50 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_4abp.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.14 Descriptive Statistics of Urinary 4-ABP - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Quantity Excreted over 24 hours (ng)	Day 1	n	39	39
		BLOQ - n (%)	1 (2.6%)	
		Geometric Mean (CV%)	8.15 (50.31%)	
		Geometric 95% CI	6.99, 9.52	
		Median	8.72	-63.18
		Q25, Q75	6.17, 11.57	-66.54, -53.05
		Min, Max	1.6, 19.0	-83.1, -24.2
		Mean (SD)	8.98 (3.740)	-60.33 (11.696)
		95% CI		-64.12, -56.53

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (1.50 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_4abp.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.14 Descriptive Statistics of Urinary 4-ABP - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Quantity Excreted over 24 hours (ng)	Day 2	n	39	39
		BLOQ - n (%)	10 (25.6%)	
		Geometric Mean (CV%)	4.57 (59.84%)	
		Geometric 95% CI	3.82, 5.48	
		Median	4.85	-78.88
		Q25, Q75	3.04, 6.69	-84.99, -70.07
		Min, Max	1.4, 15.1	-91.1, -50.7
		Mean (SD)	5.27 (2.822)	-76.72 (9.844)
		95% CI		-79.91, -73.52

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (1.50 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_4abp.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.14 Descriptive Statistics of Urinary 4-ABP - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Quantity Excreted over 24 hours (ng)	Day 3	n	39	39
		BLOQ - n (%)	19 (48.7%)	
		Geometric Mean (CV%)	3.14 (49.46%)	
		Geometric 95% CI	2.69, 3.65	
		Median	3.09	-85.11
		Q25, Q75	2.00, 4.52	-87.65, -80.11
		Min, Max	1.6, 8.9	-95.6, -63.8
		Mean (SD)	3.50 (1.732)	-83.80 (7.084)
		95% CI		-86.11, -81.50

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (1.50 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_4abp.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.14 Descriptive Statistics of Urinary 4-ABP - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Quantity Excreted over 24 hours (ng)	Day 4	n	39	39
		BLOQ - n (%)	27 (69.2%)	
		Geometric Mean (CV%)	2.70 (49.49%)	
		Geometric 95% CI	2.31, 3.15	
		Median	2.40	-87.38
		Q25, Q75	1.87, 3.57	-92.07, -81.54
		Min, Max	1.4, 9.5	-94.2, -65.3
		Mean (SD)	3.03 (1.663)	-85.70 (7.440)
		95% CI		-88.12, -83.29

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (1.50 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_4abp.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.14 Descriptive Statistics of Urinary 4-ABP - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Quantity Excreted over 24 hours (ng)	Day 5	n	39	39
		BLOQ - n (%)	29 (74.4%)	
		Geometric Mean (CV%)	2.51 (36.82%)	
		Geometric 95% CI	2.23, 2.83	
		Median	2.31	-87.56
		Q25, Q75	1.95, 3.14	-92.24, -85.04
		Min, Max	1.4, 6.3	-96.0, -74.3
		Mean (SD)	2.68 (1.075)	-86.98 (5.937)
		95% CI		-88.91, -85.05

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (1.50 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_4abp.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.14 Descriptive Statistics of Urinary 4-ABP - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Baseline	n	39	
		BLOQ - n (%)	0	
		Geometric Mean (CV%)	13.53 (46.71%)	
		Geometric 95% CI	11.71, 15.63	
		Median	12.85	
		Q25, Q75	10.18, 18.26	
		Min, Max	4.9, 37.6	
		Mean (SD)	14.92 (7.050)	
		95% CI		

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (1.50 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_4abp.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.14 Descriptive Statistics of Urinary 4-ABP - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Day 1	n	39	39
		BLOQ - n (%)	1 (2.6%)	
		Geometric Mean (CV%)	5.49 (50.97%)	
		Geometric 95% CI	4.69, 6.42	
		Median	5.33	-58.12
		Q25, Q75	3.87, 8.45	-65.99, -49.24
		Min, Max	1.5, 15.0	-83.8, -32.9
		Mean (SD)	6.12 (2.961)	-57.86 (10.904)
		95% CI		-61.40, -54.32

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (1.50 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_4abp.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.14 Descriptive Statistics of Urinary 4-ABP - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Day 2	n	39	39
		BLOQ - n (%)	10 (25.6%)	
		Geometric Mean (CV%)	2.97 (56.34%)	
		Geometric 95% CI	2.50, 3.53	
		Median	3.12	-77.46
		Q25, Q75	2.18, 4.37	-83.20, -71.54
		Min, Max	1.1, 7.9	-91.6, -57.2
		Mean (SD)	3.38 (1.715)	-76.65 (8.034)
		95% CI		-79.26, -74.04

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (1.50 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_4abp.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.14 Descriptive Statistics of Urinary 4-ABP - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Day 3	n	39	39
		BLOQ - n (%)	19 (48.7%)	
		Geometric Mean (CV%)	2.04 (50.56%)	
		Geometric 95% CI	1.75, 2.39	
		Median	1.97	-84.08
		Q25, Q75	1.42, 2.83	-89.41, -79.83
		Min, Max	0.8, 5.9	-93.5, -58.8
		Mean (SD)	2.29 (1.192)	-83.57 (7.004)
		95% CI		-85.85, -81.29

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (1.50 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_4abp.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.14 Descriptive Statistics of Urinary 4-ABP - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Day 4	n	39	39
		BLOQ - n (%)	27 (69.2%)	
		Geometric Mean (CV%)	1.83 (48.71%)	
		Geometric 95% CI	1.57, 2.14	
		Median	1.72	-87.37
		Q25, Q75	1.32, 2.51	-90.41, -83.93
		Min, Max	0.9, 4.9	-95.2, -51.4
		Mean (SD)	2.05 (1.049)	-84.71 (8.770)
		95% CI		-87.56, -81.86

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (1.50 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_4abp.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.14 Descriptive Statistics of Urinary 4-ABP - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Day 5	n	39	39
		BLOQ - n (%)	29 (74.4%)	
		Geometric Mean (CV%)	1.60 (41.10%)	
		Geometric 95% CI	1.40, 1.83	
		Median	1.60	-88.12
		Q25, Q75	1.15, 2.24	-91.37, -84.09
		Min, Max	0.8, 3.5	-96.0, -70.8
		Mean (SD)	1.73 (0.709)	-86.99 (5.774)
		95% CI		-88.87, -85.12

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (1.50 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_4abp.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.15 Analysis of Urinary 1-NA on Day 5 - FAS**

Variable	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	THS : CC Ratio (%)	THS : SA Ratio (%)
Urinary Quantity over n 24 hours (ng)		79	41	39		
	Geometric LS Mean (SE) ¹	4.80	131.10	4.02	3.66 (0.090)	119.46 (0.091)
	95% CI	4.32, 5.32	113.63, 151.27	3.46, 4.65	3.06, 4.37	99.88, 142.87
Concentration Adjusted for Creatinine (pg/mg creat)	n	79	41	39		
	Geometric LS Mean (SE) ¹	3.32	88.98	2.55	3.73 (0.085)	130.03 (0.087)
	95% CI	3.01, 3.67	77.66, 101.96	2.22, 2.94	3.15, 4.42	109.64, 154.21

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Adjusted geometric least squares (LS) means and confidence intervals (CIs) from an ANCOVA model conducted on log-transformed Day 5 values with log-transformed baseline value, study arm, sex and CC consumption reported at screening as fixed effect factors.

Baseline is defined as the last assessment prior to Day 1 product use for THS 2.2 and CC subjects and prior to 06:29 AM on Day 1 for SA subjects.

Note: 1 = Standard errors presented from analysis performed on the log_e scale.

Note: Collection over 24 hours starting on the Day stated in the title.

Appendix 15.4.4.15 and 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/tl_anlsecondbio.sas

Program Run: 27OCT14 cvn_ahedge Program Status: FINAL

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**Table 15.2.4.16 Descriptive Statistics of Urinary 1-NA - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Quantity Excreted over 24 hours (ng)	Baseline	n	80		41	
		BLOQ - n (%)	0		0	
		Geometric Mean (CV%)	118.38 (49.58%)		128.27 (49.38%)	
		Geometric 95% CI	106.65, 131.41		110.69, 148.66	
		Median	128.00		113.04	
		Q25, Q75	99.91, 150.27		83.38, 168.37	
		Min, Max	21.0, 314.6		73.3, 392.4	
		Mean (SD)	129.75 (51.873)		144.26 (79.697)	
		95% CI				

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (2.50 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_1na.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.16 Descriptive Statistics of Urinary 1-NA - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Quantity Excreted over 24 hours (ng)	Day 1	n	80	80	41	41
		BLOQ - n (%)	1 (1.3%)		0	
		Geometric Mean (CV%)	22.27 (61.47%)		114.33 (55.03%)	
		Geometric 95% CI	19.63, 25.27		97.19, 134.49	
		Median	21.73	-82.10	99.16	-5.55
		Q25, Q75	16.38, 28.74	-85.52, -77.91	73.92, 167.09	-21.23, 2.61
		Min, Max	2.8, 205.9	-91.8, 30.9	40.0, 390.0	-55.4, 41.9
		Mean (SD)	26.89 (25.351)	-79.07 (15.081)	130.81 (74.636)	-8.18 (21.708)
		95% CI		-82.43, -75.71		-15.04, -1.32

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (2.50 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_1na.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.16 Descriptive Statistics of Urinary 1-NA - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Quantity Excreted over 24 hours (ng)	Day 2	n	80	80	41	41
		BLOQ - n (%)	7 (8.8%)		0	
		Geometric Mean (CV%)	10.83 (61.16%)		114.55 (72.05%)	
		Geometric 95% CI	9.55, 12.28		93.40, 140.49	
		Median	11.42	-90.96	108.15	-5.93
		Q25, Q75	8.95, 13.29	-92.84, -89.09	77.71, 174.24	-21.61, 12.63
		Min, Max	2.6, 187.6	-97.7, 25.1	10.2, 554.3	-93.6, 67.7
		Mean (SD)	13.66 (20.248)	-89.09 (13.392)	137.90 (92.718)	-2.86 (32.620)
		95% CI		-92.07, -86.10		-13.16, 7.44

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (2.50 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_1na.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.16 Descriptive Statistics of Urinary 1-NA - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Quantity Excreted over 24 hours (ng)	Day 3	n	79	79	41	41
		BLOQ - n (%)	22 (27.8%)		1 (2.4%)	
		Geometric Mean (CV%)	6.48 (82.58%)		111.87 (87.10%)	
		Geometric 95% CI	5.51, 7.62		88.24, 141.81	
		Median	6.50	-94.57	110.94	-1.83
		Q25, Q75	3.75, 8.40	-96.27, -93.05	81.00, 176.22	-23.95, 15.38
		Min, Max	1.8, 264.0	-98.4, -16.1	3.3, 423.0	-97.1, 46.7
		Mean (SD)	11.28 (31.448)	-92.62 (11.452)	135.62 (79.639)	-4.48 (27.628)
		95% CI		-95.19, -90.05		-13.21, 4.24

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (2.50 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_1na.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.16 Descriptive Statistics of Urinary 1-NA - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Quantity Excreted over 24 hours (ng)	Day 4	n	79	79	41	41
		BLOQ - n (%)	58 (73.4%)		0	
		Geometric Mean (CV%)	4.16 (67.73%)		117.85 (52.52%)	
		Geometric 95% CI	3.62, 4.77		100.85, 137.72	
		Median	3.56	-96.86	106.25	-5.81
		Q25, Q75	3.03, 5.48	-97.64, -95.42	81.13, 171.07	-18.16, 14.26
		Min, Max	2.1, 289.8	-98.8, -7.9	45.8, 342.0	-73.4, 68.1
		Mean (SD)	7.87 (32.180)	-95.00 (10.271)	132.85 (68.495)	-2.82 (30.016)
		95% CI		-97.31, -92.69		-12.30, 6.66

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (2.50 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_1na.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.16 Descriptive Statistics of Urinary 1-NA - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Quantity Excreted over 24 hours (ng)	Day 5	n	79	79	41	41
		BLOQ - n (%)	59 (74.7%)		0	
		Geometric Mean (CV%)	4.74 (59.75%)		134.21 (53.10%)	
		Geometric 95% CI	4.19, 5.37		114.67, 157.08	
		Median	4.22	-96.13	146.47	8.37
		Q25, Q75	3.31, 6.41	-97.39, -94.97	91.50, 203.56	-4.53, 24.91
		Min, Max	1.9, 94.3	-99.0, -70.0	42.0, 384.3	-44.7, 76.2
		Mean (SD)	6.22 (10.489)	-94.94 (4.773)	150.73 (72.679)	8.25 (27.425)
		95% CI		-96.02, -93.87		-0.41, 16.91

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (2.50 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_1na.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.16 Descriptive Statistics of Urinary 1-NA - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Baseline	n	80		41	
		BLOQ - n (%)	0		0	
		Geometric Mean (CV%)	73.83 (49.82%)		77.84 (54.16%)	
		Geometric 95% CI	66.48, 81.99		66.32, 91.36	
		Median	73.86		76.03	
		Q25, Q75	60.04, 102.67		57.69, 98.15	
		Min, Max	14.4, 171.7		30.0, 316.7	
		Mean (SD)	81.45 (34.735)		89.55 (57.093)	
		95% CI				

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (2.50 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_1na.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.16 Descriptive Statistics of Urinary 1-NA - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Day 1	n	80	80	41	41
		BLOQ - n (%)	1 (1.3%)		0	
		Geometric Mean (CV%)	15.49 (60.57%)		78.41 (53.27%)	
		Geometric 95% CI	13.67, 17.54		66.97, 91.82	
		Median	15.93	-79.86	80.43	1.93
		Q25, Q75	11.21, 21.30	-83.60, -76.06	59.08, 95.30	-12.75, 18.39
		Min, Max	2.5, 98.4	-89.1, 50.7	33.0, 405.6	-54.1, 54.7
		Mean (SD)	18.41 (14.690)	-76.79 (17.438)	90.20 (61.927)	4.10 (25.557)
		95% CI			-80.68, -72.91	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (2.50 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_1na.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.16 Descriptive Statistics of Urinary 1-NA - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Day 2	n	80	80	41	41
		BLOQ - n (%)	7 (8.8%)		0	
		Geometric Mean (CV%)	7.24 (55.48%)		71.82 (69.15%)	
		Geometric 95% CI	6.45, 8.13		58.96, 87.50	
		Median	6.92	-90.32	75.53	-2.80
		Q25, Q75	5.55, 9.51	-92.41, -88.14	51.78, 104.81	-13.41, 19.42
		Min, Max	1.5, 55.4	-97.6, -17.8	6.6, 371.3	-88.5, 61.8
		Mean (SD)	8.39 (6.332)	-88.93 (8.819)	85.38 (57.668)	-1.03 (30.172)
		95% CI			-90.90, -86.97	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (2.50 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_1na.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.16 Descriptive Statistics of Urinary 1-NA - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Day 3	n	79	79	41	41
		BLOQ - n (%)	22 (27.8%)		1 (2.4%)	
		Geometric Mean (CV%)	4.38 (80.41%)		72.24 (77.65%)	
		Geometric 95% CI	3.74, 5.14		58.16, 89.74	
		Median	4.21	-94.08	75.74	2.13
		Q25, Q75	2.71, 5.94	-95.67, -92.43	50.55, 99.18	-13.09, 17.38
		Min, Max	1.4, 105.8	-98.3, 3.0	3.5, 345.6	-96.4, 76.3
		Mean (SD)	6.93 (15.052)	-92.06 (12.664)	87.03 (57.614)	1.66 (30.894)
		95% CI		-94.90, -89.22		-8.10, 11.41

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (2.50 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_1na.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.16 Descriptive Statistics of Urinary 1-NA - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Day 4	n	79	79	41	41
		BLOQ - n (%)	58 (73.4%)		0	
		Geometric Mean (CV%)	2.84 (65.58%)		77.29 (53.33%)	
		Geometric 95% CI	2.48, 3.26		66.00, 90.52	
		Median	2.67	-96.69	76.07	3.11
		Q25, Q75	2.01, 3.81	-97.45, -94.62	60.08, 106.46	-13.98, 22.70
		Min, Max	1.1, 119.3	-98.7, 16.1	29.4, 293.1	-82.7, 70.6
		Mean (SD)	4.44 (13.152)	-94.34 (12.887)	87.67 (49.229)	5.54 (32.183)
		95% CI		-97.23, -91.45		-4.62, 15.70

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (2.50 pg/mL).

Appendix 15.3.3.2

Path: /cvm/projects/prj/development/000000106324/dev/tables/t_1na.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.16 Descriptive Statistics of Urinary 1-NA - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Day 5	n	79	79	41	41
		BLOQ - n (%)	59 (74.7%)		0	
		Geometric Mean (CV%)	3.30 (64.68%)		89.37 (46.05%)	
		Geometric 95% CI	2.89, 3.78		77.81, 102.64	
		Median	3.03	-95.95	88.06	20.01
		Q25, Q75	2.13, 4.74	-97.01, -93.81	72.24, 106.55	0.09, 35.27
		Min, Max	1.0, 29.2	-99.0, -61.3	31.9, 287.1	-50.2, 106.9
		Mean (SD)	4.12 (4.154)	-94.16 (5.939)	98.61 (48.710)	19.17 (31.951)
		95% CI		-95.49, -92.82		9.08, 29.26

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (2.50 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_1na.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.16 Descriptive Statistics of Urinary 1-NA - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Quantity Excreted over 24 hours (ng)	Baseline	n	39	
		BLOQ - n (%)	0	
		Geometric Mean (CV%)	122.17 (38.79%)	
		Geometric 95% CI	108.20, 137.94	
		Median	118.91	
		Q25, Q75	91.01, 158.33	
		Min, Max	55.2, 271.2	
		Mean (SD)	130.86 (50.535)	
		95% CI		

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (2.50 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_1na.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.16 Descriptive Statistics of Urinary 1-NA - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Quantity Excreted over 24 hours (ng)	Day 1	n	39	39
		BLOQ - n (%)	0	
		Geometric Mean (CV%)	22.11 (38.00%)	
		Geometric 95% CI	19.62, 24.91	
		Median	22.41	-81.70
		Q25, Q75	17.28, 28.39	-85.01, -78.49
		Min, Max	9.3, 47.7	-89.1, -68.2
		Mean (SD)	23.59 (8.764)	-81.37 (4.638)
		95% CI		-82.88, -79.86

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (2.50 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_1na.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.16 Descriptive Statistics of Urinary 1-NA - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Quantity Excreted over 24 hours (ng)	Day 2	n	39	39
		BLOQ - n (%)	5 (12.8%)	
		Geometric Mean (CV%)	10.00 (53.05%)	
		Geometric 95% CI	8.50, 11.76	
		Median	10.78	-91.13
		Q25, Q75	7.75, 13.54	-93.70, -88.39
		Min, Max	2.4, 30.0	-97.8, -84.1
		Mean (SD)	11.16 (5.224)	-91.04 (3.528)
		95% CI		-92.18, -89.89

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (2.50 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_1na.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.16 Descriptive Statistics of Urinary 1-NA - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Quantity Excreted over 24 hours (ng)	Day 3	n	39	39
		BLOQ - n (%)	14 (35.9%)	
		Geometric Mean (CV%)	6.03 (46.74%)	
		Geometric 95% CI	5.22, 6.98	
		Median	6.59	-94.50
		Q25, Q75	4.18, 8.47	-96.77, -93.39
		Min, Max	2.7, 14.7	-98.0, -89.2
		Mean (SD)	6.61 (2.798)	-94.56 (2.367)
		95% CI		-95.33, -93.78

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (2.50 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_1na.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.16 Descriptive Statistics of Urinary 1-NA - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Quantity Excreted over 24 hours (ng)	Day 4	n	39	39
		BLOQ - n (%)	31 (79.5%)	
		Geometric Mean (CV%)	3.99 (37.15%)	
		Geometric 95% CI	3.55, 4.49	
		Median	3.47	-96.71
		Q25, Q75	3.12, 5.50	-97.88, -95.37
		Min, Max	2.3, 9.3	-98.6, -92.2
		Mean (SD)	4.26 (1.689)	-96.38 (1.732)
		95% CI		-96.94, -95.81

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (2.50 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_1na.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.16 Descriptive Statistics of Urinary 1-NA - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Quantity Excreted over 24 hours (ng)	Day 5	n	39	39
		BLOQ - n (%)	30 (76.9%)	
		Geometric Mean (CV%)	4.01 (39.87%)	
		Geometric 95% CI	3.54, 4.55	
		Median	3.75	-96.59
		Q25, Q75	3.06, 5.22	-97.73, -95.19
		Min, Max	1.9, 9.3	-98.7, -93.3
		Mean (SD)	4.33 (1.818)	-96.40 (1.503)
		95% CI		-96.89, -95.91

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (2.50 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_1na.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.16 Descriptive Statistics of Urinary 1-NA - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Baseline	n	39	
		BLOQ - n (%)	0	
		Geometric Mean (CV%)	77.31 (42.41%)	
		Geometric 95% CI	67.76, 88.21	
		Median	86.20	
		Q25, Q75	57.53, 98.78	
		Min, Max	32.4, 222.4	
		Mean (SD)	83.95 (36.968)	
		95% CI		

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (2.50 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_1na.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.16 Descriptive Statistics of Urinary 1-NA - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Day 1	n	39	39
		BLOQ - n (%)	0	
		Geometric Mean (CV%)	14.88 (46.01%)	
		Geometric 95% CI	12.91, 17.16	
		Median	14.69	-79.68
		Q25, Q75	10.92, 18.45	-83.93, -76.56
		Min, Max	6.9, 49.8	-89.7, -70.9
		Mean (SD)	16.43 (8.256)	-80.11 (4.952)
		95% CI		-81.72, -78.50

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (2.50 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_1na.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.16 Descriptive Statistics of Urinary 1-NA - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Day 2	n	39	39
		BLOQ - n (%)	5 (12.8%)	
		Geometric Mean (CV%)	6.50 (52.72%)	
		Geometric 95% CI	5.53, 7.64	
		Median	6.96	-90.86
		Q25, Q75	4.50, 9.52	-93.88, -88.86
		Min, Max	2.0, 16.7	-97.1, -83.1
		Mean (SD)	7.27 (3.433)	-90.96 (3.393)
		95% CI		-92.06, -89.86

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (2.50 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_1na.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.16 Descriptive Statistics of Urinary 1-NA - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Day 3	n	39	39
		BLOQ - n (%)	14 (35.9%)	
		Geometric Mean (CV%)	3.93 (58.12%)	
		Geometric 95% CI	3.30, 4.69	
		Median	3.94	-94.22
		Q25, Q75	2.90, 5.75	-96.36, -92.76
		Min, Max	1.3, 11.3	-98.2, -88.5
		Mean (SD)	4.51 (2.437)	-94.39 (2.413)
		95% CI		-95.18, -93.60

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (2.50 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_1na.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.16 Descriptive Statistics of Urinary 1-NA - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Day 4	n	39	39
		BLOQ - n (%)	31 (79.5%)	
		Geometric Mean (CV%)	2.71 (48.64%)	
		Geometric 95% CI	2.33, 3.15	
		Median	2.70	-96.68
		Q25, Q75	1.79, 3.48	-97.31, -95.47
		Min, Max	1.3, 7.9	-98.6, -87.9
		Mean (SD)	3.03 (1.604)	-96.09 (2.096)
		95% CI		-96.77, -95.40

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (2.50 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_1na.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.16 Descriptive Statistics of Urinary 1-NA - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Day 5	n	39	39
		BLOQ - n (%)	30 (76.9%)	
		Geometric Mean (CV%)	2.56 (39.76%)	
		Geometric 95% CI	2.25, 2.90	
		Median	2.53	-97.03
		Q25, Q75	1.92, 3.23	-97.41, -95.31
		Min, Max	1.1, 5.8	-98.4, -93.5
		Mean (SD)	2.75 (1.128)	-96.41 (1.452)
		95% CI		-96.89, -95.93

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (2.50 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_1na.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.17 Analysis of Urinary 2-NA on Day 5 - FAS**

Variable	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	THS : CC Ratio (%)	THS : SA Ratio (%)
Urinary Quantity over n 24 hours (ng)		79	41	39		
	Geometric LS Mean (SE) ¹	4.24	37.84	3.98	11.22 (0.071)	106.71 (0.072)
	95% CI	3.91, 4.61	33.78, 42.39	3.54, 4.47	9.75, 12.91	92.57, 123.02
Concentration Adjusted for Creatinine (pg/mg creat)	n	79	41	39		
	Geometric LS Mean (SE) ¹	2.95	25.55	2.53	11.54 (0.069)	116.65 (0.070)
	95% CI	2.72, 3.19	22.90, 28.50	2.25, 2.83	10.08, 13.21	101.73, 133.76

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Adjusted geometric least squares (LS) means and confidence intervals (CIs) from an ANCOVA model conducted on log-transformed Day 5 values with log-transformed baseline value, study arm, sex and CC consumption reported at screening as fixed effect factors.

Baseline is defined as the last assessment prior to Day 1 product use for THS 2.2 and CC subjects and prior to 06:29 AM on Day 1 for SA subjects.

Note: 1 = Standard errors presented from analysis performed on the log_e scale.

Note: Collection over 24 hours starting on the Day stated in the title.

Appendix 15.4.4.17 and 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/tl_anlsecondbio.sas

Program Run: 27OCT14 cvn_ahedge Program Status: FINAL

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**Table 15.2.4.18 Descriptive Statistics of Urinary 2-NA - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Quantity Excreted over 24 hours (ng)	Baseline	n	80		41	
		BLOQ - n (%)	0		0	
		Geometric Mean (CV%)	39.35 (48.21%)		39.77 (44.60%)	
		Geometric 95% CI	35.54, 43.57		34.77, 45.50	
		Median	42.29		35.52	
		Q25, Q75	32.85, 52.14		27.37, 54.40	
		Min, Max	7.2, 86.1		21.6, 125.0	
		Mean (SD)	42.86 (15.754)		43.80 (21.804)	
		95% CI				

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (2.50 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_2na.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.18 Descriptive Statistics of Urinary 2-NA - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Quantity Excreted over 24 hours (ng)	Day 1	n	80	80	41	41
		BLOQ - n (%)	9 (11.3%)		0	
		Geometric Mean (CV%)	9.23 (53.26%)		36.02 (51.71%)	
		Geometric 95% CI	8.25, 10.32		30.88, 42.00	
		Median	10.23	-77.34	36.45	-7.17
		Q25, Q75	7.33, 12.76	-80.44, -71.48	24.50, 55.94	-18.42, 2.88
		Min, Max	2.0, 35.4	-91.1, -32.4	12.5, 115.7	-48.8, 53.4
		Mean (SD)	10.32 (4.938)	-75.29 (8.749)	40.46 (20.632)	-7.41 (19.701)
		95% CI		-77.24, -73.34		-13.63, -1.18

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (2.50 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_2na.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.18 Descriptive Statistics of Urinary 2-NA - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Quantity Excreted over 24 hours (ng)	Day 2	n	80	80	41	41
		BLOQ - n (%)	51 (63.8%)		1 (2.4%)	
		Geometric Mean (CV%)	4.47 (56.77%)		34.09 (69.52%)	
		Geometric 95% CI	3.97, 5.04		27.96, 41.56	
		Median	3.67	-89.68	31.72	-11.25
		Q25, Q75	3.15, 6.50	-92.42, -84.62	25.08, 52.92	-24.75, 6.30
		Min, Max	1.8, 67.2	-95.7, 45.1	2.5, 108.9	-95.2, 88.5
		Mean (SD)	5.54 (7.317)	-85.94 (16.293)	39.80 (21.222)	-6.36 (31.675)
		95% CI		-89.57, -82.31		-16.37, 3.64

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (2.50 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_2na.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.18 Descriptive Statistics of Urinary 2-NA - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Quantity Excreted over 24 hours (ng)	Day 3	n	79	79	41	41
		BLOQ - n (%)	51 (64.6%)		1 (2.4%)	
		Geometric Mean (CV%)	4.07 (52.48%)		34.23 (65.12%)	
		Geometric 95% CI	3.64, 4.55		28.37, 41.30	
		Median	3.47	-90.50	33.88	-6.59
		Q25, Q75	2.81, 5.76	-92.57, -86.46	24.21, 49.81	-25.66, 10.00
		Min, Max	1.8, 36.6	-95.3, -21.5	3.3, 110.0	-90.7, 31.1
		Mean (SD)	4.77 (4.211)	-88.16 (9.192)	39.49 (20.343)	-8.66 (24.577)
		95% CI		-90.23, -86.10		-16.42, -0.90

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (2.50 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_2na.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.18 Descriptive Statistics of Urinary 2-NA - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Quantity Excreted over 24 hours (ng)	Day 4	n	79	79	41	41
		BLOQ - n (%)	71 (89.9%)		0	
		Geometric Mean (CV%)	3.40 (34.35%)		34.74 (51.54%)	
		Geometric 95% CI	3.15, 3.67		29.80, 40.49	
		Median	3.31	-92.01	35.56	-11.07
		Q25, Q75	2.75, 3.75	-93.93, -89.15	24.15, 49.50	-24.79, -0.46
		Min, Max	1.9, 7.5	-96.4, -54.1	11.4, 111.0	-73.7, 77.9
		Mean (SD)	3.60 (1.335)	-90.17 (6.858)	38.99 (19.991)	-9.03 (25.159)
		95% CI		-91.71, -88.63		-16.98, -1.09

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (2.50 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_2na.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.18 Descriptive Statistics of Urinary 2-NA - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Quantity Excreted over 24 hours (ng)	Day 5	n	79	79	41	41
		BLOQ - n (%)	65 (82.3%)		0	
		Geometric Mean (CV%)	4.25 (38.39%)		38.03 (55.43%)	
		Geometric 95% CI	3.91, 4.63		32.29, 44.78	
		Median	3.75	-89.59	40.35	-0.12
		Q25, Q75	3.12, 5.75	-92.19, -85.40	26.32, 59.11	-15.94, 13.61
		Min, Max	1.9, 9.0	-97.4, -48.0	12.0, 127.4	-51.1, 66.3
		Mean (SD)	4.56 (1.773)	-87.53 (8.260)	43.20 (22.617)	-1.55 (23.515)
		95% CI		-89.38, -85.67		-8.98, 5.87

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (2.50 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_2na.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.18 Descriptive Statistics of Urinary 2-NA - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Baseline	n	80		41	
		BLOQ - n (%)	0		0	
		Geometric Mean (CV%)	24.54 (49.21%)		24.14 (43.25%)	
		Geometric 95% CI	22.12, 27.22		21.18, 27.51	
		Median	24.02		23.45	
		Q25, Q75	19.98, 35.08		18.85, 31.10	
		Min, Max	5.7, 52.1		10.7, 55.1	
		Mean (SD)	27.05 (11.467)		26.28 (11.423)	
		95% CI				

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (2.50 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_2na.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.18 Descriptive Statistics of Urinary 2-NA - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Day 1	n	80	80	41	41
		BLOQ - n (%)	9 (11.3%)		0	
		Geometric Mean (CV%)	6.41 (52.63%)		24.70 (44.54%)	
		Geometric 95% CI	5.74, 7.17		21.59, 28.26	
		Median	6.59	-74.45	25.57	5.14
		Q25, Q75	4.80, 8.76	-77.55, -69.47	18.63, 32.32	-9.71, 16.34
		Min, Max	1.7, 18.0	-87.6, -42.9	11.2, 67.5	-50.6, 45.1
		Mean (SD)	7.17 (3.354)	-72.86 (7.802)	27.00 (12.080)	4.33 (19.666)
		95% CI		-74.60, -71.12		-1.88, 10.54

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (2.50 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_2na.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.18 Descriptive Statistics of Urinary 2-NA - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Day 2	n	80	80	41	41
		BLOQ - n (%)	51 (63.8%)		1 (2.4%)	
		Geometric Mean (CV%)	2.99 (52.68%)		21.37 (64.67%)	
		Geometric 95% CI	2.68, 3.35		17.73, 25.76	
		Median	2.67	-88.72	23.96	-5.52
		Q25, Q75	2.06, 4.16	-91.33, -83.78	15.39, 31.20	-16.80, 9.35
		Min, Max	1.4, 19.8	-94.9, -4.6	1.6, 52.2	-91.4, 47.8
		Mean (SD)	3.45 (2.450)	-85.72 (11.429)	24.31 (11.014)	-5.66 (25.310)
		95% CI		-88.27, -83.17		-13.66, 2.33

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (2.50 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_2na.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.18 Descriptive Statistics of Urinary 2-NA - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Day 3	n	79	79	41	41
		BLOQ - n (%)	51 (64.6%)		1 (2.4%)	
		Geometric Mean (CV%)	2.75 (51.06%)		22.10 (55.98%)	
		Geometric 95% CI	2.46, 3.07		18.74, 26.07	
		Median	2.47	-89.15	22.43	1.74
		Q25, Q75	2.00, 3.68	-92.13, -85.67	15.04, 29.36	-17.19, 7.84
		Min, Max	1.1, 28.8	-95.1, -30.7	3.5, 63.7	-88.6, 61.8
		Mean (SD)	3.23 (3.176)	-87.36 (8.559)	24.92 (12.219)	-2.91 (26.698)
		95% CI		-89.28, -85.44		-11.35, 5.52

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (2.50 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_2na.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.4.18 Descriptive Statistics of Urinary 2-NA - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Day 4	n	79	79	41	41
		BLOQ - n (%)	71 (89.9%)		0	
		Geometric Mean (CV%)	2.33 (33.86%)		22.78 (49.35%)	
		Geometric 95% CI	2.16, 2.51		19.65, 26.40	
		Median	2.24	-91.15	24.07	-4.87
		Q25, Q75	1.92, 2.98	-93.12, -87.34	15.43, 31.99	-14.72, 10.35
		Min, Max	1.0, 5.6	-95.9, -52.0	9.4, 52.5	-82.9, 64.7
		Mean (SD)	2.45 (0.820)	-89.18 (7.384)	25.21 (11.202)	-1.47 (24.708)
		95% CI		-90.84, -87.52		-9.28, 6.33

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (2.50 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_2na.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.18 Descriptive Statistics of Urinary 2-NA - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Day 5	n	79	79	41	41
		BLOQ - n (%)	65 (82.3%)		0	
		Geometric Mean (CV%)	2.96 (47.36%)		25.32 (42.35%)	
		Geometric 95% CI	2.67, 3.28		22.27, 28.79	
		Median	2.67	-88.85	24.53	3.66
		Q25, Q75	2.10, 4.29	-92.23, -83.69	18.82, 33.53	-7.68, 25.11
		Min, Max	1.4, 9.6	-95.7, -38.4	11.5, 58.4	-49.0, 49.7
		Mean (SD)	3.29 (1.634)	-85.39 (11.418)	27.47 (11.630)	7.19 (21.467)
		95% CI			-87.95, -82.82	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (2.50 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_2na.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.18 Descriptive Statistics of Urinary 2-NA - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Quantity Excreted over 24 hours (ng)	Baseline	n	39	
		BLOQ - n (%)	0	
		Geometric Mean (CV%)	38.87 (41.64%)	
		Geometric 95% CI	34.14, 44.25	
		Median	41.40	
		Q25, Q75	28.12, 50.70	
		Min, Max	16.2, 87.5	
		Mean (SD)	41.95 (16.731)	
		95% CI		

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (2.50 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_2na.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.18 Descriptive Statistics of Urinary 2-NA - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Quantity Excreted over 24 hours (ng)	Day 1	n	39	39
		BLOQ - n (%)	2 (5.1%)	
		Geometric Mean (CV%)	9.97 (44.73%)	
		Geometric 95% CI	8.68, 11.46	
		Median	10.41	-74.60
		Q25, Q75	8.35, 12.84	-77.97, -70.83
		Min, Max	2.6, 20.8	-85.3, -55.6
		Mean (SD)	10.79 (4.033)	-73.57 (6.630)
		95% CI		-75.72, -71.42

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (2.50 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_2na.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.18 Descriptive Statistics of Urinary 2-NA - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Quantity Excreted over 24 hours (ng)	Day 2	n	39	39
		BLOQ - n (%)	22 (56.4%)	
		Geometric Mean (CV%)	4.84 (49.27%)	
		Geometric 95% CI	4.15, 5.63	
		Median	4.63	-88.22
		Q25, Q75	3.31, 7.54	-90.95, -81.12
		Min, Max	2.3, 12.1	-94.4, -72.8
		Mean (SD)	5.39 (2.642)	-86.33 (5.946)
		95% CI		-88.26, -84.40

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (2.50 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_2na.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.18 Descriptive Statistics of Urinary 2-NA - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Quantity Excreted over 24 hours (ng)	Day 3	n	39	39
		BLOQ - n (%)	23 (59.0%)	
		Geometric Mean (CV%)	4.43 (38.10%)	
		Geometric 95% CI	3.93, 5.00	
		Median	4.31	-87.75
		Q25, Q75	3.18, 6.16	-91.85, -84.44
		Min, Max	2.2, 9.5	-95.1, -75.9
		Mean (SD)	4.73 (1.770)	-87.54 (5.127)
		95% CI		-89.21, -85.88

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (2.50 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_2na.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.18 Descriptive Statistics of Urinary 2-NA - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Quantity Excreted over 24 hours (ng)	Day 4	n	39	39
		BLOQ - n (%)	34 (87.2%)	
		Geometric Mean (CV%)	3.67 (30.85%)	
		Geometric 95% CI	3.32, 4.05	
		Median	3.38	-91.30
		Q25, Q75	3.06, 4.88	-93.51, -87.14
		Min, Max	2.3, 7.2	-95.3, -73.7
		Mean (SD)	3.84 (1.249)	-89.51 (5.458)
		95% CI		-91.29, -87.74

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (2.50 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_2na.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.18 Descriptive Statistics of Urinary 2-NA - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Quantity Excreted over 24 hours (ng)	Day 5	n	39	39
		BLOQ - n (%)	31 (79.5%)	
		Geometric Mean (CV%)	3.95 (36.57%)	
		Geometric 95% CI	3.52, 4.43	
		Median	3.75	-89.26
		Q25, Q75	3.06, 5.00	-92.91, -85.68
		Min, Max	2.3, 8.3	-96.1, -78.3
		Mean (SD)	4.21 (1.632)	-88.87 (4.733)
		95% CI		-90.41, -87.33

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (2.50 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_2na.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.18 Descriptive Statistics of Urinary 2-NA - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Baseline	n	39	
		BLOQ - n (%)	0	
		Geometric Mean (CV%)	24.59 (42.86%)	
		Geometric 95% CI	21.52, 28.10	
		Median	25.04	
		Q25, Q75	19.25, 32.42	
		Min, Max	9.5, 67.8	
		Mean (SD)	26.69 (11.360)	
		95% CI		

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (2.50 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_2na.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.18 Descriptive Statistics of Urinary 2-NA - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Day 1	n	39	39
		BLOQ - n (%)	2 (5.1%)	
		Geometric Mean (CV%)	6.71 (43.38%)	
		Geometric 95% CI	5.86, 7.69	
		Median	6.70	-72.41
		Q25, Q75	5.34, 9.13	-76.35, -66.50
		Min, Max	2.5, 18.0	-86.8, -59.2
		Mean (SD)	7.30 (3.124)	-71.90 (6.600)
		95% CI		-74.05, -69.76

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (2.50 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_2na.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.18 Descriptive Statistics of Urinary 2-NA - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Day 2	n	39	39
		BLOQ - n (%)	22 (56.4%)	
		Geometric Mean (CV%)	3.14 (48.89%)	
		Geometric 95% CI	2.70, 3.66	
		Median	2.97	-87.33
		Q25, Q75	2.15, 4.25	-90.71, -81.64
		Min, Max	1.3, 8.6	-95.5, -72.4
		Mean (SD)	3.50 (1.756)	-85.98 (5.936)
		95% CI		-87.91, -84.05

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (2.50 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_2na.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.18 Descriptive Statistics of Urinary 2-NA - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Day 3	n	39	39
		BLOQ - n (%)	23 (59.0%)	
		Geometric Mean (CV%)	2.89 (43.45%)	
		Geometric 95% CI	2.52, 3.31	
		Median	2.91	-87.86
		Q25, Q75	2.26, 3.51	-91.68, -83.85
		Min, Max	1.3, 8.6	-94.9, -74.2
		Mean (SD)	3.16 (1.488)	-87.27 (5.097)
		95% CI		-88.93, -85.62

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (2.50 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_2na.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.18 Descriptive Statistics of Urinary 2-NA - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Day 4	n	39	39
		BLOQ - n (%)	34 (87.2%)	
		Geometric Mean (CV%)	2.49 (40.36%)	
		Geometric 95% CI	2.19, 2.83	
		Median	2.47	-90.42
		Q25, Q75	1.70, 3.34	-92.09, -87.71
		Min, Max	1.3, 5.7	-95.7, -58.6
		Mean (SD)	2.69 (1.114)	-88.54 (6.962)
		95% CI		-90.80, -86.28

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (2.50 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_2na.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.18 Descriptive Statistics of Urinary 2-NA - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Day 5	n	39	39
		BLOQ - n (%)	31 (79.5%)	
		Geometric Mean (CV%)	2.52 (38.01%)	
		Geometric 95% CI	2.23, 2.84	
		Median	2.41	-90.15
		Q25, Q75	1.79, 3.39	-92.18, -86.10
		Min, Max	1.3, 5.9	-95.0, -78.0
		Mean (SD)	2.70 (1.069)	-88.93 (4.467)
		95% CI		-90.38, -87.47

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (2.50 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_2na.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.19 Analysis of Urinary o-toluidine on Day 5 - FAS**

Variable	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	THS : CC Ratio (%)	THS : SA Ratio (%)
Urinary Quantity over n 24 hours (ng)		79	41	39		
	Geometric LS Mean (SE) ¹	73.19	181.28	65.87	40.38 (0.074)	111.12 (0.075)
	95% CI	67.24, 79.67	161.15, 203.92	58.37, 74.33	34.92, 46.68	95.86, 128.80
Concentration Adjusted for Creatinine (pg/mg creat)	n	79	41	39		
	Geometric LS Mean (SE) ¹	50.89	122.03	41.95	41.70 (0.075)	121.31 (0.076)
	95% CI	46.70, 55.44	108.34, 137.44	37.13, 47.39	36.01, 48.29	104.50, 140.81

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Adjusted geometric least squares (LS) means and confidence intervals (CIs) from an ANCOVA model conducted on log-transformed Day 5 values with log-transformed baseline value, study arm, sex and CC consumption reported at screening as fixed effect factors.

Baseline is defined as the last assessment prior to Day 1 product use for THS 2.2 and CC subjects and prior to 06:29 AM on Day 1 for SA subjects.

Note: 1 = Standard errors presented from analysis performed on the log_e scale.

Note: Collection over 24 hours starting on the Day stated in the title.

Appendix 15.4.4.19 and 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/tl_anlsecondbio.sas

Program Run: 27OCT14 cvn_ahedge Program Status: FINAL

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**Table 15.2.4.20 Descriptive Statistics of Urinary o-toluidine - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Quantity Excreted over 24 hours (ng)	Baseline	n	80		41	
		BLOQ - n (%)	5 (6.3%)		2 (4.9%)	
		Geometric Mean (CV%)	216.78 (46.49%)		216.39 (42.99%)	
		Geometric 95% CI	196.46, 239.21		190.01, 246.43	
		Median	230.64		213.50	
		Q25, Q75	181.26, 285.02		174.02, 285.60	
		Min, Max	40.0, 450.6		71.9, 676.1	
		Mean (SD)	234.50 (81.518)		235.57 (108.007)	
		95% CI				

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (50.0 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_otol.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.20 Descriptive Statistics of Urinary o-toluidine - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Quantity Excreted over 24 hours (ng)	Day 1	n	80	80	41	41
		BLOQ - n (%)	60 (75.0%)		5 (12.2%)	
		Geometric Mean (CV%)	71.15 (44.64%)		188.50 (56.00%)	
		Geometric 95% CI	64.71, 78.23		159.84, 222.28	
		Median	61.56	-70.03	186.85	-8.51
		Q25, Q75	55.00, 95.97	-73.64, -61.15	151.05, 265.65	-21.48, 2.61
		Min, Max	35.0, 294.5	-87.1, 10.9	45.0, 490.1	-64.4, 29.1
		Mean (SD)	79.00 (44.316)	-63.24 (20.313)	212.79 (102.424)	-10.21 (20.190)
		95% CI		-67.77, -58.72		-16.59, -3.83

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (50.0 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_otoI.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.20 Descriptive Statistics of Urinary o-toluidine - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Quantity Excreted over 24 hours (ng)	Day 2	n	80	80	41	41
		BLOQ - n (%)	77 (96.3%)		5 (12.2%)	
		Geometric Mean (CV%)	64.29 (32.00%)		180.92 (53.82%)	
		Geometric 95% CI	59.97, 68.92		154.29, 212.14	
		Median	63.75	-72.79	168.84	-13.94
		Q25, Q75	55.62, 72.19	-76.58, -66.30	143.06, 266.20	-25.58, 3.34
		Min, Max	35.0, 324.8	-87.5, 46.9	50.0, 444.6	-82.3, 62.2
		Mean (SD)	68.41 (34.627)	-65.42 (25.509)	202.84 (94.796)	-11.18 (28.598)
		95% CI		-71.10, -59.74		-20.21, -2.15

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (50.0 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_otoI.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.20 Descriptive Statistics of Urinary o-toluidine - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Quantity Excreted over 24 hours (ng)	Day 3	n	79	79	41	41
		BLOQ - n (%)	71 (89.9%)		4 (9.8%)	
		Geometric Mean (CV%)	62.32 (32.88%)		193.89 (50.67%)	
		Geometric 95% CI	58.00, 66.96		166.74, 225.47	
		Median	58.75	-73.08	204.00	-5.84
		Q25, Q75	52.50, 69.23	-79.17, -65.56	147.69, 259.05	-22.16, 7.35
		Min, Max	35.6, 236.3	-87.5, -3.4	56.3, 491.4	-67.8, 62.4
		Mean (SD)	66.38 (31.155)	-67.68 (18.909)	214.62 (93.490)	-6.19 (26.123)
		95% CI		-71.92, -63.44		-14.44, 2.06

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (50.0 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_otol.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.20 Descriptive Statistics of Urinary o-toluidine - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Quantity Excreted over 24 hours (ng)	Day 4	n	79	79	41	41
		BLOQ - n (%)	77 (97.5%)		9 (22.0%)	
		Geometric Mean (CV%)	64.30 (30.71%)		182.28 (52.74%)	
		Geometric 95% CI	60.11, 68.78		155.89, 213.13	
		Median	63.75	-73.55	186.76	-12.52
		Q25, Q75	55.00, 73.13	-77.57, -63.12	127.50, 251.99	-25.66, 5.93
		Min, Max	37.5, 171.8	-87.8, 65.6	60.0, 491.4	-79.3, 73.5
		Mean (SD)	67.51 (23.859)	-65.68 (25.331)	204.19 (97.047)	-10.43 (29.398)
		95% CI		-71.36, -60.00		-19.72, -1.15

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (50.0 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_otol.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.20 Descriptive Statistics of Urinary o-toluidine - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Quantity Excreted over 24 hours (ng)	Day 5	n	79	79	41	41
		BLOQ - n (%)	78 (98.7%)		13 (31.7%)	
		Geometric Mean (CV%)	73.44 (35.00%)		181.95 (62.69%)	
		Geometric 95% CI	68.05, 79.26		151.72, 218.22	
		Median	70.00	-65.96	190.77	-9.43
		Q25, Q75	57.50, 87.50	-76.94, -55.83	116.25, 295.43	-30.40, 7.66
		Min, Max	38.8, 172.5	-90.0, 103.0	53.8, 543.7	-65.0, 50.3
		Mean (SD)	78.04 (29.522)	-59.01 (34.419)	210.86 (109.578)	-10.74 (28.398)
		95% CI		-66.72, -51.29		-19.71, -1.77

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (50.0 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_otoI.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.20 Descriptive Statistics of Urinary o-toluidine - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Baseline	n	80		41	
		BLOQ - n (%)	5 (6.3%)		2 (4.9%)	
		Geometric Mean (CV%)	135.20 (45.49%)		131.32 (41.70%)	
		Geometric 95% CI	122.75, 148.90		115.72, 149.01	
		Median	135.82		126.02	
		Q25, Q75	111.73, 183.35		98.22, 170.36	
		Min, Max	35.5, 291.8		48.1, 382.0	
		Mean (SD)	147.15 (58.453)		142.33 (62.298)	
		95% CI				

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (50.0 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_otol.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.20 Descriptive Statistics of Urinary o-toluidine - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Day 1	n	80	80	41	41
		BLOQ - n (%)	60 (75.0%)		5 (12.2%)	
		Geometric Mean (CV%)	49.46 (39.25%)		129.28 (45.31%)	
		Geometric 95% CI	45.46, 53.82		112.79, 148.17	
		Median	48.50	-65.34	133.12	1.88
		Q25, Q75	37.99, 64.53	-72.87, -52.08	97.08, 172.62	-9.40, 17.36
		Min, Max	24.0, 175.1	-87.2, 37.6	45.5, 333.3	-49.9, 32.4
		Mean (SD)	53.36 (23.527)	-59.09 (22.805)	141.04 (59.204)	1.03 (21.336)
		95% CI			-64.17, -54.01	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (50.0 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_otol.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.20 Descriptive Statistics of Urinary o-toluidine - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Day 2	n	80	80	41	41
		BLOQ - n (%)	77 (96.3%)		5 (12.2%)	
		Geometric Mean (CV%)	42.99 (34.08%)		113.44 (47.84%)	
		Geometric 95% CI	39.93, 46.28		98.29, 130.92	
		Median	41.70	-69.64	110.14	-8.27
		Q25, Q75	34.74, 53.09	-74.96, -63.25	85.79, 162.17	-21.32, 3.28
		Min, Max	15.1, 97.7	-85.5, 55.1	32.3, 252.6	-68.4, 39.0
		Mean (SD)	45.44 (16.090)	-63.27 (26.173)	124.54 (52.188)	-9.88 (23.813)
		95% CI			-69.10, -57.44	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (50.0 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_otol.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.20 Descriptive Statistics of Urinary o-toluidine - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Day 3	n	79	79	41	41
		BLOQ - n (%)	71 (89.9%)		4 (9.8%)	
		Geometric Mean (CV%)	42.15 (36.52%)		125.21 (47.93%)	
		Geometric 95% CI	38.93, 45.63		108.47, 144.54	
		Median	40.45	-70.82	133.06	-0.39
		Q25, Q75	33.37, 50.37	-78.09, -62.56	100.73, 161.68	-11.65, 13.59
		Min, Max	20.5, 172.8	-85.5, 28.7	42.8, 347.0	-60.8, 89.8
		Mean (SD)	45.30 (21.760)	-64.69 (22.405)	137.90 (61.808)	-0.19 (28.801)
		95% CI		-69.71, -59.66		-9.29, 8.90

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (50.0 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_otol.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.20 Descriptive Statistics of Urinary o-toluidine - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Day 4	n	79	79	41	41
		BLOQ - n (%)	77 (97.5%)		9 (22.0%)	
		Geometric Mean (CV%)	44.02 (31.23%)		119.54 (50.40%)	
		Geometric 95% CI	41.11, 47.14		102.87, 138.91	
		Median	43.10	-70.56	121.00	-1.79
		Q25, Q75	35.66, 55.07	-76.53, -60.44	88.09, 164.34	-16.96, 20.36
		Min, Max	19.8, 109.1	-88.1, 62.2	37.4, 303.0	-86.5, 57.1
		Mean (SD)	46.12 (14.823)	-62.23 (27.641)	132.59 (59.502)	-2.53 (30.577)
		95% CI			-68.43, -56.03	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (50.0 pg/mL).

Appendix 15.3.3.2

Path: /cvm/projects/prj/development/000000106324/dev/tables/t_otol.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.20 Descriptive Statistics of Urinary o-toluidine - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Day 5	n	79	79	41	41
		BLOQ - n (%)	78 (98.7%)		13 (31.7%)	
		Geometric Mean (CV%)	51.15 (48.96%)		121.16 (47.52%)	
		Geometric 95% CI	46.10, 56.75		105.07, 139.71	
		Median	46.04	-65.55	127.55	1.74
		Q25, Q75	35.66, 71.84	-76.21, -49.48	90.17, 159.67	-22.91, 14.06
		Min, Max	20.3, 192.3	-87.7, 152.4	45.5, 265.9	-56.8, 68.8
		Mean (SD)	57.38 (31.148)	-50.96 (48.676)	133.17 (57.542)	-3.08 (28.779)
		95% CI			-61.87, -40.05	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (50.0 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_otol.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.20 Descriptive Statistics of Urinary o-toluidine - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Quantity Excreted over 24 hours (ng)	Baseline	n	39	
		BLOQ - n (%)	6 (15.4%)	
		Geometric Mean (CV%)	209.36 (42.59%)	
		Geometric 95% CI	183.41, 238.99	
		Median	211.68	
		Q25, Q75	175.72, 285.25	
		Min, Max	70.0, 412.1	
		Mean (SD)	225.34 (82.089)	
		95% CI		

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (50.0 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_otol.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.20 Descriptive Statistics of Urinary o-toluidine - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Quantity Excreted over 24 hours (ng)	Day 1	n	39	39
		BLOQ - n (%)	23 (59.0%)	
		Geometric Mean (CV%)	75.47 (42.37%)	
		Geometric 95% CI	66.16, 86.10	
		Median	72.50	-63.03
		Q25, Q75	58.75, 104.31	-74.50, -57.14
		Min, Max	25.0, 170.0	-82.7, 91.6
		Mean (SD)	81.71 (33.725)	-59.22 (28.045)
		95% CI		-68.32, -50.13

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (50.0 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_otol.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.20 Descriptive Statistics of Urinary o-toluidine - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Quantity Excreted over 24 hours (ng)	Day 2	n	39	39
		BLOQ - n (%)	37 (94.9%)	
		Geometric Mean (CV%)	63.06 (27.12%)	
		Geometric 95% CI	57.84, 68.76	
		Median	63.75	-72.91
		Q25, Q75	50.00, 75.00	-78.08, -63.11
		Min, Max	37.5, 106.8	-86.4, 0.0
		Mean (SD)	65.27 (17.376)	-66.48 (18.717)
		95% CI		-72.55, -60.41

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (50.0 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_otol.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.20 Descriptive Statistics of Urinary o-toluidine - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Quantity Excreted over 24 hours (ng)	Day 3	n	39	39
		BLOQ - n (%)	36 (92.3%)	
		Geometric Mean (CV%)	64.71 (24.10%)	
		Geometric 95% CI	59.91, 69.89	
		Median	63.75	-73.07
		Q25, Q75	55.00, 73.75	-77.66, -64.75
		Min, Max	35.0, 103.1	-81.9, -4.3
		Mean (SD)	66.51 (15.928)	-65.53 (18.878)
		95% CI		-71.65, -59.40

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (50.0 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_otol.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.20 Descriptive Statistics of Urinary o-toluidine - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Quantity Excreted over 24 hours (ng)	Day 4	n	39	39
		BLOQ - n (%)	38 (97.4%)	
		Geometric Mean (CV%)	68.72 (42.95%)	
		Geometric 95% CI	60.14, 78.53	
		Median	63.75	-72.10
		Q25, Q75	52.50, 74.38	-77.28, -60.65
		Min, Max	40.0, 481.6	-86.4, 160.2
		Mean (SD)	78.25 (68.979)	-57.96 (43.916)
		95% CI		-72.20, -43.72

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (50.0 pg/mL).

Appendix 15.3.3.2

Path: /cvm/projects/prj/development/000000106324/dev/tables/t_otol.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.20 Descriptive Statistics of Urinary o-toluidine - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Quantity Excreted over 24 hours (ng)	Day 5	n	39	39
		BLOQ - n (%)	38 (97.4%)	
		Geometric Mean (CV%)	65.32 (26.59%)	
		Geometric 95% CI	60.00, 71.10	
		Median	70.00	-72.64
		Q25, Q75	51.25, 78.13	-78.16, -60.88
		Min, Max	37.5, 105.0	-84.5, 16.9
		Mean (SD)	67.47 (17.147)	-64.61 (21.589)
		95% CI		-71.61, -57.61

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (50.0 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_otol.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.20 Descriptive Statistics of Urinary o-toluidine - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Baseline	n	39	
		BLOQ - n (%)	6 (15.4%)	
		Geometric Mean (CV%)	132.49 (46.61%)	
		Geometric 95% CI	114.75, 152.97	
		Median	143.40	
		Q25, Q75	100.00, 168.98	
		Min, Max	41.1, 362.0	
		Mean (SD)	144.99 (62.156)	
		95% CI		

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (50.0 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_otol.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.20 Descriptive Statistics of Urinary o-toluidine - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Day 1	n	39	39
		BLOQ - n (%)	23 (59.0%)	
		Geometric Mean (CV%)	50.80 (42.90%)	
		Geometric 95% CI	44.46, 58.05	
		Median	50.71	-57.65
		Q25, Q75	34.91, 65.45	-72.20, -50.51
		Min, Max	20.3, 121.2	-82.7, 19.2
		Mean (SD)	55.17 (23.469)	-58.27 (18.409)
		95% CI		-64.24, -52.29

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (50.0 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_otol.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.20 Descriptive Statistics of Urinary o-toluidine - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Day 2	n	39	39
		BLOQ - n (%)	37 (94.9%)	
		Geometric Mean (CV%)	40.99 (33.08%)	
		Geometric 95% CI	36.92, 45.51	
		Median	39.81	-71.02
		Q25, Q75	33.92, 49.31	-76.80, -61.61
		Min, Max	21.7, 83.1	-90.8, -10.2
		Mean (SD)	43.17 (14.736)	-65.57 (17.456)
		95% CI		-71.23, -59.91

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (50.0 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_otoi.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.20 Descriptive Statistics of Urinary o-toluidine - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Day 3	n	39	39
		BLOQ - n (%)	36 (92.3%)	
		Geometric Mean (CV%)	42.18 (35.72%)	
		Geometric 95% CI	37.69, 47.20	
		Median	43.03	-68.10
		Q25, Q75	29.41, 56.06	-76.75, -62.68
		Min, Max	23.1, 89.6	-89.2, -5.3
		Mean (SD)	44.73 (15.818)	-64.59 (18.413)
		95% CI		-70.56, -58.62

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (50.0 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_otol.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.20 Descriptive Statistics of Urinary o-toluidine - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Day 4	n	39	39
		BLOQ - n (%)	38 (97.4%)	
		Geometric Mean (CV%)	46.71 (46.14%)	
		Geometric 95% CI	40.50, 53.86	
		Median	45.70	-67.63
		Q25, Q75	33.64, 57.21	-76.54, -60.02
		Min, Max	24.5, 250.7	-90.7, 190.6
		Mean (SD)	52.65 (36.747)	-53.52 (51.783)
		95% CI		-70.31, -36.73

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (50.0 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_otol.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.20 Descriptive Statistics of Urinary o-toluidine - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Day 5	n	39	39
		BLOQ - n (%)	38 (97.4%)	
		Geometric Mean (CV%)	41.64 (40.03%)	
		Geometric 95% CI	36.74, 47.18	
		Median	41.74	-70.93
		Q25, Q75	34.05, 54.00	-77.44, -57.73
		Min, Max	16.2, 100.8	-85.5, 1.5
		Mean (SD)	44.77 (17.820)	-64.96 (18.329)
		95% CI		-70.90, -59.01

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (50.0 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_otol.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.21 Analysis of Urinary CEMA on Day 5 - FAS**

Variable	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	THS : CC Ratio (%)	THS : SA Ratio (%)
Urinary Quantity over n 24 hours (µg)		79	41	39		
	Geometric LS Mean (SE) ¹	19.19	148.07	19.39	12.96 (0.076)	98.95 (0.078)
	95% CI	17.58, 20.95	131.13, 167.20	17.12, 21.97	11.15, 15.06	84.98, 115.23
Concentration Adjusted for Creatinine (ng/mg creat)	n	79	41	39		
	Geometric LS Mean (SE) ¹	13.29	101.03	12.20	13.16 (0.065)	108.96 (0.066)
	95% CI	12.34, 14.33	91.11, 112.02	10.97, 13.57	11.58, 14.95	95.73, 124.02

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Adjusted geometric least squares (LS) means and confidence intervals (CIs) from an ANCOVA model conducted on log-transformed Day 5 values with log-transformed baseline value, study arm, sex and CC consumption reported at screening as fixed effect factors.

Baseline is defined as the last assessment prior to Day 1 product use for THS 2.2 and CC subjects and prior to 06:29 AM on Day 1 for SA subjects.

Note: 1 = Standard errors presented from analysis performed on the log_e scale.

Note: Collection over 24 hours starting on the Day stated in the title.

Appendix 15.4.4.21 and 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/tl_anlsecondbio.sas

Program Run: 27OCT14 cvn_ahedge Program Status: FINAL

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**Table 15.2.4.22 Descriptive Statistics of Urinary CEMA - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Quantity Excreted over 24 hours (µg)	Baseline	n	80		41	
		BLOQ - n (%)	0		0	
		Geometric Mean (CV%)	157.19 (70.60%)		162.25 (53.22%)	
		Geometric 95% CI	136.44, 181.09		138.58, 189.95	
		Median	178.89		170.64	
		Q25, Q75	129.58, 236.03		116.44, 244.53	
		Min, Max	8.5, 407.5		55.6, 460.2	
		Mean (SD)	180.48 (77.194)		182.04 (87.590)	
		95% CI				

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_cema.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.22 Descriptive Statistics of Urinary CEMA - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Quantity Excreted over 24 hours (µg)	Day 1	n	80	80	41	41
		BLOQ - n (%)	0		0	
		Geometric Mean (CV%)	47.96 (86.34%)		150.22 (64.28%)	
		Geometric 95% CI	40.62, 56.63		124.77, 180.87	
		Median	57.42	-69.13	159.39	-10.56
		Q25, Q75	36.44, 75.36	-74.19, -61.72	96.98, 243.02	-23.40, 5.96
		Min, Max	2.4, 196.0	-87.0, -3.4	46.4, 437.9	-42.9, 69.6
		Mean (SD)	59.04 (34.454)	-67.42 (12.789)	176.40 (99.812)	-4.16 (26.770)
		95% CI		-70.27, -64.57		-12.61, 4.30

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_cema.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.22 Descriptive Statistics of Urinary CEMA - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Quantity Excreted over 24 hours (µg)	Day 2	n	80	80	41	41
		BLOQ - n (%)	0		0	
		Geometric Mean (CV%)	28.37 (86.09%)		146.92 (72.36%)	
		Geometric 95% CI	24.03, 33.49		119.70, 180.32	
		Median	30.94	-82.28	133.68	-9.09
		Q25, Q75	21.63, 43.30	-85.65, -78.99	102.65, 234.00	-22.43, 16.16
		Min, Max	1.8, 260.4	-92.5, 66.3	22.0, 430.6	-88.9, 124.2
		Mean (SD)	36.29 (31.479)	-79.36 (18.253)	176.68 (103.803)	-1.70 (37.424)
		95% CI		-83.43, -75.30		-13.52, 10.11

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_cema.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.22 Descriptive Statistics of Urinary CEMA - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Quantity Excreted over 24 hours (µg)	Day 3	n	79	79	41	41
		BLOQ - n (%)	0		0	
		Geometric Mean (CV%)	25.42 (86.82%)		150.31 (61.59%)	
		Geometric 95% CI	21.49, 30.07		125.67, 179.77	
		Median	27.60	-83.77	141.24	-5.78
		Q25, Q75	19.82, 41.28	-86.94, -80.81	109.80, 229.25	-23.19, 17.61
		Min, Max	1.6, 206.7	-93.4, -13.6	38.2, 399.6	-77.6, 72.5
		Mean (SD)	32.24 (25.838)	-82.62 (9.494)	174.19 (93.493)	-2.39 (29.848)
		95% CI		-84.75, -80.49		-11.82, 7.03

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_cema.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.22 Descriptive Statistics of Urinary CEMA - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Quantity Excreted over 24 hours (µg)	Day 4	n	79	79	41	41
		BLOQ - n (%)	0		0	
		Geometric Mean (CV%)	22.05 (81.63%)		155.61 (57.87%)	
		Geometric 95% CI	18.78, 25.88		131.33, 184.38	
		Median	24.95	-86.08	163.52	-8.03
		Q25, Q75	17.30, 31.89	-88.38, -83.49	109.74, 211.53	-18.92, 7.54
		Min, Max	1.3, 89.5	-92.9, -68.9	54.2, 458.3	-39.0, 138.7
		Mean (SD)	26.80 (15.520)	-85.38 (4.872)	179.07 (99.845)	-0.01 (33.387)
		95% CI		-86.47, -84.28		-10.56, 10.53

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_cema.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.22 Descriptive Statistics of Urinary CEMA - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Quantity Excreted over 24 hours (µg)	Day 5	n	79	79	41	41
		BLOQ - n (%)	0		0	
		Geometric Mean (CV%)	18.92 (89.97%)		149.40 (62.90%)	
		Geometric 95% CI	15.92, 22.48		124.51, 179.27	
		Median	20.88	-88.04	167.09	-11.26
		Q25, Q75	14.20, 30.52	-90.68, -84.64	100.90, 235.72	-22.49, 10.06
		Min, Max	1.5, 84.8	-97.7, -64.1	38.0, 377.7	-57.1, 89.3
		Mean (SD)	23.68 (14.871)	-87.13 (5.137)	173.62 (91.531)	-3.90 (29.187)
		95% CI		-88.29, -85.98		-13.12, 5.32

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_cema.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.22 Descriptive Statistics of Urinary CEMA - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Concentration Adjusted for Creatinine (ng/mg creat)	Baseline	n	80		41	
		BLOQ - n (%)	0		0	
		Geometric Mean (CV%)	98.03 (70.50%)		98.46 (54.54%)	
		Geometric 95% CI	85.10, 112.92		83.81, 115.67	
		Median	99.22		106.49	
		Q25, Q75	74.97, 144.96		75.06, 137.66	
		Min, Max	5.5, 246.1		24.6, 260.0	
		Mean (SD)	113.73 (54.853)		110.22 (49.785)	
		95% CI				

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_cema.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.22 Descriptive Statistics of Urinary CEMA - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Concentration Adjusted for Creatinine (ng/mg creat)	Day 1	n	80	80	41	41
		BLOQ - n (%)	0		0	
		Geometric Mean (CV%)	33.34 (77.34%)		103.03 (54.89%)	
		Geometric 95% CI	28.63, 38.84		87.62, 121.15	
		Median	36.61	-65.11	98.08	4.38
		Q25, Q75	25.30, 53.54	-72.45, -57.38	74.55, 153.46	-8.74, 19.29
		Min, Max	2.1, 107.2	-83.8, -19.2	28.8, 233.5	-36.6, 74.6
		Mean (SD)	39.88 (21.237)	-64.28 (11.315)	115.96 (54.140)	7.34 (25.034)
		95% CI		-66.80, -61.76		-0.57, 15.24

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_cema.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.22 Descriptive Statistics of Urinary CEMA - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Concentration Adjusted for Creatinine (ng/mg creat)	Day 2	n	80	80	41	41
		BLOQ - n (%)	0		0	
		Geometric Mean (CV%)	18.97 (77.43%)		92.12 (65.65%)	
		Geometric 95% CI	16.28, 22.10		76.25, 111.28	
		Median	19.14	-80.76	96.15	0.09
		Q25, Q75	12.42, 31.09	-84.34, -77.50	64.46, 147.61	-19.92, 14.12
		Min, Max	1.4, 76.9	-91.9, 9.3	14.2, 209.8	-80.1, 62.0
		Mean (SD)	23.07 (13.885)	-78.88 (12.461)	106.69 (52.081)	-1.73 (27.440)
		95% CI		-81.65, -76.10		-10.40, 6.93

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_cema.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.22 Descriptive Statistics of Urinary CEMA - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Concentration Adjusted for Creatinine (ng/mg creat)	Day 3	n	79	79	41	41
		BLOQ - n (%)	0		0	
		Geometric Mean (CV%)	17.19 (82.92%)		97.07 (52.85%)	
		Geometric 95% CI	14.62, 20.22		82.99, 113.53	
		Median	19.77	-82.53	98.89	-3.57
		Q25, Q75	11.29, 26.08	-85.68, -79.38	74.85, 147.30	-12.40, 12.85
		Min, Max	1.2, 162.9	-93.0, -23.8	31.5, 220.0	-72.8, 76.3
		Mean (SD)	21.87 (19.482)	-81.49 (8.343)	108.45 (49.102)	2.96 (28.890)
		95% CI		-83.37, -79.62		-6.17, 12.08

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_cema.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.22 Descriptive Statistics of Urinary CEMA - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Concentration Adjusted for Creatinine (ng/mg creat)	Day 4	n	79	79	41	41
		BLOQ - n (%)	0		0	
		Geometric Mean (CV%)	15.09 (78.20%)		102.05 (49.62%)	
		Geometric 95% CI	12.92, 17.62		88.00, 118.34	
		Median	16.64	-84.11	103.36	1.97
		Q25, Q75	9.74, 24.03	-87.59, -81.28	79.68, 160.51	-13.94, 19.57
		Min, Max	1.0, 46.9	-92.3, -68.9	32.1, 196.0	-38.0, 96.6
		Mean (SD)	18.21 (10.131)	-84.07 (4.776)	112.63 (47.461)	6.90 (28.738)
		95% CI		-85.14, -82.99		-2.18, 15.98

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_cema.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.22 Descriptive Statistics of Urinary CEMA - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Concentration Adjusted for Creatinine (ng/mg creat)	Day 5	n	79	79	41	41
		BLOQ - n (%)	0		0	
		Geometric Mean (CV%)	13.18 (73.49%)		99.48 (49.58%)	
		Geometric 95% CI	11.37, 15.27		85.79, 115.35	
		Median	14.58	-86.55	107.68	3.24
		Q25, Q75	8.98, 20.78	-89.49, -83.46	72.41, 141.50	-12.45, 17.72
		Min, Max	1.2, 45.4	-93.6, -72.3	31.7, 207.7	-46.6, 81.9
		Mean (SD)	15.72 (8.785)	-86.10 (4.147)	109.65 (45.982)	4.21 (26.577)
		95% CI		-87.04, -85.17		-4.18, 12.61

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_cema.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.22 Descriptive Statistics of Urinary CEMA - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Quantity Excreted over 24 hours (µg)	Baseline	n	39	
		BLOQ - n (%)	0	
		Geometric Mean (CV%)	163.57 (51.76%)	
		Geometric 95% CI	139.67, 191.56	
		Median	162.05	
		Q25, Q75	103.77, 220.50	
		Min, Max	67.8, 498.9	
		Mean (SD)	184.69 (100.043)	
		95% CI		

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_cema.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.22 Descriptive Statistics of Urinary CEMA - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Quantity Excreted over 24 hours (µg)	Day 1	n	39	39
		BLOQ - n (%)	0	
		Geometric Mean (CV%)	53.05 (60.87%)	
		Geometric 95% CI	44.22, 63.64	
		Median	57.13	-66.04
		Q25, Q75	35.65, 80.58	-74.64, -56.33
		Min, Max	10.6, 138.9	-87.4, -28.5
		Mean (SD)	60.98 (31.387)	-65.20 (13.041)
		95% CI		-69.43, -60.97

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_cema.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.22 Descriptive Statistics of Urinary CEMA - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Quantity Excreted over 24 hours (µg)	Day 2	n	39	39
		BLOQ - n (%)	0	
		Geometric Mean (CV%)	29.37 (63.90%)	
		Geometric 95% CI	24.29, 35.51	
		Median	27.50	-81.60
		Q25, Q75	22.81, 42.92	-86.59, -74.27
		Min, Max	4.7, 74.6	-94.4, -62.9
		Mean (SD)	33.96 (17.346)	-80.58 (7.515)
		95% CI		-83.02, -78.14

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_cema.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.22 Descriptive Statistics of Urinary CEMA - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Quantity Excreted over 24 hours (µg)	Day 3	n	39	39
		BLOQ - n (%)	0	
		Geometric Mean (CV%)	25.10 (64.84%)	
		Geometric 95% CI	20.71, 30.42	
		Median	27.72	-83.99
		Q25, Q75	18.40, 39.83	-87.83, -78.97
		Min, Max	3.1, 63.1	-96.3, -71.0
		Mean (SD)	28.98 (14.410)	-83.55 (5.639)
		95% CI		-85.38, -81.72

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_cema.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.22 Descriptive Statistics of Urinary CEMA - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Quantity Excreted over 24 hours (µg)	Day 4	n	39	39
		BLOQ - n (%)	0	
		Geometric Mean (CV%)	21.31 (73.93%)	
		Geometric 95% CI	17.20, 26.40	
		Median	22.41	-86.24
		Q25, Q75	15.32, 35.88	-89.77, -81.67
		Min, Max	2.5, 57.6	-97.0, -70.3
		Mean (SD)	25.45 (14.044)	-85.75 (5.900)
		95% CI		-87.67, -83.83

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_cema.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.22 Descriptive Statistics of Urinary CEMA - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Quantity Excreted over 24 hours (µg)	Day 5	n	39	39
		BLOQ - n (%)	0	
		Geometric Mean (CV%)	19.77 (74.00%)	
		Geometric 95% CI	15.95, 24.50	
		Median	19.58	-87.40
		Q25, Q75	13.20, 32.56	-91.02, -82.66
		Min, Max	2.1, 71.2	-97.5, -71.7
		Mean (SD)	23.83 (14.469)	-86.73 (5.554)
		95% CI		-88.53, -84.92

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_cema.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.22 Descriptive Statistics of Urinary CEMA - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Concentration Adjusted for Creatinine (ng/mg creat)	Baseline	n	39	
		BLOQ - n (%)	0	
		Geometric Mean (CV%)	103.51 (47.80%)	
		Geometric 95% CI	89.35, 119.91	
		Median	111.65	
		Q25, Q75	70.04, 159.46	
		Min, Max	39.8, 237.7	
		Mean (SD)	114.08 (50.465)	
		95% CI		

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_cema.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.22 Descriptive Statistics of Urinary CEMA - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Concentration Adjusted for Creatinine (ng/mg creat)	Day 1	n	39	39
		BLOQ - n (%)	0	
		Geometric Mean (CV%)	35.71 (52.41%)	
		Geometric 95% CI	30.43, 41.90	
		Median	38.41	-62.57
		Q25, Q75	26.75, 51.32	-72.59, -56.35
		Min, Max	10.3, 100.4	-85.3, -34.7
		Mean (SD)	39.92 (18.879)	-63.45 (11.950)
		95% CI		-67.32, -59.57

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_cema.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.22 Descriptive Statistics of Urinary CEMA - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Concentration Adjusted for Creatinine (ng/mg creat)	Day 2	n	39	39
		BLOQ - n (%)	0	
		Geometric Mean (CV%)	19.09 (58.96%)	
		Geometric 95% CI	15.99, 22.79	
		Median	18.23	-79.31
		Q25, Q75	14.17, 26.95	-85.69, -75.01
		Min, Max	5.1, 59.9	-92.8, -67.8
		Mean (SD)	22.00 (12.286)	-80.47 (6.204)
		95% CI		-82.48, -78.45

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_cema.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.22 Descriptive Statistics of Urinary CEMA - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Concentration Adjusted for Creatinine (ng/mg creat)	Day 3	n	39	39
		BLOQ - n (%)	0	
		Geometric Mean (CV%)	16.36 (62.87%)	
		Geometric 95% CI	13.57, 19.73	
		Median	16.39	-83.05
		Q25, Q75	12.07, 24.30	-87.27, -78.07
		Min, Max	3.1, 53.0	-95.5, -72.6
		Mean (SD)	19.04 (10.679)	-83.19 (5.470)
		95% CI		-84.97, -81.42

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_cema.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.22 Descriptive Statistics of Urinary CEMA - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Concentration Adjusted for Creatinine (ng/mg creat)	Day 4	n	39	39
		BLOQ - n (%)	0	
		Geometric Mean (CV%)	14.48 (70.03%)	
		Geometric 95% CI	11.79, 17.78	
		Median	13.92	-84.82
		Q25, Q75	10.59, 21.93	-89.00, -80.23
		Min, Max	2.4, 48.5	-96.6, -72.6
		Mean (SD)	17.32 (10.440)	-84.92 (5.432)
		95% CI		-86.68, -83.15

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_cema.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.22 Descriptive Statistics of Urinary CEMA - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Concentration Adjusted for Creatinine (ng/mg creat)	Day 5	n	39	39
		BLOQ - n (%)	0	
		Geometric Mean (CV%)	12.60 (76.28%)	
		Geometric 95% CI	10.12, 15.70	
		Median	12.79	-87.19
		Q25, Q75	9.33, 18.29	-90.68, -82.35
		Min, Max	1.3, 44.4	-98.2, -77.9
		Mean (SD)	15.27 (9.243)	-86.74 (4.804)
		95% CI		-88.30, -85.17

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_cema.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.23 Analysis of Urinary HEMA on Day 5 - FAS**

Variable	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	THS : CC Ratio (%)	THS : SA Ratio (%)
Urinary Quantity over n 24 hours (ng)		79	41	39		
	Geometric LS Mean (SE) ¹	1966.21	6268.43	2036.38	31.37 (0.086)	96.55 (0.087)
	95% CI	1782.78, 2168.51	5469.94, 7183.49	1771.37, 2341.04	26.51, 37.11	81.43, 114.49
Concentration Adjusted for Creatinine (pg/mg creat)	n	79	41	39		
	Geometric LS Mean (SE) ¹	1363.28	4260.11	1286.99	32.00 (0.085)	105.93 (0.086)
	95% CI	1237.33, 1502.04	3722.95, 4874.78	1121.14, 1477.36	27.10, 37.79	89.49, 125.38

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Adjusted geometric least squares (LS) means and confidence intervals (CIs) from an ANCOVA model conducted on log-transformed Day 5 values with log-transformed baseline value, study arm, sex and CC consumption reported at screening as fixed effect factors.

Baseline is defined as the last assessment prior to Day 1 product use for THS 2.2 and CC subjects and prior to 06:29 AM on Day 1 for SA subjects.

Note: 1 = Standard errors presented from analysis performed on the log_e scale.

Note: Collection over 24 hours starting on the Day stated in the title.

Appendix 15.4.4.23 and 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/tl_anlsecondbio.sas

Program Run: 27OCT14 cvn_ahedge Program Status: FINAL

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**Table 15.2.4.24 Descriptive Statistics of Urinary HEMA - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Quantity Excreted over Baseline 24 hours (ng)	n		80		41	
	BLOQ - n (%)		1 (1.3%)		0	
	Geometric Mean (CV%)		6673.02 (99.83%)		7775.33 (105.57%)	
	Geometric 95% CI		5545.77, 8029.39		5917.01, 10217.28	
	Median		6681.25		7568.00	
	Q25, Q75		4288.50, 11514.50		4012.00, 14962.50	
	Min, Max		105.0, 31500.0		1082.8, 38915.0	
	Mean (SD)		8755.34 (6404.969)		10828.17 (8792.124)	
	95% CI					

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.100 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hema.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.24 Descriptive Statistics of Urinary HEMA - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Quantity Excreted over Day 1 24 hours (ng)	n		80	80	41	41
	BLOQ - n (%)		1 (1.3%)		0	
	Geometric Mean (CV%)		3455.29 (88.55%)		6645.36 (99.01%)	
	Geometric 95% CI		2917.12, 4092.75		5119.38, 8626.19	
	Median		3664.50	-48.63	7144.00	-18.21
	Q25, Q75		2314.25, 5181.25	-59.73, -31.16	3417.00, 13030.50	-27.07, -7.70
	Min, Max		85.0, 17729.5	-75.8, 7.4	1140.0, 25575.0	-49.6, 62.4
	Mean (SD)		4372.97 (3039.484)	-44.97 (19.509)	8936.06 (6550.942)	-11.15 (26.978)
	95% CI			-49.32, -40.62		-19.67, -2.63

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.100 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hema.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.24 Descriptive Statistics of Urinary HEMA - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Quantity Excreted over Day 2 24 hours (ng)	n		80	80	41	41
	BLOQ - n (%)		1 (1.3%)		0	
	Geometric Mean (CV%)		2481.01 (79.96%)		7510.42 (117.74%)	
	Geometric 95% CI		2121.68, 2901.19		5595.34, 10080.95	
	Median		2405.90	-67.30	9072.00	-2.44
	Q25, Q75		1592.32, 3956.50	-74.95, -45.37	3850.00, 15427.75	-26.47, 18.71
	Min, Max		97.5, 10522.5	-85.6, 46.5	682.0, 40162.5	-90.3, 265.5
	Mean (SD)		3073.71 (2064.876)	-56.72 (26.905)	10738.04 (8731.853)	7.76 (54.585)
	95% CI			-62.71, -50.73		-9.47, 25.00

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.100 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hema.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.24 Descriptive Statistics of Urinary HEMA - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Quantity Excreted over Day 3 24 hours (ng)	n		79	79	41	41
	BLOQ - n (%)		1 (1.3%)		0	
	Geometric Mean (CV%)		1872.97 (82.48%)		6646.31 (88.19%)	
	Geometric 95% CI		1593.87, 2200.95		5231.25, 8444.13	
	Median		1845.60	-74.82	6456.00	-15.02
	Q25, Q75		1260.25, 2925.00	-80.63, -57.01	3780.00, 10900.50	-31.55, 3.04
	Min, Max		87.5, 8880.0	-89.3, -14.2	1279.8, 22440.0	-58.5, 93.0
	Mean (SD)		2340.16 (1581.002)	-68.01 (18.007)	8590.55 (5955.996)	-10.28 (29.621)
	95% CI			-72.05, -63.97		-19.64, -0.93

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.100 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hema.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.24 Descriptive Statistics of Urinary HEMA - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Quantity Excreted over Day 4 24 hours (ng)	n		79	79	41	41
	BLOQ - n (%)		1 (1.3%)		0	
	Geometric Mean (CV%)		2308.74 (79.83%)		7609.37 (93.54%)	
	Geometric 95% CI		1972.75, 2701.95		5924.72, 9773.04	
	Median		2475.00	-66.82	7429.00	-0.41
	Q25, Q75		1482.75, 3881.25	-76.65, -48.91	4125.00, 13879.25	-14.72, 20.05
	Min, Max		100.0, 10062.0	-90.6, 22.7	1151.4, 36037.5	-80.6, 103.3
	Mean (SD)		2827.11 (1738.167)	-60.96 (20.889)	10128.94 (7839.813)	5.37 (39.109)
	95% CI			-65.64, -56.27		-6.98, 17.72

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.100 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hema.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.24 Descriptive Statistics of Urinary HEMA - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Quantity Excreted over Day 5 24 hours (ng)	n		79	79	41	41
	BLOQ - n (%)		1 (1.3%)		0	
	Geometric Mean (CV%)		1927.33 (72.36%)		6764.14 (90.44%)	
	Geometric 95% CI		1666.61, 2228.83		5299.48, 8633.58	
	Median		1922.40	-72.02	7650.00	-9.97
	Q25, Q75		1275.55, 3084.25	-80.38, -60.87	3720.60, 11210.00	-26.61, 5.70
	Min, Max		225.0, 6240.0	-91.2, 114.3	1159.2, 32240.0	-55.0, 123.9
	Mean (SD)		2329.57 (1408.346)	-65.75 (26.699)	8832.40 (6505.239)	-7.26 (36.024)
	95% CI			-71.73, -59.76		-18.63, 4.12

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.100 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hema.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.24 Descriptive Statistics of Urinary HEMA - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Baseline	n	80		41	
		BLOQ - n (%)	1 (1.3%)		0	
		Geometric Mean (CV%)	4161.66 (110.90%)		4718.48 (106.89%)	
		Geometric 95% CI	3409.70, 5079.44		3582.18, 6215.23	
		Median	4500.83		4883.72	
		Q25, Q75	2416.50, 7909.24		2402.82, 9913.80	
		Min, Max	68.8, 25495.8		990.3, 30887.4	
		Mean (SD)	5718.28 (4509.109)		6752.57 (6141.565)	
		95% CI				

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.100 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hema.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.4.24 Descriptive Statistics of Urinary HEMA - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Day 1	n	80	80	41	41
		BLOQ - n (%)	1 (1.3%)		0	
		Geometric Mean (CV%)	2402.18 (97.96%)		4557.69 (96.93%)	
		Geometric 95% CI	2001.41, 2883.19		3525.14, 5892.68	
		Median	2582.94	-42.83	4743.20	-3.36
		Q25, Q75	1371.90, 4388.04	-52.80, -29.05	2521.24, 8198.85	-17.47, 10.07
		Min, Max	68.5, 12897.0	-69.2, 6.1	1061.1, 22398.8	-44.0, 88.8
		Mean (SD)	3202.11 (2507.209)	-39.54 (18.712)	6200.94 (5039.279)	-0.38 (26.102)
		95% CI		-43.71, -35.37		-8.63, 7.86

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.100 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hema.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.24 Descriptive Statistics of Urinary HEMA - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Day 2	n	80	80	41	41
		BLOQ - n (%)	1 (1.3%)		0	
		Geometric Mean (CV%)	1658.91 (85.86%)		4709.04 (113.50%)	
		Geometric 95% CI	1406.05, 1957.24		3533.62, 6275.46	
		Median	1695.47	-65.36	4501.31	1.76
		Q25, Q75	1090.31, 2578.41	-72.72, -41.67	2187.98, 9320.89	-19.00, 17.82
		Min, Max	70.5, 7033.3	-83.9, 25.6	440.6, 22495.3	-82.7, 168.0
		Mean (SD)	2116.49 (1538.759)	-54.38 (25.660)	6725.26 (5546.472)	7.70 (42.253)
		95% CI			-60.10, -48.67	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.100 pg/mL).

Appendix 15.3.3.2

Path: /cvm/projects/prj/development/000000106324/dev/tables/t_hema.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.24 Descriptive Statistics of Urinary HEMA - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Day 3	n	79	79	41	41
		BLOQ - n (%)	1 (1.3%)		0	
		Geometric Mean (CV%)	1266.78 (86.70%)		4292.17 (91.91%)	
		Geometric 95% CI	1071.20, 1498.08		3352.80, 5494.71	
		Median	1338.03	-72.18	3898.64	-8.12
		Q25, Q75	725.46, 2061.86	-79.38, -55.30	2602.73, 7514.02	-24.21, 12.51
		Min, Max	73.2, 6997.6	-87.9, 6.4	950.2, 19465.6	-56.2, 113.1
		Mean (SD)	1632.54 (1221.454)	-65.79 (18.279)	5737.36 (4553.117)	-4.13 (33.005)
		95% CI		-69.89, -61.69		-14.56, 6.29

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.100 pg/mL).

Appendix 15.3.3.2

Path: /cvm/projects/prj/development/000000106324/dev/tables/t_hema.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.24 Descriptive Statistics of Urinary HEMA - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Day 4	n	79	79	41	41
		BLOQ - n (%)	1 (1.3%)		0	
		Geometric Mean (CV%)	1580.48 (89.22%)		4990.40 (101.16%)	
		Geometric 95% CI	1331.53, 1875.98		3828.76, 6504.47	
		Median	1852.23	-63.79	4736.84	8.87
		Q25, Q75	862.06, 2789.12	-74.15, -44.11	2429.02, 9760.69	-14.06, 29.58
		Min, Max	78.7, 7543.9	-90.4, 22.9	1024.7, 29026.2	-87.4, 138.0
		Mean (SD)	2033.41 (1429.083)	-56.92 (23.497)	6970.35 (6115.553)	14.38 (41.141)
		95% CI		-62.19, -51.65		1.39, 27.37

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.100 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hema.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.24 Descriptive Statistics of Urinary HEMA - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Day 5	n	79	79	41	41
		BLOQ - n (%)	1 (1.3%)		0	
		Geometric Mean (CV%)	1342.40 (83.59%)		4504.00 (93.56%)	
		Geometric 95% CI	1140.44, 1580.12		3506.73, 5784.88	
		Median	1276.74	-69.31	4751.13	-3.61
		Q25, Q75	766.45, 2553.70	-79.73, -53.32	2451.26, 8038.38	-18.92, 14.01
		Min, Max	183.8, 6280.2	-89.9, 167.3	1092.3, 23853.2	-63.9, 87.5
		Mean (SD)	1721.74 (1241.124)	-60.71 (33.908)	6064.19 (4979.859)	0.74 (33.567)
		95% CI		-68.31, -53.11		-9.86, 11.34

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.100 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hema.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.24 Descriptive Statistics of Urinary HEMA - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Quantity Excreted over 24 hours (ng)	Baseline	n	39	
		BLOQ - n (%)	0	
		Geometric Mean (CV%)	6502.25 (97.40%)	
		Geometric 95% CI	4989.76, 8473.20	
		Median	6675.00	
		Q25, Q75	3812.75, 10378.50	
		Min, Max	1628.1, 36480.0	
		Mean (SD)	9027.88 (8101.137)	
		95% CI		

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.100 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hema.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.24 Descriptive Statistics of Urinary HEMA - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Quantity Excreted over 24 hours (ng)	Day 1	n	39	39
		BLOQ - n (%)	0	
		Geometric Mean (CV%)	4103.30 (75.05%)	
		Geometric 95% CI	3303.97, 5096.01	
		Median	4150.00	-42.04
		Q25, Q75	2560.00, 5938.50	-52.60, -9.40
		Min, Max	918.0, 17050.0	-65.3, 80.0
		Mean (SD)	5090.68 (3610.664)	-31.21 (31.560)
		95% CI		-41.44, -20.97

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.100 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hema.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.24 Descriptive Statistics of Urinary HEMA - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Quantity Excreted over 24 hours (ng)	Day 2	n	39	39
		BLOQ - n (%)	0	
		Geometric Mean (CV%)	2884.55 (77.79%)	
		Geometric 95% CI	2308.04, 3605.07	
		Median	3225.00	-56.86
		Q25, Q75	1632.00, 4558.00	-71.45, -32.48
		Min, Max	555.5, 11100.0	-84.4, 66.2
		Mean (SD)	3564.20 (2299.459)	-47.24 (34.447)
		95% CI		-58.41, -36.07

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.100 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hema.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.24 Descriptive Statistics of Urinary HEMA - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Quantity Excreted over 24 hours (ng)	Day 3	n	39	39
		BLOQ - n (%)	0	
		Geometric Mean (CV%)	2128.20 (66.55%)	
		Geometric 95% CI	1748.92, 2589.74	
		Median	2313.30	-68.81
		Q25, Q75	1344.00, 2882.00	-79.49, -49.59
		Min, Max	507.8, 7395.0	-89.8, -10.4
		Mean (SD)	2533.24 (1593.320)	-61.60 (22.859)
		95% CI		-69.01, -54.18

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.100 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hema.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.24 Descriptive Statistics of Urinary HEMA - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Quantity Excreted over 24 hours (ng)	Day 4	n	39	39
		BLOQ - n (%)	1 (2.6%)	
		Geometric Mean (CV%)	2329.30 (88.12%)	
		Geometric 95% CI	1821.84, 2978.09	
		Median	2519.83	-65.40
		Q25, Q75	1447.50, 3956.00	-76.64, -40.13
		Min, Max	220.0, 9555.0	-87.5, 21.4
		Mean (SD)	2967.84 (2067.576)	-57.33 (26.838)
		95% CI		-66.03, -48.62

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.100 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hema.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.24 Descriptive Statistics of Urinary HEMA - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Quantity Excreted over 24 hours (ng)	Day 5	n	39	39
		BLOQ - n (%)	0	
		Geometric Mean (CV%)	1958.16 (77.89%)	
		Geometric 95% CI	1566.43, 2447.86	
		Median	2057.18	-73.72
		Q25, Q75	1213.59, 2775.00	-80.52, -47.19
		Min, Max	371.9, 11760.0	-90.4, -9.4
		Mean (SD)	2497.24 (2097.155)	-63.56 (23.203)
		95% CI		-71.09, -56.04

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.100 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hema.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.24 Descriptive Statistics of Urinary HEMA - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Baseline	n	39	
		BLOQ - n (%)	0	
		Geometric Mean (CV%)	4114.78 (102.33%)	
		Geometric 95% CI	3127.42, 5413.84	
		Median	4758.06	
		Q25, Q75	2450.65, 7021.28	
		Min, Max	797.6, 25494.1	
		Mean (SD)	5812.33 (5544.552)	
		95% CI		

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.100 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hema.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.24 Descriptive Statistics of Urinary HEMA - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Day 1	n	39	39
		BLOQ - n (%)	0	
		Geometric Mean (CV%)	2762.05 (84.93%)	
		Geometric 95% CI	2175.15, 3507.31	
		Median	2822.38	-37.40
		Q25, Q75	1676.47, 4447.48	-50.91, -16.02
		Min, Max	564.6, 14187.6	-60.8, 103.7
		Mean (SD)	3582.88 (2864.770)	-27.52 (33.116)
		95% CI		-38.26, -16.78

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.100 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hema.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.24 Descriptive Statistics of Urinary HEMA - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Day 2	n	39	39
		BLOQ - n (%)	0	
		Geometric Mean (CV%)	1875.03 (84.51%)	
		Geometric 95% CI	1477.96, 2378.77	
		Median	1991.47	-57.60
		Q25, Q75	1087.61, 3170.74	-67.88, -36.97
		Min, Max	321.7, 8434.7	-82.9, 57.1
		Mean (SD)	2367.72 (1598.271)	-47.45 (31.566)
		95% CI		-57.69, -37.21

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.100 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hema.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.24 Descriptive Statistics of Urinary HEMA - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Day 3	n	39	39
		BLOQ - n (%)	0	
		Geometric Mean (CV%)	1387.25 (76.06%)	
		Geometric 95% CI	1114.40, 1726.90	
		Median	1437.50	-69.78
		Q25, Q75	963.13, 2068.97	-76.04, -50.31
		Min, Max	283.3, 5979.4	-84.8, -4.7
		Mean (SD)	1724.08 (1225.061)	-61.56 (21.241)
		95% CI		-68.45, -54.67

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.100 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hema.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.24 Descriptive Statistics of Urinary HEMA - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Day 4	n	39	39
		BLOQ - n (%)	1 (2.6%)	
		Geometric Mean (CV%)	1583.02 (98.12%)	
		Geometric 95% CI	1213.06, 2065.80	
		Median	1749.43	-62.40
		Q25, Q75	921.38, 2352.95	-75.52, -41.54
		Min, Max	177.9, 7414.4	-86.5, 4.0
		Mean (SD)	2142.09 (1753.784)	-56.16 (23.022)
		95% CI		-63.63, -48.69

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.100 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hema.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.24 Descriptive Statistics of Urinary HEMA - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Day 5	n	39	39
		BLOQ - n (%)	0	
		Geometric Mean (CV%)	1248.27 (86.06%)	
		Geometric 95% CI	980.62, 1588.98	
		Median	1173.08	-74.52
		Q25, Q75	746.87, 2104.84	-81.56, -49.00
		Min, Max	219.7, 6494.8	-86.4, -8.3
		Mean (SD)	1643.48 (1403.807)	-64.48 (21.004)
		95% CI		-71.29, -57.66

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (0.100 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hema.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.25 Analysis of Urinary 3-hydroxy(a)benzopyrene on Day 5 - FAS**

Variable	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	THS : CC Ratio (%)	THS : SA Ratio (%)
Urinary Quantity over n 24 hours (pg)	n	79	41	39		
	Geometric LS Mean (SE) ¹	53.12	199.50	51.80	26.62 (0.089)	102.54 (0.091)
	95% CI	47.94, 58.85	173.01, 230.03	44.76, 59.94	22.33, 31.74	85.80, 122.56
Concentration Adjusted for Creatinine (fg/mg creat)	n	79	41	39		
	Geometric LS Mean (SE) ¹	36.89	134.13	33.05	27.50 (0.085)	111.62 (0.087)
	95% CI	33.45, 40.68	117.08, 153.67	28.74, 37.99	23.25, 32.52	94.16, 132.32

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Adjusted geometric least squares (LS) means and confidence intervals (CIs) from an ANCOVA model conducted on log-transformed Day 5 values with log-transformed baseline value, study arm, sex and CC consumption reported at screening as fixed effect factors.

Baseline is defined as the last assessment prior to Day 1 product use for THS 2.2 and CC subjects and prior to 06:29 AM on Day 1 for SA subjects.

Note: 1 = Standard errors presented from analysis performed on the log_e scale.

Note: Collection over 24 hours starting on the Day stated in the title.

Appendix 15.4.4.25 and 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/tl_anl3OH_BAP.sas

Program Run: 05NOV15 cvn_ahedge Program Status: FINAL

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**Table 15.2.4.26 Descriptive Statistics of Urinary B[a]P - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Quantity Excreted over Baseline 24 hours (pg)	n		80		41	
	BLOQ - n (%)		1 (1.3%)		0	
	Geometric Mean (CV%)		258.43 (61.68%)		246.30 (50.33%)	
	Geometric 95% CI		227.75, 293.24		211.99, 286.15	
	Median		278.25		241.50	
	Q25, Q75		180.52, 373.43		161.69, 324.83	
	Min, Max		27.5, 806.8		121.7, 771.8	
	Mean (SD)		297.63 (154.252)		276.43 (145.179)	
	95% CI					

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (25.0 fg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_bap.sas

Program Run: 05NOV15 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.26 Descriptive Statistics of Urinary B[a]P - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Quantity Excreted over Day 1 24 hours (pg)	n		80	80	41	41
	BLOQ - n (%)		7 (8.8%)		1 (2.4%)	
	Geometric Mean (CV%)		108.19 (61.05%)		224.06 (57.62%)	
	Geometric 95% CI		95.45, 122.63		189.22, 265.31	
	Median		112.94	-59.20	245.10	-5.72
	Q25, Q75		77.97, 157.79	-67.62, -48.21	163.59, 300.30	-19.03, 3.88
	Min, Max		25.6, 389.2	-85.1, 11.8	37.5, 754.8	-69.7, 63.6
	Mean (SD)		125.31 (69.412)	-55.28 (17.097)	254.67 (132.210)	-6.39 (21.129)
	95% CI			-59.09, -51.47		-13.06, 0.28

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (25.0 fg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_bap.sas

Program Run: 05NOV15 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.26 Descriptive Statistics of Urinary B[a]P - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Quantity Excreted over Day 2 24 hours (pg)	n		80	80	41	41
	BLOQ - n (%)		25 (31.3%)		1 (2.4%)	
	Geometric Mean (CV%)		67.03 (64.24%)		216.89 (57.73%)	
	Geometric 95% CI		58.81, 76.40		183.11, 256.90	
	Median		74.18	-74.20	214.20	-7.20
	Q25, Q75		37.03, 102.85	-81.70, -63.05	159.68, 310.50	-25.87, 10.48
	Min, Max		19.4, 442.4	-94.4, 58.4	37.2, 725.0	-76.9, 93.5
	Mean (SD)		79.78 (55.885)	-68.73 (22.693)	246.13 (125.136)	-6.20 (31.320)
	95% CI			-73.79, -63.68		-16.09, 3.69

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (25.0 fg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_bap.sas

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Program Run: 05NOV15 cvn_kbooth Program Status: FINAL

**Table 15.2.4.26 Descriptive Statistics of Urinary B[a]P - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Quantity Excreted over Day 3 24 hours (pg)	n		79	79	41	41
	BLOQ - n (%)		23 (29.1%)		1 (2.4%)	
	Geometric Mean (CV%)		63.68 (61.82%)		208.56 (59.57%)	
	Geometric 95% CI		56.06, 72.34		175.26, 248.18	
	Median		71.60	-75.74	215.90	-10.50
	Q25, Q75		34.37, 98.40	-82.76, -63.69	154.77, 278.40	-26.65, -2.08
	Min, Max		17.8, 147.2	-95.3, -3.4	33.1, 762.9	-72.8, 65.4
	Mean (SD)		73.46 (36.123)	-71.38 (16.582)	239.63 (134.409)	-12.23 (22.882)
	95% CI			-75.09, -67.66		-19.45, -5.00

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (25.0 fg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_bap.sas

Program Run: 05NOV15 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.26 Descriptive Statistics of Urinary B[a]P - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Quantity Excreted over Day 4 24 hours (pg)	n		79	79	41	41
	BLOQ - n (%)		44 (55.7%)		0	
	Geometric Mean (CV%)		49.22 (59.93%)		213.36 (51.51%)	
	Geometric 95% CI		43.47, 55.72		183.07, 248.66	
	Median		45.00	-82.73	225.44	-15.64
	Q25, Q75		32.50, 74.50	-86.43, -73.88	153.01, 295.80	-24.63, 2.54
	Min, Max		20.0, 434.8	-94.4, 106.1	82.1, 791.4	-49.2, 107.2
	Mean (SD)		58.97 (51.006)	-76.13 (26.280)	240.17 (130.103)	-10.53 (25.717)
	95% CI			-82.02, -70.24		-18.66, -2.41

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (25.0 fg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_bap.sas

Program Run: 05NOV15 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.26 Descriptive Statistics of Urinary B[a]P - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Quantity Excreted over Day 5 24 hours (pg)	n		79	79	41	41
	BLOQ - n (%)		49 (62.0%)		2 (4.9%)	
	Geometric Mean (CV%)		53.22 (49.47%)		195.67 (63.86%)	
	Geometric 95% CI		47.92, 59.10		162.68, 235.34	
	Median		55.63	-78.31	191.54	-15.12
	Q25, Q75		35.00, 72.88	-85.55, -72.42	142.74, 283.99	-31.94, -1.54
	Min, Max		19.4, 153.7	-96.7, 22.7	53.1, 759.9	-71.0, 68.7
	Mean (SD)		59.36 (29.003)	-75.55 (17.235)	230.71 (141.707)	-16.23 (27.064)
	95% CI			-79.42, -71.69		-24.78, -7.68

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (25.0 fg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_bap.sas

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Program Run: 05NOV15 cvn_kbooth Program Status: FINAL

**Table 15.2.4.26 Descriptive Statistics of Urinary B[a]P - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Concentration Adjusted for Creatinine (fg/mg creat)	Baseline	n	80		41	
		BLOQ - n (%)	1 (1.3%)		0	
		Geometric Mean (CV%)	161.17 (60.71%)		149.47 (49.43%)	
		Geometric 95% CI	142.28, 182.57		128.96, 173.23	
		Median	162.56		152.10	
		Q25, Q75	115.86, 251.98		106.91, 205.36	
		Min, Max	18.6, 602.6		50.2, 477.8	
		Mean (SD)	185.63 (99.277)		166.51 (83.370)	
		95% CI				

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (25.0 fg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_bap.sas

Program Run: 05NOV15 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.26 Descriptive Statistics of Urinary B[a]P - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Concentration Adjusted for Creatinine (fg/mg creat)	Day 1	n	80	80	41	41
		BLOQ - n (%)	7 (8.8%)		1 (2.4%)	
		Geometric Mean (CV%)	75.21 (54.98%)		153.67 (53.25%)	
		Geometric 95% CI	67.08, 84.33		131.24, 179.92	
		Median	78.75	-51.93	158.26	5.81
		Q25, Q75	51.26, 118.00	-63.49, -43.04	117.85, 216.77	-5.78, 18.39
		Min, Max	19.0, 208.7	-83.7, 36.5	42.7, 478.3	-63.0, 44.0
		Mean (SD)	84.90 (40.655)	-50.54 (17.978)	172.65 (86.287)	5.52 (21.636)
		95% CI		-54.54, -46.53		-1.31, 12.36

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (25.0 fg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_bap.sas

Program Run: 05NOV15 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.26 Descriptive Statistics of Urinary B[a]P - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Concentration Adjusted for Creatinine (fg/mg creat)	Day 2	n	80	80	41	41
		BLOQ - n (%)	25 (31.3%)		1 (2.4%)	
		Geometric Mean (CV%)	44.82 (58.02%)		135.99 (55.68%)	
		Geometric 95% CI	39.75, 50.53		115.42, 160.23	
		Median	49.46	-70.51	144.93	-1.95
		Q25, Q75	31.76, 61.04	-81.37, -57.64	114.71, 197.03	-15.61, 9.24
		Min, Max	14.2, 133.6	-93.4, 32.0	32.5, 372.4	-71.8, 40.8
		Mean (SD)	51.25 (26.308)	-67.36 (20.077)	153.30 (73.906)	-5.42 (23.206)
		95% CI		-71.84, -62.89		-12.75, 1.91

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (25.0 fg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_bap.sas

Program Run: 05NOV15 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.26 Descriptive Statistics of Urinary B[a]P - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Concentration Adjusted for Creatinine (fg/mg creat)	Day 3	n	79	79	41	41
		BLOQ - n (%)	23 (29.1%)		1 (2.4%)	
		Geometric Mean (CV%)	43.07 (55.73%)		134.68 (55.48%)	
		Geometric 95% CI	38.33, 48.39		114.36, 158.61	
		Median	47.59	-72.85	136.30	-6.64
		Q25, Q75	29.78, 61.42	-81.92, -58.81	94.96, 192.26	-21.52, 6.89
		Min, Max	14.9, 105.1	-94.5, 7.9	35.3, 390.7	-66.8, 35.7
		Mean (SD)	48.67 (22.779)	-68.83 (18.069)	152.73 (78.792)	-7.07 (21.479)
		95% CI		-72.89, -64.78		-13.86, -0.29

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (25.0 fg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_bap.sas

Program Run: 05NOV15 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.26 Descriptive Statistics of Urinary B[a]P - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Concentration Adjusted for Creatinine (fg/mg creat)	Day 4	n	79	79	41	41
		BLOQ - n (%)	44 (55.7%)		0	
		Geometric Mean (CV%)	33.69 (54.16%)		139.93 (50.98%)	
		Geometric 95% CI	30.07, 37.75		120.23, 162.85	
		Median	32.31	-81.33	131.27	-9.13
		Q25, Q75	22.40, 47.48	-85.75, -71.06	96.37, 186.62	-16.52, 12.83
		Min, Max	12.5, 204.3	-93.6, 124.6	56.7, 445.5	-54.5, 43.9
		Mean (SD)	38.79 (25.724)	-73.74 (28.808)	157.41 (84.681)	-4.22 (20.442)
		95% CI		-80.19, -67.28		-10.68, 2.23

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (25.0 fg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_bap.sas

Program Run: 05NOV15 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.26 Descriptive Statistics of Urinary B[a]P - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Concentration Adjusted for Creatinine (fg/mg creat)	Day 5	n	79	79	41	41
		BLOQ - n (%)	49 (62.0%)		2 (4.9%)	
		Geometric Mean (CV%)	37.07 (51.43%)		130.29 (57.10%)	
		Geometric 95% CI	33.25, 41.32		110.17, 154.07	
		Median	40.06	-76.35	125.65	-7.99
		Q25, Q75	23.06, 57.17	-84.27, -70.16	104.04, 182.99	-27.89, 8.62
		Min, Max	12.5, 96.2	-94.4, 46.9	36.2, 520.5	-62.8, 52.1
		Mean (SD)	41.41 (19.100)	-71.43 (23.294)	149.21 (86.583)	-8.92 (25.583)
		95% CI		-76.65, -66.21		-17.00, -0.84

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (25.0 fg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_bap.sas

Program Run: 05NOV15 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.26 Descriptive Statistics of Urinary B[a]P - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Quantity Excreted over 24 hours (pg)	Baseline	n	39	
		BLOQ - n (%)	0	
		Geometric Mean (CV%)	267.97 (54.58%)	
		Geometric 95% CI	227.09, 316.21	
		Median	260.70	
		Q25, Q75	201.13, 359.93	
		Min, Max	81.0, 960.8	
		Mean (SD)	303.70 (163.558)	
		95% CI		

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (25.0 fg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_bap.sas

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Program Run: 05NOV15 cvn_kbooth Program Status: FINAL

**Table 15.2.4.26 Descriptive Statistics of Urinary B[a]P - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Quantity Excreted over 24 hours (pg)	Day 1	n	39	39
		BLOQ - n (%)	3 (7.7%)	
		Geometric Mean (CV%)	116.35 (84.46%)	
		Geometric 95% CI	91.71, 147.60	
		Median	113.73	-57.62
		Q25, Q75	79.38, 150.26	-68.77, -49.78
		Min, Max	26.3, 1131.0	-78.4, 280.2
		Mean (SD)	160.55 (190.725)	-48.09 (55.884)
		95% CI		-66.21, -29.97

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (25.0 fg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_bap.sas

Program Run: 05NOV15 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.26 Descriptive Statistics of Urinary B[a]P - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Quantity Excreted over 24 hours (pg)	Day 2	n	39	39
		BLOQ - n (%)	15 (38.5%)	
		Geometric Mean (CV%)	66.29 (78.72%)	
		Geometric 95% CI	52.93, 83.03	
		Median	68.75	-72.81
		Q25, Q75	36.25, 117.51	-81.67, -58.71
		Min, Max	19.4, 201.9	-92.6, -22.8
		Mean (SD)	82.68 (53.758)	-70.87 (16.325)
		95% CI		-76.16, -65.57

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (25.0 fg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_bap.sas

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Program Run: 05NOV15 cvn_kbooth Program Status: FINAL

**Table 15.2.4.26 Descriptive Statistics of Urinary B[a]P - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Quantity Excreted over 24 hours (pg)	Day 3	n	39	39
		BLOQ - n (%)	13 (33.3%)	
		Geometric Mean (CV%)	70.35 (81.55%)	
		Geometric 95% CI	55.81, 88.67	
		Median	72.54	-74.22
		Q25, Q75	41.58, 113.88	-82.28, -58.21
		Min, Max	13.1, 368.3	-93.4, -6.6
		Mean (SD)	89.69 (68.543)	-68.50 (19.362)
		95% CI		-74.78, -62.22

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (25.0 fg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_bap.sas

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Program Run: 05NOV15 cvn_kbooth Program Status: FINAL

**Table 15.2.4.26 Descriptive Statistics of Urinary B[a]P - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Quantity Excreted over 24 hours (pg)	Day 4	n	39	39
		BLOQ - n (%)	25 (64.1%)	
		Geometric Mean (CV%)	48.49 (70.19%)	
		Geometric 95% CI	39.49, 59.53	
		Median	42.50	-81.58
		Q25, Q75	30.00, 75.94	-89.07, -69.65
		Min, Max	20.0, 291.6	-92.9, -52.5
		Mean (SD)	61.02 (52.672)	-78.79 (12.329)
		95% CI		-82.79, -74.79

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (25.0 fg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_bap.sas

Program Run: 05NOV15 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.26 Descriptive Statistics of Urinary B[a]P - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Quantity Excreted over 24 hours (pg)	Day 5	n	39	39
		BLOQ - n (%)	20 (51.3%)	
		Geometric Mean (CV%)	52.77 (65.11%)	
		Geometric 95% CI	43.52, 63.99	
		Median	46.25	-80.58
		Q25, Q75	35.00, 81.00	-87.47, -67.76
		Min, Max	19.1, 198.7	-93.7, -23.7
		Mean (SD)	63.57 (43.756)	-76.80 (14.238)
		95% CI		-81.42, -72.18

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (25.0 fg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_bap.sas

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Program Run: 05NOV15 cvn_kbooth Program Status: FINAL

**Table 15.2.4.26 Descriptive Statistics of Urinary B[a]P - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Concentration Adjusted for Creatinine (fg/mg creat)	Baseline	n	39	
		BLOQ - n (%)	0	
		Geometric Mean (CV%)	169.58 (46.20%)	
		Geometric 95% CI	147.04, 195.57	
		Median	168.82	
		Q25, Q75	133.33, 214.15	
		Min, Max	63.2, 451.5	
		Mean (SD)	186.43 (86.526)	
		95% CI		

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (25.0 fg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_bap.sas

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Program Run: 05NOV15 cvn_kbooth Program Status: FINAL

**Table 15.2.4.26 Descriptive Statistics of Urinary B[a]P - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Concentration Adjusted for Creatinine (fg/mg creat)	Day 1	n	39	39
		BLOQ - n (%)	3 (7.7%)	
		Geometric Mean (CV%)	78.32 (64.13%)	
		Geometric 95% CI	64.75, 94.73	
		Median	69.56	-54.50
		Q25, Q75	54.01, 97.45	-63.40, -43.70
		Min, Max	25.4, 526.5	-85.4, 375.0
		Mean (SD)	96.32 (86.367)	-43.25 (70.215)
		95% CI		-66.02, -20.48

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (25.0 fg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_bap.sas

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Program Run: 05NOV15 cvn_kbooth Program Status: FINAL

**Table 15.2.4.26 Descriptive Statistics of Urinary B[a]P - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Concentration Adjusted for Creatinine (fg/mg creat)	Day 2	n	39	39
		BLOQ - n (%)	15 (38.5%)	
		Geometric Mean (CV%)	43.09 (63.33%)	
		Geometric 95% CI	35.69, 52.02	
		Median	44.21	-71.16
		Q25, Q75	26.65, 74.65	-82.08, -58.58
		Min, Max	14.3, 137.3	-93.4, -23.7
		Mean (SD)	50.41 (28.357)	-70.09 (16.018)
		95% CI		-75.28, -64.89

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (25.0 fg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_bap.sas

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Program Run: 05NOV15 cvn_kbooth Program Status: FINAL

**Table 15.2.4.26 Descriptive Statistics of Urinary B[a]P - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Concentration Adjusted for Creatinine (fg/mg creat)	Day 3	n	39	39
		BLOQ - n (%)	13 (33.3%)	
		Geometric Mean (CV%)	45.85 (58.90%)	
		Geometric 95% CI	38.41, 54.73	
		Median	44.80	-71.35
		Q25, Q75	31.19, 64.36	-81.08, -55.28
		Min, Max	12.4, 146.5	-94.9, -27.5
		Mean (SD)	53.19 (31.615)	-68.02 (17.445)
		95% CI		-73.68, -62.36

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (25.0 fg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_bap.sas

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Program Run: 05NOV15 cvn_kbooth Program Status: FINAL

**Table 15.2.4.26 Descriptive Statistics of Urinary B[a]P - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Concentration Adjusted for Creatinine (fg/mg creat)	Day 4	n	39	39
		BLOQ - n (%)	25 (64.1%)	
		Geometric Mean (CV%)	32.95 (56.32%)	
		Geometric 95% CI	27.79, 39.07	
		Median	31.35	-80.23
		Q25, Q75	22.08, 47.35	-88.11, -70.48
		Min, Max	14.7, 114.6	-93.1, -37.8
		Mean (SD)	38.01 (22.736)	-76.96 (14.254)
		95% CI		-81.59, -72.34

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (25.0 fg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_bap.sas

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Program Run: 05NOV15 cvn_kbooth Program Status: FINAL

**Table 15.2.4.26 Descriptive Statistics of Urinary B[a]P - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Concentration Adjusted for Creatinine (fg/mg creat)	Day 5	n	39	39
		BLOQ - n (%)	20 (51.3%)	
		Geometric Mean (CV%)	33.64 (50.27%)	
		Geometric 95% CI	28.84, 39.24	
		Median	33.88	-79.48
		Q25, Q75	22.40, 48.73	-87.92, -66.95
		Min, Max	15.5, 90.3	-93.6, -45.5
		Mean (SD)	37.66 (19.055)	-77.24 (12.023)
		95% CI		-81.14, -73.34

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (25.0 fg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_bap.sas

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Program Run: 05NOV15 cvn_kbooth Program Status: FINAL

**Table 15.2.4.27 Analysis of Urinary HMPMA on Day 5 - FAS**

Variable	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	THS : CC Ratio (%)	THS : SA Ratio (%)
Urinary Quantity over n 24 hours (µg)		78	41	39		
	Geometric LS Mean (SE) ¹	122.82	562.07	102.12	21.85 (0.076)	120.28 (0.077)
	95% CI	112.55, 134.03	498.42, 633.86	90.21, 115.59	18.83, 25.36	103.31, 140.02
Concentration Adjusted for Creatinine (ng/mg creat)	n	78	41	39		
	Geometric LS Mean (SE) ¹	85.52	379.36	64.57	22.54 (0.059)	132.46 (0.060)
	95% CI	79.95, 91.49	345.74, 416.25	58.69, 71.03	20.10, 25.29	117.83, 148.90

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Adjusted geometric least squares (LS) means and confidence intervals (CIs) from an ANCOVA model conducted on log-transformed Day 5 values with log-transformed baseline value, study arm, sex and CC consumption reported at screening as fixed effect factors.

Baseline is defined as the last assessment prior to Day 1 product use for THS 2.2 and CC subjects and prior to 06:29 AM on Day 1 for SA subjects.

Note: 1 = Standard errors presented from analysis performed on the log_e scale.

Note: Collection over 24 hours starting on the Day stated in the title.

Appendix 15.4.4.27 and 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/tl_anlsecondbio.sas

Program Run: 27OCT14 cvn_ahedge Program Status: FINAL

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**Table 15.2.4.28 Descriptive Statistics of Urinary HMPMA - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Quantity Excreted over 24 hours (µg)	Baseline	n	80		41	
		BLOQ - n (%)	0		0	
		Geometric Mean (CV%)	768.61 (44.60%)		758.87 (39.04%)	
		Geometric 95% CI	699.10, 845.02		673.82, 854.67	
		Median	788.75		690.00	
		Q25, Q75	643.27, 994.85		607.20, 892.70	
		Min, Max	198.5, 1876.8		420.5, 2348.2	
		Mean (SD)	833.84 (326.122)		819.68 (375.701)	
		95% CI				

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (20.0 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hmpma.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.28 Descriptive Statistics of Urinary HMPMA - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Quantity Excreted over 24 hours (µg)	Day 1	n	80	80	41	41
		BLOQ - n (%)	0		0	
		Geometric Mean (CV%)	226.85 (38.56%)		641.90 (51.79%)	
		Geometric 95% CI	208.81, 246.45		550.36, 748.66	
		Median	243.68	-71.94	620.40	-14.50
		Q25, Q75	175.04, 281.50	-77.87, -59.69	458.85, 952.30	-38.80, 4.04
		Min, Max	61.3, 620.5	-89.3, -25.5	235.5, 1853.1	-54.8, 85.0
		Mean (SD)	242.05 (87.774)	-67.97 (13.405)	722.57 (376.863)	-10.74 (30.504)
		95% CI		-70.96, -64.98		-20.37, -1.10

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (20.0 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hmpma.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.28 Descriptive Statistics of Urinary HMPMA - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Quantity Excreted over 24 hours (µg)	Day 2	n	80	80	41	41
		BLOQ - n (%)	1 (1.3%)		0	
		Geometric Mean (CV%)	316.66 (58.75%)		754.00 (47.71%)	
		Geometric 95% CI	280.52, 357.46		653.57, 869.86	
		Median	342.46	-57.16	720.00	1.32
		Q25, Q75	250.02, 437.25	-67.64, -45.88	526.50, 996.00	-18.22, 19.20
		Min, Max	29.3, 1526.0	-85.7, 100.9	280.0, 1755.6	-61.6, 136.3
		Mean (SD)	359.54 (190.164)	-54.01 (25.021)	834.47 (394.196)	6.31 (41.383)
		95% CI		-59.58, -48.43		-6.76, 19.38

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (20.0 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hmpma.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.28 Descriptive Statistics of Urinary HMPMA - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Quantity Excreted over 24 hours (µg)	Day 3	n	78	78	41	41
		BLOQ - n (%)	0		0	
		Geometric Mean (CV%)	226.83 (41.05%)		632.40 (54.54%)	
		Geometric 95% CI	207.52, 247.94		538.32, 742.92	
		Median	224.20	-69.07	642.40	-19.76
		Q25, Q75	183.60, 290.40	-78.98, -61.77	456.00, 839.55	-34.66, 11.95
		Min, Max	72.4, 612.0	-91.9, -31.3	114.0, 1640.9	-75.0, 58.7
		Mean (SD)	244.45 (97.158)	-68.02 (12.963)	712.27 (357.112)	-11.59 (30.226)
		95% CI		-70.95, -65.10		-21.14, -2.05

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (20.0 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hmpma.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.28 Descriptive Statistics of Urinary HMPMA - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Quantity Excreted over 24 hours (µg)	Day 4	n	79	79	41	41
		BLOQ - n (%)	0		0	
		Geometric Mean (CV%)	144.87 (34.60%)		570.19 (56.73%)	
		Geometric 95% CI	134.35, 156.20		482.62, 673.64	
		Median	137.20	-82.29	527.85	-23.00
		Q25, Q75	116.55, 177.48	-86.60, -76.99	416.67, 708.90	-44.59, -7.42
		Min, Max	75.9, 543.4	-92.5, -26.4	214.5, 2049.6	-58.9, 127.1
		Mean (SD)	154.30 (66.006)	-78.88 (12.735)	660.92 (412.374)	-20.04 (32.396)
		95% CI		-81.74, -76.02		-30.27, -9.81

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (20.0 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hmpma.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.28 Descriptive Statistics of Urinary HMPMA - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Quantity Excreted over 24 hours (µg)	Day 5	n	78	78	41	41
		BLOQ - n (%)	6 (7.7%)		0	
		Geometric Mean (CV%)	124.63 (51.90%)		565.85 (58.14%)	
		Geometric 95% CI	111.63, 139.14		477.24, 670.92	
		Median	133.04	-83.42	559.00	-27.40
		Q25, Q75	95.40, 171.36	-86.87, -79.00	393.00, 802.50	-40.31, -9.36
		Min, Max	23.0, 300.7	-95.0, -48.3	140.6, 1567.5	-69.9, 91.2
		Mean (SD)	137.91 (57.799)	-82.40 (7.878)	650.15 (354.000)	-20.24 (30.922)
		95% CI		-84.19, -80.62		-30.01, -10.48

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (20.0 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hmpma.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.28 Descriptive Statistics of Urinary HMPMA - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Concentration Adjusted for Creatinine (ng/mg creat)	Baseline	n	80		41	
		BLOQ - n (%)	0		0	
		Geometric Mean (CV%)	479.34 (45.31%)		460.52 (40.35%)	
		Geometric 95% CI	435.40, 527.72		407.39, 520.59	
		Median	481.07		452.63	
		Q25, Q75	369.88, 661.80		356.80, 617.25	
		Min, Max	147.2, 1364.1		197.9, 1326.7	
		Mean (SD)	522.92 (217.708)		496.28 (206.475)	
		95% CI				

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (20.0 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hmpma.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.28 Descriptive Statistics of Urinary HMPMA - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Concentration Adjusted for Creatinine (ng/mg creat)	Day 1	n	80	80	41	41
		BLOQ - n (%)	0		0	
		Geometric Mean (CV%)	157.71 (32.39%)		440.24 (42.85%)	
		Geometric 95% CI	147.00, 169.20		386.73, 501.16	
		Median	159.87	-68.51	433.52	-4.49
		Q25, Q75	129.92, 202.72	-74.69, -57.15	341.26, 582.14	-22.30, 24.05
		Min, Max	60.3, 286.7	-85.8, -14.3	201.9, 1041.1	-50.7, 69.4
		Mean (SD)	165.29 (49.508)	-64.51 (14.712)	477.12 (194.409)	-0.26 (29.085)
		95% CI		-67.79, -61.23		-9.44, 8.93

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (20.0 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hmpma.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.28 Descriptive Statistics of Urinary HMPMA - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Concentration Adjusted for Creatinine (ng/mg creat)	Day 2	n	80	80	41	41
		BLOQ - n (%)	1 (1.3%)		0	
		Geometric Mean (CV%)	211.73 (41.85%)		472.76 (36.37%)	
		Geometric 95% CI	193.62, 231.54		422.99, 528.39	
		Median	218.85	-55.50	477.55	2.42
		Q25, Q75	181.75, 271.42	-63.12, -43.67	369.04, 563.67	-17.12, 22.37
		Min, Max	33.2, 450.4	-84.9, 32.1	180.9, 937.4	-46.6, 116.4
		Mean (SD)	226.39 (75.895)	-51.94 (20.323)	501.43 (173.290)	7.36 (33.736)
		95% CI	-56.47, -47.41		-3.29, 18.01	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (20.0 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hmpma.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.4.28 Descriptive Statistics of Urinary HMPMA - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Concentration Adjusted for Creatinine (ng/mg creat)	Day 3	n	78	78	41	41
		BLOQ - n (%)	0		0	
		Geometric Mean (CV%)	153.34 (32.21%)		408.40 (47.31%)	
		Geometric 95% CI	142.85, 164.60		354.39, 470.64	
		Median	154.57	-66.43	401.49	-12.55
		Q25, Q75	122.70, 181.93	-75.49, -58.57	306.66, 531.92	-27.06, 11.22
		Min, Max	76.7, 482.3	-91.3, -20.0	121.5, 998.2	-69.6, 71.7
		Mean (SD)	161.37 (57.313)	-65.50 (13.827)	448.71 (195.609)	-6.45 (31.054)
		95% CI		-68.62, -62.38		-16.25, 3.36

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (20.0 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hmpma.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.28 Descriptive Statistics of Urinary HMPMA - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Concentration Adjusted for Creatinine (ng/mg creat)	Day 4	n	79	79	41	41
		BLOQ - n (%)	0		0	
		Geometric Mean (CV%)	99.17 (36.17%)		373.94 (46.78%)	
		Geometric 95% CI	91.68, 107.28		324.96, 430.31	
		Median	100.86	-80.91	384.30	-21.69
		Q25, Q75	77.62, 123.04	-85.25, -74.11	273.58, 515.16	-35.80, 1.87
		Min, Max	48.7, 345.0	-91.2, 17.6	127.0, 815.3	-54.1, 51.4
		Mean (SD)	105.82 (43.290)	-76.49 (16.537)	410.03 (174.256)	-15.26 (25.718)
		95% CI		-80.20, -72.78		-23.38, -7.13

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (20.0 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hmpma.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.28 Descriptive Statistics of Urinary HMPMA - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Concentration Adjusted for Creatinine (ng/mg creat)	Day 5	n	78	78	41	41
		BLOQ - n (%)	6 (7.7%)		0	
		Geometric Mean (CV%)	86.65 (34.65%)		376.78 (44.43%)	
		Geometric 95% CI	80.31, 93.49		329.54, 430.80	
		Median	90.74	-82.66	391.40	-16.72
		Q25, Q75	70.11, 112.40	-86.27, -76.04	272.92, 512.96	-29.76, 4.25
		Min, Max	36.8, 164.4	-93.3, -38.1	158.5, 712.1	-62.6, 43.8
		Mean (SD)	91.37 (28.857)	-80.58 (8.423)	409.52 (163.281)	-14.53 (25.193)
		95% CI			-82.48, -78.68	
					-22.49, -6.57	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (20.0 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hmpma.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.28 Descriptive Statistics of Urinary HMPMA - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Quantity Excreted over 24 hours (µg)	Baseline	n	39	
		BLOQ - n (%)	0	
		Geometric Mean (CV%)	701.16 (51.42%)	
		Geometric 95% CI	599.27, 820.36	
		Median	702.00	
		Q25, Q75	473.96, 945.40	
		Min, Max	275.2, 2013.1	
		Mean (SD)	786.03 (393.566)	
		95% CI		

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (20.0 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hmpma.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.28 Descriptive Statistics of Urinary HMPMA - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Quantity Excreted over 24 hours (µg)	Day 1	n	39	39
		BLOQ - n (%)	0	
		Geometric Mean (CV%)	219.05 (32.54%)	
		Geometric 95% CI	197.64, 242.77	
		Median	221.00	-72.45
		Q25, Q75	171.10, 278.00	-77.27, -57.24
		Min, Max	108.1, 426.0	-86.6, -6.7
		Mean (SD)	229.88 (72.130)	-64.87 (18.898)
		95% CI		-71.00, -58.74

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (20.0 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hmpma.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.28 Descriptive Statistics of Urinary HMPMA - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Quantity Excreted over 24 hours (µg)	Day 2	n	39	39
		BLOQ - n (%)	0	
		Geometric Mean (CV%)	304.61 (50.31%)	
		Geometric 95% CI	261.14, 355.32	
		Median	335.80	-53.80
		Q25, Q75	229.77, 456.50	-72.83, -31.71
		Min, Max	79.7, 591.7	-93.0, 56.3
		Mean (SD)	335.45 (135.923)	-47.38 (32.431)
		95% CI		-57.90, -36.87

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (20.0 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hmpma.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.28 Descriptive Statistics of Urinary HMPMA - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Quantity Excreted over 24 hours (µg)	Day 3	n	39	39
		BLOQ - n (%)	0	
		Geometric Mean (CV%)	204.56 (47.39%)	
		Geometric 95% CI	176.78, 236.69	
		Median	207.70	-70.58
		Q25, Q75	148.40, 286.35	-80.91, -61.59
		Min, Max	82.8, 511.5	-87.9, -3.3
		Mean (SD)	225.85 (105.244)	-66.24 (19.895)
		95% CI		-72.70, -59.79

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (20.0 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hmpma.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.28 Descriptive Statistics of Urinary HMPMA - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Quantity Excreted over 24 hours (µg)	Day 4	n	39	39
		BLOQ - n (%)	4 (10.3%)	
		Geometric Mean (CV%)	95.80 (45.72%)	
		Geometric 95% CI	83.18, 110.34	
		Median	106.40	-86.99
		Q25, Q75	73.16, 132.09	-89.34, -81.72
		Min, Max	34.0, 245.1	-96.9, -66.1
		Mean (SD)	104.65 (44.452)	-84.76 (7.226)
		95% CI		-87.10, -82.41

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (20.0 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hmpma.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.28 Descriptive Statistics of Urinary HMPMA - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Quantity Excreted over 24 hours (µg)	Day 5	n	39	39
		BLOQ - n (%)	2 (5.1%)	
		Geometric Mean (CV%)	99.22 (38.68%)	
		Geometric 95% CI	87.90, 111.99	
		Median	95.76	-84.47
		Q25, Q75	85.12, 133.20	-90.20, -81.31
		Min, Max	26.0, 180.4	-95.7, -58.4
		Mean (SD)	105.17 (32.870)	-83.96 (8.301)
		95% CI		-86.66, -81.27

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (20.0 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hmpma.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.28 Descriptive Statistics of Urinary HMPMA - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Concentration Adjusted for Creatinine (ng/mg creat)	Baseline	n	39	
		BLOQ - n (%)	0	
		Geometric Mean (CV%)	443.71 (48.75%)	
		Geometric 95% CI	382.03, 515.35	
		Median	471.14	
		Q25, Q75	314.81, 616.53	
		Min, Max	161.7, 1266.9	
		Mean (SD)	490.00 (221.983)	
		95% CI		

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (20.0 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hmpma.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.28 Descriptive Statistics of Urinary HMPMA - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Concentration Adjusted for Creatinine (ng/mg creat)	Day 1	n	39	39
		BLOQ - n (%)	0	
		Geometric Mean (CV%)	147.45 (27.74%)	
		Geometric 95% CI	134.99, 161.06	
		Median	144.34	-69.04
		Q25, Q75	119.67, 180.69	-73.27, -52.81
		Min, Max	89.1, 289.2	-87.2, -12.8
		Mean (SD)	153.02 (43.950)	-63.05 (18.105)
		95% CI		-68.92, -57.17

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (20.0 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hmpma.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.28 Descriptive Statistics of Urinary HMPMA - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Concentration Adjusted for Creatinine (ng/mg creat)	Day 2	n	39	39
		BLOQ - n (%)	0	
		Geometric Mean (CV%)	198.01 (39.91%)	
		Geometric 95% CI	174.80, 224.29	
		Median	207.46	-47.41
		Q25, Q75	166.09, 257.55	-73.86, -25.06
		Min, Max	64.4, 373.6	-86.7, 43.4
		Mean (SD)	211.07 (70.157)	-47.41 (28.843)
		95% CI		-56.77, -38.06

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (20.0 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hmpma.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.28 Descriptive Statistics of Urinary HMPMA - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Concentration Adjusted for Creatinine (ng/mg creat)	Day 3	n	39	39
		BLOQ - n (%)	0	
		Geometric Mean (CV%)	133.34 (38.64%)	
		Geometric 95% CI	118.15, 150.48	
		Median	124.09	-69.80
		Q25, Q75	107.70, 159.22	-79.11, -57.57
		Min, Max	57.6, 325.1	-88.5, 2.9
		Mean (SD)	143.15 (58.807)	-65.85 (18.834)
		95% CI		-71.96, -59.74

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (20.0 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hmpma.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.28 Descriptive Statistics of Urinary HMPMA - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Concentration Adjusted for Creatinine (ng/mg creat)	Day 4	n	39	39
		BLOQ - n (%)	4 (10.3%)	
		Geometric Mean (CV%)	65.11 (40.67%)	
		Geometric 95% CI	57.35, 73.92	
		Median	64.10	-85.17
		Q25, Q75	52.35, 87.76	-88.39, -80.54
		Min, Max	27.8, 159.7	-95.2, -64.9
		Mean (SD)	70.13 (28.154)	-83.80 (7.384)
		95% CI		-86.19, -81.40

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (20.0 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hmpma.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.28 Descriptive Statistics of Urinary HMPMA - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Concentration Adjusted for Creatinine (ng/mg creat)	Day 5	n	39	39
		BLOQ - n (%)	2 (5.1%)	
		Geometric Mean (CV%)	63.25 (28.36%)	
		Geometric 95% CI	57.79, 69.22	
		Median	63.76	-84.98
		Q25, Q75	54.31, 79.62	-88.81, -81.50
		Min, Max	30.3, 108.7	-95.8, -65.6
		Mean (SD)	65.58 (17.521)	-84.39 (6.744)
		95% CI		-86.58, -82.20

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (20.0 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hmpma.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.29 Analysis of Urinary S-BMA on Day 5 - FAS**

Variable	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	THS : CC Ratio (%)	THS : SA Ratio (%)
Urinary Quantity over n 24 hours (ng)		79	41	39		
	Geometric LS Mean (SE) ¹	11888.39	10970.22	13389.43	108.37 (0.113)	88.79 (0.115)
	95% CI	10441.09, 13536.29	9161.62, 13135.86	11130.81, 16106.37	86.78, 135.33	70.83, 111.30
Concentration Adjusted for Creatinine (pg/mg creat)	n	79	41	39		
	Geometric LS Mean (SE) ¹	8241.59	7462.65	8449.62	110.44 (0.099)	97.54 (0.101)
	95% CI	7355.01, 9235.03	6371.75, 8740.31	7186.58, 9934.65	90.88, 134.20	80.02, 118.89

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Adjusted geometric least squares (LS) means and confidence intervals (CIs) from an ANCOVA model conducted on log-transformed Day 5 values with log-transformed baseline value, study arm, sex and CC consumption reported at screening as fixed effect factors.

Baseline is defined as the last assessment prior to Day 1 product use for THS 2.2 and CC subjects and prior to 06:29 AM on Day 1 for SA subjects.

Note: 1 = Standard errors presented from analysis performed on the log_e scale.

Note: Collection over 24 hours starting on the Day stated in the title.

Appendix 15.4.4.29 and 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/tl_anlsecondbio.sas

Program Run: 27OCT14 cvn_ahedge Program Status: FINAL

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**Table 15.2.4.30 Descriptive Statistics of Urinary S-BMA - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)	
			Raw value	%Change (*)
Quantity Excreted over 24 hours (ng)	Baseline	n	80	
		BLOQ - n (%)	0	
		Geometric Mean (CV%)	11892.33 (44.84%)	
		Geometric 95% CI	10811.84, 13080.80	
		Median	11751.00	
		Q25, Q75	9171.00, 16308.50	
		Min, Max	4152.0, 32718.0	
		Mean (SD)	12981.54 (5486.017)	
		95% CI		

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_bma.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.30 Descriptive Statistics of Urinary S-BMA - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)	
			Raw value	%Change (*)
Quantity Excreted over 24 hours (ng)	Day 1	n	80	80
		BLOQ - n (%)	0	
		Geometric Mean (CV%)	8820.59 (47.46%)	
		Geometric 95% CI	7978.81, 9751.18	
		Median	9385.00	-26.36
		Q25, Q75	6564.25, 12161.25	-40.13, -16.73
		Min, Max	2360.0, 29120.0	-62.1, 115.0
		Mean (SD)	9712.42 (4361.599)	-22.16 (27.865)
		95% CI		-28.36, -15.95

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_bma.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.30 Descriptive Statistics of Urinary S-BMA - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)	
			Raw value	%Change (*)
Quantity Excreted over 24 hours (ng)	Day 2	n	80	80
		BLOQ - n (%)	0	
		Geometric Mean (CV%)	7467.93 (54.12%)	
		Geometric 95% CI	6671.44, 8359.50	
		Median	7843.75	-40.56
		Q25, Q75	5522.00, 9870.75	-50.22, -27.80
		Min, Max	2373.0, 51520.0	-73.6, 186.9
		Mean (SD)	8595.54 (6048.143)	-31.82 (34.775)
		95% CI		-39.56, -24.07

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_bma.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.30 Descriptive Statistics of Urinary S-BMA - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)	
			Raw value	%Change (*)
Quantity Excreted over 24 hours (ng)	Day 3	n	79	79
		BLOQ - n (%)	0	
		Geometric Mean (CV%)	8878.39 (55.29%)	
		Geometric 95% CI	7908.64, 9967.06	
		Median	9135.00	-28.40
		Q25, Q75	6025.00, 12531.50	-41.19, -10.31
		Min, Max	2442.0, 25173.8	-63.4, 157.8
		Mean (SD)	10102.08 (5306.904)	-19.95 (35.951)
		95% CI		-28.00, -11.89

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_bma.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.30 Descriptive Statistics of Urinary S-BMA - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)	
			Raw value	%Change (*)
Quantity Excreted over 24 hours (ng)	Day 4	n	79	79
		BLOQ - n (%)	0	
		Geometric Mean (CV%)	8901.81 (51.40%)	
		Geometric 95% CI	7987.02, 9921.38	
		Median	9152.00	-30.99
		Q25, Q75	6042.00, 12048.75	-41.63, -13.13
		Min, Max	2314.0, 32640.0	-53.5, 232.6
		Mean (SD)	10002.20 (5257.837)	-18.93 (46.193)
		95% CI		-29.28, -8.58

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_bma.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.30 Descriptive Statistics of Urinary S-BMA - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)	
			Raw value	%Change (*)
Quantity Excreted over 24 hours (ng)	Day 5	n	79	79
		BLOQ - n (%)	0	
		Geometric Mean (CV%)	12052.57 (88.00%)	
		Geometric 95% CI	10172.28, 14280.43	
		Median	12638.00	-0.49
		Q25, Q75	7422.80, 21148.50	-35.45, 49.48
		Min, Max	956.8, 59850.0	-84.7, 436.5
		Mean (SD)	15556.63 (11227.495)	24.49 (93.398)
		95% CI		3.56, 45.41

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_bma.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.30 Descriptive Statistics of Urinary S-BMA - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)	
			Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Baseline	n	80	
		BLOQ - n (%)	0	
		Geometric Mean (CV%)	7416.70 (50.20%)	
		Geometric 95% CI	6674.19, 8241.82	
		Median	7193.66	
		Q25, Q75	5342.03, 9481.21	
		Min, Max	2467.4, 25420.9	
		Mean (SD)	8322.81 (4399.338)	
		95% CI		

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_bma.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.30 Descriptive Statistics of Urinary S-BMA - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)	
			Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Day 1	n	80	80
		BLOQ - n (%)	0	
		Geometric Mean (CV%)	6132.23 (49.56%)	
		Geometric 95% CI	5524.98, 6806.21	
		Median	6026.10	-18.99
		Q25, Q75	4844.70, 7905.07	-29.45, -4.03
		Min, Max	2099.6, 26563.7	-55.3, 159.6
		Mean (SD)	6871.97 (3749.192)	-14.07 (27.962)
		95% CI		-20.30, -7.84

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_bma.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.30 Descriptive Statistics of Urinary S-BMA - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)	
			Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Day 2	n	80	80
		BLOQ - n (%)	0	
		Geometric Mean (CV%)	4993.38 (51.32%)	
		Geometric 95% CI	4483.98, 5560.64	
		Median	4807.79	-35.09
		Q25, Q75	3790.21, 6428.34	-44.00, -21.01
		Min, Max	1611.8, 17366.7	-65.4, 88.6
		Mean (SD)	5624.83 (3015.160)	-28.89 (27.312)
		95% CI		-34.97, -22.80

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_bma.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.30 Descriptive Statistics of Urinary S-BMA - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)	
			Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Day 3	n	79	79
		BLOQ - n (%)	0	
		Geometric Mean (CV%)	6004.90 (53.46%)	
		Geometric 95% CI	5367.07, 6718.53	
		Median	5954.95	-24.77
		Q25, Q75	4625.19, 8220.07	-36.27, -1.39
		Min, Max	1776.0, 17032.3	-62.3, 159.8
		Mean (SD)	6779.32 (3448.167)	-13.73 (35.910)
		95% CI		-21.78, -5.68

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_bma.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.30 Descriptive Statistics of Urinary S-BMA - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)	
			Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Day 4	n	79	79
		BLOQ - n (%)	0	
		Geometric Mean (CV%)	6093.87 (52.28%)	
		Geometric 95% CI	5458.61, 6803.07	
		Median	5981.31	-24.31
		Q25, Q75	4415.58, 8428.36	-35.19, -7.99
		Min, Max	1846.5, 21062.8	-53.4, 319.0
		Mean (SD)	6869.49 (3559.555)	-10.08 (56.802)
		95% CI		-22.81, 2.65

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_bma.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.30 Descriptive Statistics of Urinary S-BMA - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)	
			Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Day 5	n	79	79
		BLOQ - n (%)	0	
		Geometric Mean (CV%)	8394.71 (74.08%)	
		Geometric 95% CI	7238.83, 9735.15	
		Median	8446.77	6.70
		Q25, Q75	5235.23, 15150.95	-26.63, 48.32
		Min, Max	1598.7, 27536.2	-54.1, 384.1
		Mean (SD)	10249.14 (6310.555)	32.22 (86.063)
		95% CI		12.93, 51.50

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_bma.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.30 Descriptive Statistics of Urinary S-BMA - FAS**

Variable	Timepoint	Statistic	CC (N=41)	
			Raw value	%Change (*)
Quantity Excreted over 24 hours (ng)	Baseline	n	41	
		BLOQ - n (%)	0	
		Geometric Mean (CV%)	11554.68 (44.21%)	
		Geometric 95% CI	10112.16, 13202.98	
		Median	11224.00	
		Q25, Q75	8624.00, 14229.00	
		Min, Max	4978.0, 26860.0	
		Mean (SD)	12612.78 (5535.402)	
		95% CI		

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_bma.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.30 Descriptive Statistics of Urinary S-BMA - FAS**

Variable	Timepoint	Statistic	CC (N=41)	
			Raw value	%Change (*)
Quantity Excreted over 24 hours (ng)	Day 1	n	41	41
		BLOQ - n (%)	0	
		Geometric Mean (CV%)	8577.21 (44.07%)	
		Geometric 95% CI	7509.36, 9796.90	
		Median	8100.00	-27.16
		Q25, Q75	6372.00, 10780.00	-38.33, -15.33
		Min, Max	4313.0, 22496.8	-54.7, 42.8
		Mean (SD)	9424.06 (4594.400)	-23.09 (21.929)
		95% CI		-30.02, -16.17

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_bma.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.30 Descriptive Statistics of Urinary S-BMA - FAS**

Variable	Timepoint	Statistic	CC (N=41)	
			Raw value	%Change (*)
Quantity Excreted over 24 hours (ng)	Day 2	n	41	41
		BLOQ - n (%)	0	
		Geometric Mean (CV%)	7561.97 (53.56%)	
		Geometric 95% CI	6453.48, 8860.84	
		Median	8112.00	-36.30
		Q25, Q75	5217.00, 9462.00	-43.65, -24.33
		Min, Max	2697.5, 20575.0	-68.1, 70.9
		Mean (SD)	8564.43 (4539.697)	-30.83 (25.730)
		95% CI		-38.96, -22.71

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_bma.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.30 Descriptive Statistics of Urinary S-BMA - FAS**

Variable	Timepoint	Statistic	CC (N=41)	
			Raw value	%Change (*)
Quantity Excreted over 24 hours (ng)	Day 3	n	41	41
		BLOQ - n (%)	0	
		Geometric Mean (CV%)	8894.49 (68.08%)	
		Geometric 95% CI	7320.37, 10807.10	
		Median	9506.00	-26.50
		Q25, Q75	5830.50, 10736.00	-37.91, -7.14
		Min, Max	2400.0, 57960.0	-72.8, 157.4
		Mean (SD)	10945.36 (9186.877)	-14.91 (43.771)
		95% CI		-28.73, -1.09

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_bma.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.30 Descriptive Statistics of Urinary S-BMA - FAS**

Variable	Timepoint	Statistic	CC (N=41)	
			Raw value	%Change (*)
Quantity Excreted over 24 hours (ng)	Day 4	n	41	41
		BLOQ - n (%)	0	
		Geometric Mean (CV%)	8660.82 (68.88%)	
		Geometric 95% CI	7114.55, 10543.15	
		Median	8109.00	-27.47
		Q25, Q75	5460.00, 12441.00	-44.62, -17.92
		Min, Max	3425.0, 100005.0	-62.4, 344.2
		Mean (SD)	11372.40 (14888.584)	-14.99 (63.888)
		95% CI		-35.16, 5.18

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_bma.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.30 Descriptive Statistics of Urinary S-BMA - FAS**

Variable	Timepoint	Statistic	CC (N=41)	
			Raw value	%Change (*)
Quantity Excreted over 24 hours (ng)	Day 5	n	41	41
		BLOQ - n (%)	0	
		Geometric Mean (CV%)	10886.58 (63.93%)	
		Geometric 95% CI	9050.00, 13095.86	
		Median	10109.75	-8.88
		Q25, Q75	6960.00, 15966.00	-37.78, 23.60
		Min, Max	4592.3, 39780.0	-54.0, 471.1
		Mean (SD)	13029.96 (8677.301)	12.88 (95.977)
		95% CI		-17.42, 43.18

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_bma.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.30 Descriptive Statistics of Urinary S-BMA - FAS**

Variable	Timepoint	Statistic	CC (N=41)	
			Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Baseline	n	41	
		BLOQ - n (%)	0	
		Geometric Mean (CV%)	7011.99 (43.94%)	
		Geometric 95% CI	6141.12, 8006.37	
		Median	7153.85	
		Q25, Q75	5182.10, 9485.30	
		Min, Max	2531.2, 15887.4	
		Mean (SD)	7604.13 (2997.427)	
		95% CI		

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_bma.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.30 Descriptive Statistics of Urinary S-BMA - FAS**

Variable	Timepoint	Statistic	CC (N=41)	
			Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Day 1	n	41	41
		BLOQ - n (%)	0	
		Geometric Mean (CV%)	5882.64 (49.38%)	
		Geometric 95% CI	5076.30, 6817.07	
		Median	5802.62	-17.03
		Q25, Q75	4518.41, 7570.10	-28.47, 0.64
		Min, Max	1857.3, 19219.7	-56.0, 44.9
		Mean (SD)	6568.49 (3451.116)	-13.45 (21.689)
		95% CI		-20.30, -6.60

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_bma.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.30 Descriptive Statistics of Urinary S-BMA - FAS**

Variable	Timepoint	Statistic	CC (N=41)	
			Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Day 2	n	41	41
		BLOQ - n (%)	0	
		Geometric Mean (CV%)	4741.37 (55.33%)	
		Geometric 95% CI	4027.78, 5581.37	
		Median	5062.84	-29.76
		Q25, Q75	3875.00, 6480.00	-40.96, -18.67
		Min, Max	1315.1, 14499.1	-70.1, 0.3
		Mean (SD)	5341.88 (2580.299)	-30.28 (16.337)
		95% CI		-35.45, -25.12

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_bma.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.30 Descriptive Statistics of Urinary S-BMA - FAS**

Variable	Timepoint	Statistic	CC (N=41)	
			Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Day 3	n	41	41
		BLOQ - n (%)	0	
		Geometric Mean (CV%)	5744.04 (74.50%)	
		Geometric 95% CI	4657.38, 7084.23	
		Median	5841.68	-21.25
		Q25, Q75	4092.46, 8505.54	-34.32, -4.46
		Min, Max	1343.3, 22500.0	-76.1, 176.3
		Mean (SD)	7036.24 (4688.040)	-9.27 (46.674)
		95% CI		-24.01, 5.47

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_bma.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.30 Descriptive Statistics of Urinary S-BMA - FAS**

Variable	Timepoint	Statistic	CC (N=41)	
			Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Day 4	n	41	41
		BLOQ - n (%)	0	
		Geometric Mean (CV%)	5679.96 (70.82%)	
		Geometric 95% CI	4644.69, 6945.99	
		Median	5372.81	-24.87
		Q25, Q75	3856.50, 7852.26	-33.50, -3.24
		Min, Max	1631.7, 40745.2	-73.8, 400.4
		Mean (SD)	7079.88 (6296.091)	-7.19 (72.168)
		95% CI		-29.97, 15.59

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_bma.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.30 Descriptive Statistics of Urinary S-BMA - FAS**

Variable	Timepoint	Statistic	CC (N=41)	
			Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Day 5	n	41	41
		BLOQ - n (%)	0	
		Geometric Mean (CV%)	7248.99 (63.53%)	
		Geometric 95% CI	6032.10, 8711.37	
		Median	6823.08	-4.22
		Q25, Q75	4711.11, 10532.32	-22.86, 38.69
		Min, Max	2096.8, 23409.1	-64.3, 378.2
		Mean (SD)	8530.27 (5087.066)	18.26 (79.426)
		95% CI		-6.82, 43.33

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_bma.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.30 Descriptive Statistics of Urinary S-BMA - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Quantity Excreted over 24 hours (ng)	Baseline	n	39	
		BLOQ - n (%)	0	
		Geometric Mean (CV%)	11464.44 (67.42%)	
		Geometric 95% CI	9401.22, 13980.45	
		Median	10741.50	
		Q25, Q75	7366.00, 15652.00	
		Min, Max	4050.0, 74967.5	
		Mean (SD)	14301.38 (12845.197)	
		95% CI		

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_bma.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.30 Descriptive Statistics of Urinary S-BMA - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Quantity Excreted over 24 hours (ng)	Day 1	n	39	39
		BLOQ - n (%)	0	
		Geometric Mean (CV%)	9323.77 (66.49%)	
		Geometric 95% CI	7663.29, 11344.04	
		Median	9527.50	-22.76
		Q25, Q75	6048.00, 12824.00	-39.32, -4.16
		Min, Max	2960.0, 47100.0	-53.0, 142.1
		Mean (SD)	11302.62 (8382.930)	-11.69 (41.667)
		95% CI		-25.20, 1.82

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_bma.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.30 Descriptive Statistics of Urinary S-BMA - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Quantity Excreted over 24 hours (ng)	Day 2	n	39	39
		BLOQ - n (%)	0	
		Geometric Mean (CV%)	7958.97 (53.89%)	
		Geometric 95% CI	6757.29, 9374.34	
		Median	7912.50	-35.67
		Q25, Q75	5487.00, 10860.00	-46.85, -12.95
		Min, Max	2646.0, 38125.0	-77.9, 80.0
		Mean (SD)	9105.89 (5849.462)	-23.55 (37.247)
		95% CI		-35.63, -11.47

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_bma.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.30 Descriptive Statistics of Urinary S-BMA - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Quantity Excreted over 24 hours (ng)	Day 3	n	39	39
		BLOQ - n (%)	0	
		Geometric Mean (CV%)	9672.26 (83.06%)	
		Geometric 95% CI	7648.11, 12232.10	
		Median	9838.75	-19.86
		Q25, Q75	5304.00, 14280.00	-46.30, -0.87
		Min, Max	2675.0, 72105.0	-78.3, 415.7
		Mean (SD)	13152.73 (14240.053)	8.23 (104.587)
		95% CI		-25.68, 42.14

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_bma.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.30 Descriptive Statistics of Urinary S-BMA - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Quantity Excreted over 24 hours (ng)	Day 4	n	39	39
		BLOQ - n (%)	0	
		Geometric Mean (CV%)	9885.45 (88.25%)	
		Geometric 95% CI	7729.66, 12642.49	
		Median	9369.00	-28.41
		Q25, Q75	5658.00, 14904.00	-41.82, 3.31
		Min, Max	3472.0, 146280.0	-66.1, 655.6
		Mean (SD)	15209.09 (24810.205)	14.75 (131.459)
		95% CI		-27.87, 57.37

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_bma.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.30 Descriptive Statistics of Urinary S-BMA - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Quantity Excreted over 24 hours (ng)	Day 5	n	39	39
		BLOQ - n (%)	0	
		Geometric Mean (CV%)	13173.34 (72.88%)	
		Geometric 95% CI	10661.44, 16277.04	
		Median	12833.00	7.89
		Q25, Q75	8697.00, 19014.75	-15.85, 42.88
		Min, Max	4181.3, 66780.0	-65.9, 301.7
		Mean (SD)	16463.52 (13064.412)	34.91 (88.309)
		95% CI		6.28, 63.54

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_bma.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.30 Descriptive Statistics of Urinary S-BMA - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Baseline	n	39	
		BLOQ - n (%)	0	
		Geometric Mean (CV%)	7254.96 (61.47%)	
		Geometric 95% CI	6038.57, 8716.38	
		Median	6797.02	
		Q25, Q75	4338.62, 11449.71	
		Min, Max	3049.2, 29339.5	
		Mean (SD)	8549.72 (5466.464)	
		95% CI		

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_bma.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.30 Descriptive Statistics of Urinary S-BMA - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Day 1	n	39	39
		BLOQ - n (%)	0	
		Geometric Mean (CV%)	6276.10 (62.76%)	
		Geometric 95% CI	5206.78, 7565.02	
		Median	5975.61	-17.17
		Q25, Q75	4301.67, 9110.00	-32.13, 2.67
		Min, Max	2287.0, 28846.2	-47.5, 108.8
		Mean (SD)	7494.56 (5354.827)	-8.57 (34.473)
		95% CI		-19.75, 2.61

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_bma.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.30 Descriptive Statistics of Urinary S-BMA - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Day 2	n	39	39
		BLOQ - n (%)	0	
		Geometric Mean (CV%)	5173.53 (54.36%)	
		Geometric 95% CI	4386.96, 6101.13	
		Median	5182.85	-29.09
		Q25, Q75	3456.26, 7122.74	-45.51, -9.44
		Min, Max	2038.2, 19290.1	-58.2, 42.3
		Mean (SD)	5909.26 (3440.135)	-25.12 (24.286)
		95% CI		-33.00, -17.24

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_bma.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.30 Descriptive Statistics of Urinary S-BMA - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Day 3	n	39	39
		BLOQ - n (%)	0	
		Geometric Mean (CV%)	6304.79 (77.06%)	
		Geometric 95% CI	5053.14, 7866.45	
		Median	5963.09	-22.44
		Q25, Q75	4070.45, 9402.40	-38.45, 13.76
		Min, Max	1885.9, 58781.4	-76.9, 325.4
		Mean (SD)	8604.29 (10650.999)	2.87 (75.979)
		95% CI		-21.77, 27.50

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_bma.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.30 Descriptive Statistics of Urinary S-BMA - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Day 4	n	39	39
		BLOQ - n (%)	0	
		Geometric Mean (CV%)	6718.27 (85.49%)	
		Geometric 95% CI	5284.28, 8541.40	
		Median	5930.52	-16.32
		Q25, Q75	4634.14, 8423.33	-36.06, 8.29
		Min, Max	2508.8, 88333.3	-55.9, 408.0
		Mean (SD)	10261.05 (16581.352)	15.39 (105.826)
		95% CI		-18.92, 49.70

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_bma.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.30 Descriptive Statistics of Urinary S-BMA - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Day 5	n	39	39
		BLOQ - n (%)	0	
		Geometric Mean (CV%)	8397.62 (67.56%)	
		Geometric 95% CI	6883.93, 10244.14	
		Median	8680.27	15.45
		Q25, Q75	5062.76, 11158.65	-15.39, 54.78
		Min, Max	2974.5, 47425.5	-58.2, 243.2
		Mean (SD)	10332.89 (8350.523)	30.80 (70.587)
		95% CI		7.92, 53.69

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_bma.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.31 Analysis of Urinary Total NNAL on Day 5 - FAS**

Variable	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	THS : CC Ratio (%)	THS : SA Ratio (%)
Urinary Quantity over n 24 hours (ng)		79	41	39		
	Geometric LS Mean (SE) ¹	71.56	167.25	61.96	42.79 (0.069)	115.49 (0.070)
	95% CI	66.15, 77.41	149.95, 186.53	55.40, 69.30	37.40, 48.95	100.73, 132.41
Concentration Adjusted for Creatinine (pg/mg creat)	n	79	41	39		
	Geometric LS Mean (SE) ¹	49.61	113.93	38.99	43.54 (0.052)	127.25 (0.053)
	95% CI	46.75, 52.64	104.93, 123.71	35.82, 42.42	39.34, 48.19	114.77, 141.08

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Adjusted geometric least squares (LS) means and confidence intervals (CIs) from an ANCOVA model conducted on log-transformed Day 5 values with log-transformed baseline value, study arm, sex and CC consumption reported at screening as fixed effect factors.

Baseline is defined as the last assessment prior to Day 1 product use for THS 2.2 and CC subjects and prior to 06:29 AM on Day 1 for SA subjects.

Note: 1 = Standard errors presented from analysis performed on the log_e scale.

Note: Collection over 24 hours starting on the Day stated in the title.

Appendix 15.4.4.31 and 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/tl_anlsecondbio.sas

Program Run: 27OCT14 cvn_ahedge Program Status: FINAL

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**Table 15.2.4.32 Descriptive Statistics of Urinary Total NNAL - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Quantity Excreted over 24 hours (ng)	Baseline	n	80		41	
		BLOQ - n (%)	0		0	
		Geometric Mean (CV%)	178.00 (75.89%)		173.10 (71.59%)	
		Geometric 95% CI	153.19, 206.83		141.29, 212.07	
		Median	200.16		170.58	
		Q25, Q75	136.37, 272.01		115.44, 289.00	
		Min, Max	11.3, 600.6		43.1, 749.7	
		Mean (SD)	212.82 (115.775)		210.36 (138.636)	
		95% CI				

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (5.00 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_nnal.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.32 Descriptive Statistics of Urinary Total NNAL - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Quantity Excreted over 24 hours (ng)	Day 1	n	80	80	41	41
		BLOQ - n (%)	1 (1.3%)		0	
		Geometric Mean (CV%)	116.54 (79.21%)		159.95 (73.67%)	
		Geometric 95% CI	99.77, 136.13		129.93, 196.91	
		Median	137.84	-34.01	161.54	-5.72
		Q25, Q75	91.10, 173.50	-43.21, -22.91	105.60, 256.20	-18.14, 1.86
		Min, Max	5.6, 473.2	-68.0, 46.6	28.8, 675.7	-33.1, 66.2
		Mean (SD)	140.72 (78.988)	-32.71 (16.473)	194.47 (124.561)	-6.39 (16.323)
		95% CI		-36.38, -29.04		-11.55, -1.23

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (5.00 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_nnal.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.32 Descriptive Statistics of Urinary Total NNAL - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Quantity Excreted over 24 hours (ng)	Day 2	n	80	80	41	41
		BLOQ - n (%)	1 (1.3%)		0	
		Geometric Mean (CV%)	94.25 (81.01%)		163.88 (77.07%)	
		Geometric 95% CI	80.47, 110.40		132.10, 203.30	
		Median	106.88	-46.56	175.80	-2.13
		Q25, Q75	69.13, 150.72	-53.67, -37.64	101.32, 257.61	-15.74, 7.03
		Min, Max	6.1, 351.0	-80.0, 52.5	39.6, 741.0	-71.5, 107.0
		Mean (SD)	114.99 (65.294)	-44.76 (17.442)	202.72 (135.855)	-1.83 (26.605)
		95% CI		-48.65, -40.88		-10.23, 6.57

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (5.00 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_nnal.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.32 Descriptive Statistics of Urinary Total NNAL - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Quantity Excreted over 24 hours (ng)	Day 3	n	79	79	41	41
		BLOQ - n (%)	1 (1.3%)		0	
		Geometric Mean (CV%)	89.25 (82.91%)		157.51 (75.07%)	
		Geometric 95% CI	75.89, 104.95		127.55, 194.52	
		Median	100.80	-49.92	179.79	-3.91
		Q25, Q75	67.32, 139.27	-57.15, -41.65	95.92, 244.20	-19.54, 2.33
		Min, Max	5.3, 348.0	-80.0, 58.8	37.8, 696.2	-58.5, 53.5
		Mean (SD)	109.85 (65.873)	-48.04 (17.354)	193.31 (128.364)	-6.92 (19.460)
		95% CI		-51.93, -44.15		-13.07, -0.77

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (5.00 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_nnal.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.32 Descriptive Statistics of Urinary Total NNAL - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Quantity Excreted over 24 hours (ng)	Day 4	n	79	79	41	41
		BLOQ - n (%)	1 (1.3%)		0	
		Geometric Mean (CV%)	81.59 (75.97%)		159.34 (72.80%)	
		Geometric 95% CI	70.14, 94.92		129.70, 195.76	
		Median	94.33	-53.79	166.32	-5.14
		Q25, Q75	65.75, 122.63	-61.59, -47.43	111.75, 248.90	-16.08, 4.52
		Min, Max	7.3, 300.9	-80.0, 0.2	32.2, 717.6	-56.9, 133.2
		Mean (SD)	98.00 (54.646)	-52.84 (12.914)	193.50 (126.757)	-4.59 (28.480)
		95% CI		-55.73, -49.94		-13.58, 4.40

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (5.00 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_nnal.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.32 Descriptive Statistics of Urinary Total NNAL - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Quantity Excreted over 24 hours (ng)	Day 5	n	79	79	41	41
		BLOQ - n (%)	5 (6.3%)		0	
		Geometric Mean (CV%)	71.29 (94.99%)		160.75 (84.18%)	
		Geometric 95% CI	59.56, 85.32		127.58, 202.53	
		Median	84.55	-56.92	169.05	-10.96
		Q25, Q75	60.27, 119.00	-65.20, -48.33	94.98, 268.81	-16.31, 5.20
		Min, Max	5.8, 264.0	-95.0, -26.9	24.7, 759.5	-42.6, 110.4
		Mean (SD)	89.86 (52.557)	-57.55 (13.177)	203.76 (142.190)	-4.26 (26.195)
		95% CI		-60.51, -54.60		-12.53, 4.01

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (5.00 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_nnal.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.32 Descriptive Statistics of Urinary Total NNAL - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Baseline	n	80		41	
		BLOQ - n (%)	0		0	
		Geometric Mean (CV%)	111.01 (76.55%)		105.05 (80.18%)	
		Geometric 95% CI	95.44, 129.13		84.10, 131.21	
		Median	127.91		118.45	
		Q25, Q75	80.02, 181.38		60.50, 174.53	
		Min, Max	7.7, 302.7		24.2, 323.9	
		Mean (SD)	132.93 (69.766)		130.95 (83.856)	
		95% CI				

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (5.00 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_nnal.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.4.32 Descriptive Statistics of Urinary Total NNAL - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Day 1	n	80	80	41	41
		BLOQ - n (%)	1 (1.3%)		0	
		Geometric Mean (CV%)	81.02 (77.47%)		109.70 (77.58%)	
		Geometric 95% CI	69.55, 94.38		88.33, 136.25	
		Median	90.48	-24.29	115.03	4.04
		Q25, Q75	58.08, 140.74	-33.58, -17.73	70.25, 171.19	-3.19, 14.05
		Min, Max	5.1, 236.4	-59.1, 20.2	23.1, 331.0	-26.7, 34.2
		Mean (SD)	97.35 (51.589)	-25.84 (12.903)	135.32 (85.341)	5.30 (13.534)
		95% CI			-28.72, -22.97	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (5.00 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_nnal.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.32 Descriptive Statistics of Urinary Total NNAL - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Day 2	n	80	80	41	41
		BLOQ - n (%)	1 (1.3%)		0	
		Geometric Mean (CV%)	63.02 (79.50%)		102.75 (79.91%)	
		Geometric 95% CI	53.93, 73.65		82.31, 128.27	
		Median	73.54	-40.74	122.42	0.38
		Q25, Q75	46.89, 109.24	-49.93, -33.58	61.11, 168.53	-11.04, 9.39
		Min, Max	4.9, 201.0	-75.5, 4.9	21.9, 341.2	-49.0, 28.1
		Mean (SD)	76.71 (42.643)	-41.67 (13.392)	128.57 (84.953)	-0.92 (15.176)
		95% CI	-44.65, -38.68		-5.71, 3.88	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (5.00 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_nnal.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.32 Descriptive Statistics of Urinary Total NNAL - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Day 3	n	79	79	41	41
		BLOQ - n (%)	1 (1.3%)		0	
		Geometric Mean (CV%)	60.36 (82.86%)		101.72 (77.70%)	
		Geometric 95% CI	51.33, 70.98		81.88, 126.37	
		Median	72.99	-45.84	122.43	2.69
		Q25, Q75	42.31, 106.33	-53.99, -35.08	58.75, 155.04	-13.93, 10.87
		Min, Max	4.0, 274.2	-78.9, 40.2	21.1, 354.7	-57.9, 43.3
		Mean (SD)	74.77 (46.057)	-44.17 (15.638)	126.55 (84.711)	-0.89 (19.850)
		95% CI			-47.67, -40.66	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (5.00 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_nnal.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.32 Descriptive Statistics of Urinary Total NNAL - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Day 4	n	79	79	41	41
		BLOQ - n (%)	1 (1.3%)		0	
		Geometric Mean (CV%)	55.86 (76.91%)		104.50 (78.93%)	
		Geometric 95% CI	47.94, 65.08		83.89, 130.17	
		Median	66.67	-49.61	114.35	2.08
		Q25, Q75	36.62, 94.39	-57.49, -39.47	69.57, 164.53	-10.74, 18.24
		Min, Max	5.5, 188.1	-79.1, -8.0	22.6, 330.9	-72.0, 45.9
		Mean (SD)	67.61 (37.782)	-48.47 (12.958)	130.14 (85.755)	2.36 (21.486)
		95% CI			-51.37, -45.56	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (5.00 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_nnal.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.32 Descriptive Statistics of Urinary Total NNAL - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Day 5	n	79	79	41	41
		BLOQ - n (%)	5 (6.3%)		0	
		Geometric Mean (CV%)	49.65 (79.09%)		107.04 (79.06%)	
		Geometric 95% CI	42.47, 58.05		85.90, 133.37	
		Median	58.47	-54.66	120.87	-0.59
		Q25, Q75	32.17, 82.52	-61.57, -47.05	60.91, 174.21	-10.71, 16.26
		Min, Max	5.5, 164.7	-84.4, -22.6	20.5, 402.5	-29.6, 73.8
		Mean (SD)	60.46 (33.824)	-53.98 (12.083)	133.36 (89.108)	3.85 (21.172)
		95% CI		-56.69, -51.27		-2.84, 10.54

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (5.00 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_nnal.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.32 Descriptive Statistics of Urinary Total NNAL - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Quantity Excreted over 24 hours (ng)	Baseline	n	39	
		BLOQ - n (%)	0	
		Geometric Mean (CV%)	188.67 (74.51%)	
		Geometric 95% CI	152.11, 234.02	
		Median	196.95	
		Q25, Q75	117.60, 309.23	
		Min, Max	30.2, 554.8	
		Mean (SD)	227.81 (133.914)	
		95% CI		

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (5.00 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_nnal.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.32 Descriptive Statistics of Urinary Total NNAL - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Quantity Excreted over 24 hours (ng)	Day 1	n	39	39
		BLOQ - n (%)	0	
		Geometric Mean (CV%)	119.56 (80.79%)	
		Geometric 95% CI	95.01, 150.45	
		Median	122.12	-35.60
		Q25, Q75	78.54, 208.00	-45.59, -26.62
		Min, Max	16.8, 373.5	-68.1, 5.9
		Mean (SD)	146.96 (87.437)	-35.24 (13.587)
		95% CI		-39.65, -30.83

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (5.00 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_nnal.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.32 Descriptive Statistics of Urinary Total NNAL - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Quantity Excreted over 24 hours (ng)	Day 2	n	39	39
		BLOQ - n (%)	1 (2.6%)	
		Geometric Mean (CV%)	88.41 (94.59%)	
		Geometric 95% CI	68.22, 114.56	
		Median	110.25	-52.63
		Q25, Q75	61.38, 146.88	-61.20, -41.02
		Min, Max	4.8, 382.5	-84.2, -10.0
		Mean (SD)	112.18 (72.198)	-51.03 (14.253)
		95% CI		-55.66, -46.41

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (5.00 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_nnal.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.32 Descriptive Statistics of Urinary Total NNAL - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Quantity Excreted over 24 hours (ng)	Day 3	n	39	39
		BLOQ - n (%)	1 (2.6%)	
		Geometric Mean (CV%)	80.28 (91.64%)	
		Geometric 95% CI	62.32, 103.40	
		Median	93.66	-56.37
		Q25, Q75	60.06, 133.53	-65.20, -47.15
		Min, Max	5.8, 303.5	-80.7, -16.9
		Mean (SD)	101.76 (65.170)	-55.57 (12.993)
		95% CI		-59.79, -51.35

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (5.00 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_nnal.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.32 Descriptive Statistics of Urinary Total NNAL - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Quantity Excreted over 24 hours (ng)	Day 4	n	39	39
		BLOQ - n (%)	1 (2.6%)	
		Geometric Mean (CV%)	69.83 (90.67%)	
		Geometric 95% CI	54.32, 89.77	
		Median	86.94	-62.87
		Q25, Q75	45.15, 114.56	-70.55, -47.18
		Min, Max	6.5, 252.0	-85.9, -38.5
		Mean (SD)	88.85 (57.200)	-60.96 (12.418)
		95% CI		-64.99, -56.93

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (5.00 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_nnal.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.32 Descriptive Statistics of Urinary Total NNAL - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Quantity Excreted over 24 hours (ng)	Day 5	n	39	39
		BLOQ - n (%)	1 (2.6%)	
		Geometric Mean (CV%)	65.12 (89.41%)	
		Geometric 95% CI	50.79, 83.50	
		Median	76.47	-64.53
		Q25, Q75	39.95, 111.07	-72.47, -58.44
		Min, Max	7.7, 256.5	-82.1, -22.9
		Mean (SD)	82.20 (51.378)	-63.83 (11.547)
		95% CI		-67.57, -60.08

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (5.00 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_nnal.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.32 Descriptive Statistics of Urinary Total NNAL - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Baseline	n	39	
		BLOQ - n (%)	0	
		Geometric Mean (CV%)	119.40 (80.36%)	
		Geometric 95% CI	94.97, 150.10	
		Median	124.58	
		Q25, Q75	77.41, 207.05	
		Min, Max	25.0, 387.1	
		Mean (SD)	147.67 (90.845)	
		95% CI		

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (5.00 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_nnal.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.32 Descriptive Statistics of Urinary Total NNAL - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Day 1	n	39	39
		BLOQ - n (%)	0	
		Geometric Mean (CV%)	80.48 (85.35%)	
		Geometric 95% CI	63.31, 102.29	
		Median	91.36	-33.16
		Q25, Q75	47.12, 140.91	-38.03, -20.54
		Min, Max	16.2, 308.2	-65.7, -5.9
		Mean (SD)	102.02 (68.022)	-31.28 (12.811)
		95% CI		-35.44, -27.12

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (5.00 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_nnal.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.32 Descriptive Statistics of Urinary Total NNAL - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Day 2	n	39	39
		BLOQ - n (%)	1 (2.6%)	
		Geometric Mean (CV%)	57.47 (94.85%)	
		Geometric 95% CI	44.32, 74.51	
		Median	62.50	-50.74
		Q25, Q75	36.92, 110.56	-58.54, -40.80
		Min, Max	5.1, 252.2	-79.7, -23.8
		Mean (SD)	74.45 (50.870)	-50.31 (11.882)
		95% CI		-54.17, -46.45

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (5.00 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_nnal.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.32 Descriptive Statistics of Urinary Total NNAL - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Day 3	n	39	39
		BLOQ - n (%)	1 (2.6%)	
		Geometric Mean (CV%)	52.33 (96.09%)	
		Geometric 95% CI	40.25, 68.02	
		Median	58.78	-54.99
		Q25, Q75	30.46, 97.47	-61.12, -46.65
		Min, Max	5.8, 232.9	-76.8, -33.1
		Mean (SD)	68.76 (49.411)	-54.78 (10.924)
		95% CI		-58.33, -51.24

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (5.00 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_nnal.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.32 Descriptive Statistics of Urinary Total NNAL - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Day 4	n	39	39
		BLOQ - n (%)	1 (2.6%)	
		Geometric Mean (CV%)	47.46 (90.69%)	
		Geometric 95% CI	36.92, 61.01	
		Median	50.77	-59.92
		Q25, Q75	28.66, 90.56	-65.47, -50.34
		Min, Max	6.2, 193.5	-85.3, -34.4
		Mean (SD)	61.03 (41.764)	-58.63 (11.050)
		95% CI		-62.22, -55.04

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (5.00 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_nnal.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.32 Descriptive Statistics of Urinary Total NNAL - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Day 5	n	39	39
		BLOQ - n (%)	1 (2.6%)	
		Geometric Mean (CV%)	41.51 (98.88%)	
		Geometric 95% CI	31.76, 54.26	
		Median	44.69	-64.24
		Q25, Q75	25.33, 80.39	-69.86, -56.91
		Min, Max	4.5, 183.7	-82.1, -43.4
		Mean (SD)	54.77 (38.683)	-63.95 (9.331)
		95% CI		-66.98, -60.92

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification (5.00 pg/mL).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_nnal.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.33 Analysis of Urinary NEQ on Day 5 - FAS**

Variable	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	THS : CC Ratio (%)	THS : SA Ratio (%)
Urinary Quantity over n 24 hours (mg)		79	41	39		
	Geometric LS Mean (SE) ¹	15.30	14.89	0.22	102.80 (0.082)	7043.21 (0.083)
	95% CI	13.92, 16.82	13.06, 16.97	0.19, 0.25	87.50, 120.78	5978.67, 8297.29
Concentration Adjusted for Creatinine (mg/g creat)	n	79	41	39		
	Geometric LS Mean (SE) ¹	10.61	10.12	0.14	104.89 (0.067)	7737.73 (0.068)
	95% CI	9.83, 11.46	9.09, 11.26	0.12, 0.16	92.03, 119.55	6773.93, 8838.66

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Adjusted geometric least squares (LS) means and confidence intervals (CIs) from an ANCOVA model conducted on log-transformed Day 5 values with log-transformed baseline value, study arm, sex and CC consumption reported at screening as fixed effect factors.

Baseline is defined as the last assessment prior to Day 1 product use for THS 2.2 and CC subjects and prior to 06:29 AM on Day 1 for SA subjects.

Note: 1 = Standard errors presented from analysis performed on the log_e scale.

Note: Collection over 24 hours starting on the Day stated in the title.

Appendix 15.4.4.33 and 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/tl_anlsecondbio.sas

Program Run: 27OCT14 cvn_ahedge Program Status: FINAL

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**Table 15.2.4.34 Descriptive Statistics of Urinary Total NEQ - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Quantity Excreted over 24 hours (mg)	Baseline	n	80		41	
		BLOQ - n (%)	1 (1.3%)		2 (4.9%)	
		Geometric Mean (CV%)	14.44 (48.67%)		14.31 (45.16%)	
		Geometric 95% CI	13.03, 16.01		12.49, 16.40	
		Median	15.36		14.05	
		Q25, Q75	12.11, 19.56		9.77, 20.20	
		Min, Max	2.4, 29.0		6.8, 38.2	
		Mean (SD)	15.74 (5.741)		15.70 (7.184)	
		95% CI				

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification for one or more component variables.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_neq.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.34 Descriptive Statistics of Urinary Total NEQ - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Quantity Excreted over 24 hours (mg)	Day 1	n	80	80	41	41
		BLOQ - n (%)	3 (3.8%)		1 (2.4%)	
		Geometric Mean (CV%)	13.22 (52.93%)		14.98 (47.74%)	
		Geometric 95% CI	11.83, 14.78		12.98, 17.29	
		Median	15.04	-8.30	14.33	3.38
		Q25, Q75	10.64, 18.28	-18.76, 5.79	10.69, 20.13	-4.66, 13.85
		Min, Max	2.2, 36.4	-57.1, 102.5	6.3, 37.0	-30.5, 93.7
		Mean (SD)	14.63 (5.914)	-5.63 (24.342)	16.57 (7.741)	6.55 (22.005)
		95% CI		-11.05, -0.20		-0.41, 13.50

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification for one or more component variables.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_neq.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.34 Descriptive Statistics of Urinary Total NEQ - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Quantity Excreted over 24 hours (mg)	Day 2	n	80	80	41	41
		BLOQ - n (%)	5 (6.3%)		3 (7.3%)	
		Geometric Mean (CV%)	12.94 (68.35%)		14.21 (49.83%)	
		Geometric 95% CI	11.27, 14.86		12.24, 16.49	
		Median	15.79	-8.04	12.81	-0.17
		Q25, Q75	9.53, 18.90	-28.90, 18.18	9.94, 20.17	-17.43, 15.66
		Min, Max	1.7, 32.9	-79.1, 162.3	6.6, 38.1	-56.8, 158.6
		Mean (SD)	14.99 (6.833)	-3.08 (38.433)	15.90 (8.060)	4.03 (34.878)
		95% CI		-11.64, 5.48		-6.98, 15.05

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification for one or more component variables.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_neq.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.34 Descriptive Statistics of Urinary Total NEQ - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Quantity Excreted over 24 hours (mg)	Day 3	n	79	79	41	41
		BLOQ - n (%)	2 (2.5%)		2 (4.9%)	
		Geometric Mean (CV%)	14.57 (70.80%)		14.59 (52.35%)	
		Geometric 95% CI	12.63, 16.81		12.49, 17.05	
		Median	18.38	5.03	13.79	-2.69
		Q25, Q75	11.73, 21.18	-19.57, 29.06	10.81, 19.48	-16.17, 18.37
		Min, Max	1.1, 34.9	-71.1, 126.3	6.2, 60.5	-41.2, 167.7
		Mean (SD)	16.87 (7.335)	7.65 (40.145)	16.62 (10.035)	6.61 (36.888)
		95% CI		-1.35, 16.64		-5.04, 18.25

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification for one or more component variables.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_neq.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.34 Descriptive Statistics of Urinary Total NEQ - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Quantity Excreted over 24 hours (mg)	Day 4	n	79	79	41	41
		BLOQ - n (%)	3 (3.8%)		3 (7.3%)	
		Geometric Mean (CV%)	15.38 (67.30%)		15.11 (49.10%)	
		Geometric 95% CI	13.41, 17.65		13.04, 17.50	
		Median	17.97	5.53	15.40	5.50
		Q25, Q75	12.34, 22.42	-13.56, 36.18	10.95, 20.81	-3.21, 14.60
		Min, Max	0.8, 42.5	-65.2, 158.4	5.5, 39.7	-39.6, 133.8
		Mean (SD)	17.66 (7.907)	13.00 (43.149)	16.83 (8.366)	8.71 (28.824)
		95% CI		3.33, 22.67		-0.39, 17.81

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification for one or more component variables.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_neq.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.34 Descriptive Statistics of Urinary Total NEQ - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Quantity Excreted over 24 hours (mg)	Day 5	n	79	79	41	41
		BLOQ - n (%)	7 (8.9%)		2 (4.9%)	
		Geometric Mean (CV%)	15.22 (76.42%)		14.66 (53.95%)	
		Geometric 95% CI	13.07, 17.73		12.49, 17.20	
		Median	18.75	9.03	14.05	-0.59
		Q25, Q75	12.30, 24.09	-22.72, 48.45	10.00, 22.18	-10.54, 13.70
		Min, Max	0.9, 48.5	-75.7, 141.0	4.7, 41.2	-48.6, 110.3
		Mean (SD)	18.04 (8.783)	13.95 (45.906)	16.51 (8.091)	6.22 (30.619)
		95% CI		3.67, 24.24		-3.45, 15.89

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification for one or more component variables.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_neq.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.34 Descriptive Statistics of Urinary Total NEQ - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Concentration Adjusted for Creatinine (mg/g creat)	Baseline	n	80		41	
		BLOQ - n (%)	1 (1.3%)		2 (4.9%)	
		Geometric Mean (CV%)	9.01 (51.17%)		8.69 (48.40%)	
		Geometric 95% CI	8.09, 10.03		7.51, 10.04	
		Median	9.14		9.06	
		Q25, Q75	6.96, 12.59		6.24, 11.87	
		Min, Max	1.6, 25.8		3.5, 20.6	
		Mean (SD)	9.99 (4.381)		9.58 (4.210)	
		95% CI				

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification for one or more component variables.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_neq.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.34 Descriptive Statistics of Urinary Total NEQ - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Concentration Adjusted for Creatinine (mg/g creat)	Day 1	n	80	80	41	41
		BLOQ - n (%)	3 (3.8%)		1 (2.4%)	
		Geometric Mean (CV%)	9.19 (49.69%)		10.27 (43.54%)	
		Geometric 95% CI	8.28, 10.21		9.00, 11.72	
		Median	9.44	1.64	11.26	20.51
		Q25, Q75	7.72, 13.20	-9.39, 14.69	7.54, 13.54	9.20, 29.49
		Min, Max	2.0, 23.7	-46.4, 70.2	4.0, 22.4	-14.4, 84.7
		Mean (SD)	10.12 (4.155)	4.00 (20.045)	11.14 (4.491)	19.63 (18.661)
		95% CI		-0.46, 8.47		13.74, 25.53

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification for one or more component variables.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_neq.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.34 Descriptive Statistics of Urinary Total NEQ - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Concentration Adjusted for Creatinine (mg/g creat)	Day 2	n	80	80	41	41
		BLOQ - n (%)	5 (6.3%)		3 (7.3%)	
		Geometric Mean (CV%)	8.66 (59.35%)		8.91 (45.03%)	
		Geometric 95% CI	7.65, 9.79		7.78, 10.21	
		Median	10.08	1.65	9.51	5.31
		Q25, Q75	6.81, 12.75	-22.00, 19.96	6.30, 12.48	-8.02, 17.37
		Min, Max	1.4, 23.0	-74.3, 120.9	3.3, 18.4	-48.0, 66.9
		Mean (SD)	9.77 (4.176)	1.90 (34.137)	9.71 (3.966)	4.62 (20.504)
		95% CI			-5.70, 9.50	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification for one or more component variables.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_neq.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.34 Descriptive Statistics of Urinary Total NEQ - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Concentration Adjusted for Creatinine (mg/g creat)	Day 3	n	79	79	41	41
		BLOQ - n (%)	2 (2.5%)		2 (4.9%)	
		Geometric Mean (CV%)	9.86 (68.24%)		9.42 (45.54%)	
		Geometric 95% CI	8.58, 11.32		8.21, 10.81	
		Median	11.57	18.27	10.27	9.19
		Q25, Q75	7.68, 14.34	-17.52, 36.43	7.08, 12.23	-4.81, 26.78
		Min, Max	0.8, 28.8	-69.5, 190.6	3.3, 19.5	-24.1, 78.1
		Mean (SD)	11.43 (5.461)	17.17 (45.777)	10.27 (4.158)	10.40 (21.176)
		95% CI		6.91, 27.43		3.71, 17.09

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification for one or more component variables.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_neq.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.34 Descriptive Statistics of Urinary Total NEQ - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Concentration Adjusted for Creatinine (mg/g creat)	Day 4	n	79	79	41	41
		BLOQ - n (%)	3 (3.8%)		3 (7.3%)	
		Geometric Mean (CV%)	10.53 (65.38%)		9.91 (45.70%)	
		Geometric 95% CI	9.21, 12.04		8.63, 11.38	
		Median	12.09	19.97	11.67	17.59
		Q25, Q75	8.08, 15.32	-10.59, 46.32	6.95, 13.72	-1.80, 31.61
		Min, Max	0.6, 27.7	-61.4, 154.9	4.1, 19.3	-25.1, 74.2
		Mean (SD)	12.06 (5.480)	23.39 (43.685)	10.80 (4.293)	16.18 (22.108)
		95% CI	13.60, 33.18		9.19, 23.16	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification for one or more component variables.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_neq.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.34 Descriptive Statistics of Urinary Total NEQ - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Concentration Adjusted for Creatinine (mg/g creat)	Day 5	n	79	79	41	41
		BLOQ - n (%)	7 (8.9%)		2 (4.9%)	
		Geometric Mean (CV%)	10.60 (61.33%)		9.76 (43.90%)	
		Geometric 95% CI	9.34, 12.04		8.54, 11.15	
		Median	11.78	17.02	10.70	9.35
		Q25, Q75	7.54, 15.04	-8.96, 52.14	7.18, 12.49	-1.79, 29.41
		Min, Max	0.8, 24.8	-53.3, 128.7	3.7, 22.7	-23.5, 80.2
		Mean (SD)	11.99 (5.153)	22.95 (40.299)	10.61 (4.392)	14.78 (24.526)
		95% CI	13.92, 31.98		7.04, 22.53	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification for one or more component variables.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_neq.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.34 Descriptive Statistics of Urinary Total NEQ - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Quantity Excreted over 24 hours (mg)	Baseline	n	39	
		BLOQ - n (%)	0	
		Geometric Mean (CV%)	15.06 (37.27%)	
		Geometric 95% CI	13.39, 16.93	
		Median	15.96	
		Q25, Q75	11.60, 18.32	
		Min, Max	7.5, 35.7	
		Mean (SD)	16.08 (6.266)	
		95% CI		

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification for one or more component variables.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_neq.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.34 Descriptive Statistics of Urinary Total NEQ - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Quantity Excreted over 24 hours (mg)	Day 1	n	39	39
		BLOQ - n (%)	3 (7.7%)	
		Geometric Mean (CV%)	8.47 (39.90%)	
		Geometric 95% CI	7.47, 9.60	
		Median	8.97	-43.37
		Q25, Q75	6.74, 10.76	-48.99, -36.33
		Min, Max	3.4, 19.6	-84.1, 7.9
		Mean (SD)	9.07 (3.380)	-41.76 (14.539)
		95% CI		-46.48, -37.04

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification for one or more component variables.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_neq.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.34 Descriptive Statistics of Urinary Total NEQ - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Quantity Excreted over 24 hours (mg)	Day 2	n	39	39
		BLOQ - n (%)	38 (97.4%)	
		Geometric Mean (CV%)	3.20 (39.04%)	
		Geometric 95% CI	2.83, 3.62	
		Median	3.31	-79.21
		Q25, Q75	2.50, 4.20	-83.62, -72.68
		Min, Max	1.3, 6.9	-89.4, -44.0
		Mean (SD)	3.42 (1.244)	-77.21 (9.558)
		95% CI		-80.31, -74.11

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification for one or more component variables.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_neq.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.34 Descriptive Statistics of Urinary Total NEQ - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Quantity Excreted over 24 hours (mg)	Day 3	n	39	39
		BLOQ - n (%)	39 (100%)	
		Geometric Mean (CV%)	1.20 (56.73%)	
		Geometric 95% CI	1.01, 1.43	
		Median	1.34	-92.76
		Q25, Q75	0.84, 1.85	-94.14, -89.71
		Min, Max	0.2, 2.8	-98.0, -71.3
		Mean (SD)	1.36 (0.616)	-90.79 (5.444)
		95% CI		-92.56, -89.02

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification for one or more component variables.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_neq.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.34 Descriptive Statistics of Urinary Total NEQ - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Quantity Excreted over 24 hours (mg)	Day 4	n	39	39
		BLOQ - n (%)	39 (100%)	
		Geometric Mean (CV%)	0.50 (66.51%)	
		Geometric 95% CI	0.40, 0.61	
		Median	0.53	-96.96
		Q25, Q75	0.30, 0.73	-97.84, -95.55
		Min, Max	0.1, 1.6	-99.3, -84.9
		Mean (SD)	0.59 (0.348)	-96.00 (2.985)
		95% CI		-96.97, -95.02

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification for one or more component variables.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_neq.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.34 Descriptive Statistics of Urinary Total NEQ - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Quantity Excreted over 24 hours (mg)	Day 5	n	39	39
		BLOQ - n (%)	39 (100%)	
		Geometric Mean (CV%)	0.22 (55.72%)	
		Geometric 95% CI	0.18, 0.27	
		Median	0.22	-98.77
		Q25, Q75	0.14, 0.35	-99.03, -97.95
		Min, Max	0.1, 0.7	-99.3, -92.9
		Mean (SD)	0.26 (0.146)	-98.20 (1.377)
		95% CI		-98.65, -97.75

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification for one or more component variables.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_neq.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.34 Descriptive Statistics of Urinary Total NEQ - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Concentration Adjusted for Creatinine (mg/g creat)	Baseline	n	39	
		BLOQ - n (%)	0	
		Geometric Mean (CV%)	9.53 (38.53%)	
		Geometric 95% CI	8.44, 10.76	
		Median	9.83	
		Q25, Q75	7.50, 12.94	
		Min, Max	4.4, 20.7	
		Mean (SD)	10.17 (3.699)	
		95% CI		

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification for one or more component variables.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_neq.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.34 Descriptive Statistics of Urinary Total NEQ - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Concentration Adjusted for Creatinine (mg/g creat)	Day 1	n	39	39
		BLOQ - n (%)	3 (7.7%)	
		Geometric Mean (CV%)	5.70 (37.29%)	
		Geometric 95% CI	5.07, 6.42	
		Median	5.43	-36.93
		Q25, Q75	4.65, 7.16	-44.70, -29.01
		Min, Max	2.8, 15.1	-83.0, -10.5
		Mean (SD)	6.09 (2.408)	-38.38 (13.076)
		95% CI		-42.62, -34.14

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification for one or more component variables.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_neq.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.34 Descriptive Statistics of Urinary Total NEQ - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Concentration Adjusted for Creatinine (mg/g creat)	Day 2	n	39	39
		BLOQ - n (%)	38 (97.4%)	
		Geometric Mean (CV%)	2.08 (31.10%)	
		Geometric 95% CI	1.88, 2.30	
		Median	2.23	-79.26
		Q25, Q75	1.64, 2.67	-82.30, -70.97
		Min, Max	1.2, 4.1	-88.5, -59.4
		Mean (SD)	2.18 (0.662)	-77.23 (6.863)
		95% CI		-79.45, -75.00

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification for one or more component variables.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_neq.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.34 Descriptive Statistics of Urinary Total NEQ - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Concentration Adjusted for Creatinine (mg/g creat)	Day 3	n	39	39
		BLOQ - n (%)	39 (100%)	
		Geometric Mean (CV%)	0.78 (41.83%)	
		Geometric 95% CI	0.68, 0.90	
		Median	0.82	-92.42
		Q25, Q75	0.65, 1.07	-93.67, -88.97
		Min, Max	0.2, 1.5	-97.0, -69.0
		Mean (SD)	0.84 (0.293)	-90.80 (5.084)
		95% CI		-92.46, -89.15

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification for one or more component variables.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_neq.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.34 Descriptive Statistics of Urinary Total NEQ - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Concentration Adjusted for Creatinine (mg/g creat)	Day 4	n	39	39
		BLOQ - n (%)	39 (100%)	
		Geometric Mean (CV%)	0.34 (47.75%)	
		Geometric 95% CI	0.29, 0.40	
		Median	0.33	-96.64
		Q25, Q75	0.26, 0.46	-97.56, -95.24
		Min, Max	0.1, 0.8	-98.9, -83.4
		Mean (SD)	0.37 (0.160)	-95.85 (2.866)
		95% CI		-96.78, -94.92

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification for one or more component variables.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_neq.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.34 Descriptive Statistics of Urinary Total NEQ - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Concentration Adjusted for Creatinine (mg/g creat)	Day 5	n	39	39
		BLOQ - n (%)	39 (100%)	
		Geometric Mean (CV%)	0.14 (42.70%)	
		Geometric 95% CI	0.12, 0.17	
		Median	0.15	-98.66
		Q25, Q75	0.10, 0.18	-98.92, -97.99
		Min, Max	0.1, 0.3	-99.4, -92.3
		Mean (SD)	0.15 (0.067)	-98.25 (1.279)
		95% CI		-98.67, -97.83

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: BLOQ = number of observations imputed using half limit of quantification for one or more component variables.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_neq.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.35 Descriptive Statistics of Plasma Nicotine Concentrations (ng/mL) - FAS**

Timepoint	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)
Baseline	n	80	41	39
	BLOQ - n (%)	0	0	0
	Geometric Mean (CV%)	14.16 (55.24%)	14.03 (55.69%)	14.39 (41.76%)
	95% CI	12.62, 15.89	11.90, 16.53	12.63, 16.40
	Median	15.25	15.70	14.50
	Q25, Q75	9.66, 20.25	10.20, 20.00	11.30, 18.60
	Min, Max	2.6, 35.1	2.9, 32.2	4.9, 28.4
	Mean (SD)	15.96 (7.507)	15.76 (7.159)	15.50 (5.950)

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: T₀ = Time of the first product use on Day 5.

Note: BLOQ values are imputed by LLOQ/2.

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.3.4

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_pkconc.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.35 Descriptive Statistics of Plasma Nicotine Concentrations (ng/mL) - FAS**

Timepoint	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)
Day 1	n	80	41	37
	BLOQ - n (%)	0	0	8 (21.6%)
	Geometric Mean (CV%)	9.52 (82.59%)	12.28 (56.34%)	0.29 (77.84%)
	95% CI	8.11, 11.19	10.40, 14.50	0.23, 0.37
	Median	11.25	12.50	0.31
	Q25, Q75	7.00, 15.60	9.62, 17.40	0.23, 0.44
	Min, Max	1.0, 33.5	2.9, 31.5	0.1, 1.0
	Mean (SD)	11.62 (6.377)	13.94 (7.067)	0.36 (0.235)

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: T₀ = Time of the first product use on Day 5.

Note: BLOQ values are imputed by LLOQ/2.

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.3.4

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_pkconc.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.4.35 Descriptive Statistics of Plasma Nicotine Concentrations (ng/mL) - FAS**

Timepoint	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)
Day 2	n	80	41	39
	BLOQ - n (%)	0	0	19 (48.7%)
	Geometric Mean (CV%)	14.19 (77.04%)	13.17 (46.48%)	0.17 (61.11%)
	95% CI	12.18, 16.52	11.45, 15.15	0.14, 0.21
	Median	17.70	13.80	0.20
	Q25, Q75	11.15, 22.25	9.90, 18.20	0.10, 0.26
	Min, Max	1.9, 33.4	5.5, 30.9	0.1, 0.6
	Mean (SD)	16.83 (7.901)	14.44 (6.180)	0.20 (0.127)

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: T₀ = Time of the first product use on Day 5.

Note: BLOQ values are imputed by LLOQ/2.

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.3.4

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_pkconc.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.35 Descriptive Statistics of Plasma Nicotine Concentrations (ng/mL) - FAS**

Timepoint	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)
Day 3	n	79	41	39
	BLOQ - n (%)	1 (1.3%)	0	38 (97.4%)
	Geometric Mean (CV%)	14.43 (81.37%)	13.41 (47.88%)	0.10 (17.68%)
	95% CI	12.29, 16.93	11.61, 15.48	0.09, 0.11
	Median	16.20	14.50	0.10
	Q25, Q75	12.20, 22.50	10.30, 18.00	0.10, 0.10
	Min, Max	0.5, 39.8	4.6, 33.9	0.1, 0.3
	Mean (SD)	17.21 (8.503)	14.74 (6.346)	0.11 (0.032)

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: T₀ = Time of the first product use on Day 5.

Note: BLOQ values are imputed by LLOQ/2.

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.3.4

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_pkconc.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.35 Descriptive Statistics of Plasma Nicotine Concentrations (ng/mL) - FAS**

Timepoint	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)
Day 4	n	78	41	38
	BLOQ - n (%)	1 (1.3%)	0	30 (78.9%)
	Geometric Mean (CV%)	13.52 (82.28%)	12.54 (43.62%)	0.12 (42.45%)
	95% CI	11.49, 15.91	10.99, 14.32	0.10, 0.14
	Median	15.25	12.50	0.10
	Q25, Q75	9.82, 20.80	10.30, 15.80	0.10, 0.10
	Min, Max	0.5, 44.0	4.9, 31.1	0.1, 0.4
	Mean (SD)	16.38 (8.980)	13.67 (6.032)	0.13 (0.077)

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: T₀ = Time of the first product use on Day 5.

Note: BLOQ values are imputed by LLOQ/2.

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.3.4

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_pkconc.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.35 Descriptive Statistics of Plasma Nicotine Concentrations (ng/mL) - FAS**

Timepoint	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)
Day 5 , 08:00-10:00 PM	n	0	0	38
	BLOQ - n (%)			36 (94.7%)
	Geometric Mean (CV%)			0.10 (20.54%)
	95% CI			0.09, 0.12
	Median			0.10
	Q25, Q75			0.10, 0.10
	Min, Max			0.1, 0.3
	Mean (SD)			0.11 (0.033)

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: T₀ = Time of the first product use on Day 5.

Note: BLOQ values are imputed by LLOQ/2.

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.3.4

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_pkconc.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.35 Descriptive Statistics of Plasma Nicotine Concentrations (ng/mL) - FAS**

Timepoint	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)
Day 5, 15 min < T ₀	n	79	41	0
	BLOQ - n (%)	14 (17.7%)	5 (12.2%)	
	Geometric Mean (CV%)	1.87 (90.57%)	1.54 (62.30%)	
	95% CI	1.57, 2.23	1.28, 1.85	
	Median	2.01	1.39	
	Q25, Q75	1.34, 3.30	1.17, 2.47	
	Min, Max	0.5, 13.6	0.5, 4.2	
	Mean (SD)	2.45 (1.949)	1.78 (0.917)	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: T₀ = Time of the first product use on Day 5.

Note: BLOQ values are imputed by LLOQ/2.

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.3.4

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_pkconc.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.35 Descriptive Statistics of Plasma Nicotine Concentrations (ng/mL) - FAS**

Timepoint	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)
Day 5, T ₀ + 2 h	n	79	41	0
	BLOQ - n (%)	1 (1.3%)	0	
	Geometric Mean (CV%)	10.00 (90.11%)	8.40 (55.99%)	
	95% CI	8.41, 11.88	7.12, 9.91	
	Median	12.30	8.24	
	Q25, Q75	7.41, 16.80	6.18, 12.70	
	Min, Max	0.5, 30.7	2.3, 19.4	
	Mean (SD)	12.50 (7.169)	9.47 (4.348)	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: T₀ = Time of the first product use on Day 5.

Note: BLOQ values are imputed by LLOQ/2.

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.3.4

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_pkconc.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.35 Descriptive Statistics of Plasma Nicotine Concentrations (ng/mL) - FAS**

Timepoint	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)
Day 5, T ₀ + 4 h	n	79	40	0
	BLOQ - n (%)	2 (2.5%)	0	
	Geometric Mean (CV%)	7.94 (96.61%)	7.02 (95.97%)	
	95% CI	6.62, 9.53	5.42, 9.10	
	Median	9.42	8.08	
	Q25, Q75	5.40, 14.60	3.89, 12.05	
	Min, Max	0.5, 22.4	1.2, 36.9	
	Mean (SD)	10.11 (5.865)	9.43 (7.614)	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: T₀ = Time of the first product use on Day 5.

Note: BLOQ values are imputed by LLOQ/2.

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.3.4

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_pkconc.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.35 Descriptive Statistics of Plasma Nicotine Concentrations (ng/mL) - FAS**

Timepoint	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)
Day 5, T ₀ + 6 h	n	79	41	0
	BLOQ - n (%)	1 (1.3%)	0	
	Geometric Mean (CV%)	14.10 (76.40%)	12.14 (50.16%)	
	95% CI	12.11, 16.42	10.44, 14.10	
	Median	16.80	13.60	
	Q25, Q75	11.30, 21.80	9.01, 16.60	
	Min, Max	0.5, 32.6	3.7, 25.7	
	Mean (SD)	16.40 (7.177)	13.38 (5.545)	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: T₀ = Time of the first product use on Day 5.

Note: BLOQ values are imputed by LLOQ/2.

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.3.4

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_pkconc.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.35 Descriptive Statistics of Plasma Nicotine Concentrations (ng/mL) - FAS**

Timepoint	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)
Day 5, T ₀ + 8 h	n	79	41	0
	BLOQ - n (%)	0	0	
	Geometric Mean (CV%)	12.70 (67.19%)	11.15 (45.27%)	
	95% CI	11.07, 14.57	9.73, 12.79	
	Median	14.50	12.10	
	Q25, Q75	10.10, 19.00	8.59, 14.60	
	Min, Max	1.3, 29.4	4.2, 25.9	
	Mean (SD)	14.62 (6.527)	12.18 (5.171)	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: T₀ = Time of the first product use on Day 5.

Note: BLOQ values are imputed by LLOQ/2.

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.3.4

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_pkconc.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.35 Descriptive Statistics of Plasma Nicotine Concentrations (ng/mL) - FAS**

Timepoint	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)
Day 5, T ₀ + 10 h	n	79	41	0
	BLOQ - n (%)	1 (1.3%)	0	
	Geometric Mean (CV%)	15.24 (79.93%)	14.68 (35.07%)	
	95% CI	13.01, 17.84	13.18, 16.35	
	Median	16.70	14.50	
	Q25, Q75	12.30, 23.40	12.70, 17.70	
	Min, Max	0.5, 41.5	6.6, 35.3	
	Mean (SD)	18.15 (9.086)	15.54 (5.576)	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: T₀ = Time of the first product use on Day 5.

Note: BLOQ values are imputed by LLOQ/2.

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.3.4

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_pkconc.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.35 Descriptive Statistics of Plasma Nicotine Concentrations (ng/mL) - FAS**

Timepoint	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)
Day 5, T ₀ + 12 h	n	79	41	0
	BLOQ - n (%)	1 (1.3%)	0	
	Geometric Mean (CV%)	15.68 (88.00%)	14.31 (48.20%)	
	95% CI	13.23, 18.59	12.38, 16.54	
	Median	18.90	15.70	
	Q25, Q75	10.90, 25.90	11.40, 18.50	
	Min, Max	0.5, 47.9	5.2, 40.7	
	Mean (SD)	19.17 (10.127)	15.78 (7.142)	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: T₀ = Time of the first product use on Day 5.

Note: BLOQ values are imputed by LLOQ/2.

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.3.4

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_pkconc.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.35 Descriptive Statistics of Plasma Nicotine Concentrations (ng/mL) - FAS**

Timepoint	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)
Day 5, T ₀ + 14 h	n	79	41	0
	BLOQ - n (%)	1 (1.3%)	0	
	Geometric Mean (CV%)	16.78 (76.36%)	15.29 (46.34%)	
	95% CI	14.41, 19.54	13.30, 17.58	
	Median	19.20	16.10	
	Q25, Q75	13.80, 25.10	12.40, 18.60	
	Min, Max	0.5, 47.9	6.0, 40.8	
	Mean (SD)	19.65 (9.265)	16.77 (7.429)	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: T₀ = Time of the first product use on Day 5.

Note: BLOQ values are imputed by LLOQ/2.

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.3.4

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_pkconc.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.35 Descriptive Statistics of Plasma Nicotine Concentrations (ng/mL) - FAS**

Timepoint	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)
Day 5, T ₀ + 16 h	n	79	41	0
	BLOQ - n (%)	1 (1.3%)	0	
	Geometric Mean (CV%)	20.74 (89.40%)	19.01 (46.63%)	
	95% CI	17.46, 24.62	16.52, 21.87	
	Median	23.90	20.70	
	Q25, Q75	17.30, 32.50	16.40, 24.50	
	Min, Max	0.5, 73.0	6.2, 41.6	
	Mean (SD)	25.07 (12.534)	20.72 (8.125)	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: T₀ = Time of the first product use on Day 5.

Note: BLOQ values are imputed by LLOQ/2.

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.3.4

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_pkconc.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.35 Descriptive Statistics of Plasma Nicotine Concentrations (ng/mL) - FAS**

Timepoint	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)
% Change from Baseline at Day 5, sample closest to 08:00 PM	n	79	41	38
	Mean (SD)	35.98 (74.949)	19.68 (56.831)	-99.19 (0.613)
	Median	10.98	6.44	-99.30
	Q25, Q75	-14.50, 73.88	-11.25, 35.52	-99.47, -99.11
	Min, Max	-80.7, 279.4	-58.7, 224.6	-99.6, -95.9
	95% CI	19.19, 52.77	1.74, 37.62	-99.40, -98.98

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: T₀ = Time of the first product use on Day 5.

Note: BLOQ values are imputed by LLOQ/2.

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.3.4

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_pkconc.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.4.35 Descriptive Statistics of Plasma Nicotine Concentrations (ng/mL) - FAS**

Timepoint	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)
Day 6/Discharge , 08:00-10:00 AM	n	0	0	39
	BLOQ - n (%)			35 (89.7%)
	Geometric Mean (CV%)			0.11 (25.94%)
	95% CI			0.10, 0.12
	Median			0.10
	Q25, Q75			0.10, 0.10
	Min, Max			0.1, 0.2
	Mean (SD)			0.11 (0.040)

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: T₀ = Time of the first product use on Day 5.

Note: BLOQ values are imputed by LLOQ/2.

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.3.4

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_pkconc.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.4.35 Descriptive Statistics of Plasma Nicotine Concentrations (ng/mL) - FAS**

Timepoint	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)
Day 6/Discharge, T ₀ + 20 h	n	79	41	0
	BLOQ - n (%)	3 (3.8%)	0	
	Geometric Mean (CV%)	7.67 (92.18%)	6.71 (43.14%)	
	95% CI	6.43, 9.15	5.89, 7.65	
	Median	9.40	7.50	
	Q25, Q75	6.05, 12.20	5.29, 8.95	
	Min, Max	0.5, 21.5	2.2, 12.3	
	Mean (SD)	9.40 (4.730)	7.22 (2.550)	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: T₀ = Time of the first product use on Day 5.

Note: BLOQ values are imputed by LLOQ/2.

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.3.4

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_pkconc.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.35 Descriptive Statistics of Plasma Nicotine Concentrations (ng/mL) - FAS**

Timepoint	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)
Day 6/Discharge, T ₀ + 24 h	n	79	41	0
	BLOQ - n (%)	5 (6.3%)	1 (2.4%)	
	Geometric Mean (CV%)	3.09 (81.95%)	2.40 (50.21%)	
	95% CI	2.63, 3.64	2.06, 2.79	
	Median	3.66	2.63	
	Q25, Q75	2.24, 5.40	1.72, 3.42	
	Min, Max	0.5, 11.3	0.5, 4.8	
	Mean (SD)	3.81 (2.272)	2.64 (1.095)	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: T₀ = Time of the first product use on Day 5.

Note: BLOQ values are imputed by LLOQ/2.

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.3.4

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_pkconc.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.36 Descriptive Statistics of Plasma Cotinine Concentrations (ng/mL) - FAS**

Timepoint	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)
Baseline	n	80	41	39
	BLOQ - n (%)	0	0	0
	Geometric Mean (CV%)	208.54 (47.54%)	211.26 (47.86%)	220.60 (27.52%)
	95% CI	188.61, 230.58	183.05, 243.82	202.10, 240.79
	Median	231.00	219.00	212.00
	Q25, Q75	174.50, 279.50	161.00, 286.00	185.00, 277.00
	Min, Max	45.1, 491.0	81.2, 619.0	120.0, 368.0
	Mean (SD)	227.41 (86.286)	232.97 (106.678)	228.38 (60.052)

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: T₀ = Time of the first product use on Day 5.

Note: BLOQ values are imputed by LLOQ/2.

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: LLOQ = 1 ng/mL.

Appendix 15.3.3.6

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_pkconc2.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.36 Descriptive Statistics of Plasma Cotinine Concentrations (ng/mL) - FAS**

Timepoint	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)
Day 1	n	80	41	39
	BLOQ - n (%)	0	0	0
	Geometric Mean (CV%)	177.05 (55.61%)	205.66 (48.05%)	87.61 (41.08%)
	95% CI	157.73, 198.74	178.10, 237.48	77.08, 99.58
	Median	204.50	202.00	92.30
	Q25, Q75	151.50, 246.00	149.00, 264.00	68.00, 116.00
	Min, Max	36.3, 358.0	91.3, 668.0	30.5, 232.0
	Mean (SD)	197.75 (81.156)	228.55 (115.698)	94.47 (38.778)

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: T₀ = Time of the first product use on Day 5.

Note: BLOQ values are imputed by LLOQ/2.

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: LLOQ = 1 ng/mL.

Appendix 15.3.3.6

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_pkconc2.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.36 Descriptive Statistics of Plasma Cotinine Concentrations (ng/mL) - FAS**

Timepoint	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)
Day 2	n	80	41	39
	BLOQ - n (%)	0	0	0
	Geometric Mean (CV%)	184.60 (64.25%)	185.05 (51.09%)	28.83 (60.77%)
	95% CI	161.96, 210.40	158.95, 215.43	24.04, 34.59
	Median	209.50	193.00	28.30
	Q25, Q75	164.50, 268.50	124.00, 273.00	20.80, 42.30
	Min, Max	23.5, 409.0	77.5, 609.0	4.8, 90.3
	Mean (SD)	210.81 (90.240)	207.14 (104.256)	33.24 (18.361)

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: T₀ = Time of the first product use on Day 5.

Note: BLOQ values are imputed by LLOQ/2.

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: LLOQ = 1 ng/mL.

Appendix 15.3.3.6

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_pkconc2.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.36 Descriptive Statistics of Plasma Cotinine Concentrations (ng/mL) - FAS**

Timepoint	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)
Day 3	n	79	41	39
	BLOQ - n (%)	0	0	0
	Geometric Mean (CV%)	227.49 (64.89%)	208.59 (46.73%)	11.45 (77.74%)
	95% CI	199.19, 259.79	181.29, 240.01	9.15, 14.31
	Median	260.00	216.00	10.60
	Q25, Q75	205.00, 326.00	168.00, 277.00	7.86, 18.60
	Min, Max	18.0, 526.0	82.4, 582.0	1.3, 45.8
	Mean (SD)	259.40 (111.272)	229.41 (104.679)	14.20 (9.966)

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: T₀ = Time of the first product use on Day 5.

Note: BLOQ values are imputed by LLOQ/2.

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: LLOQ = 1 ng/mL.

Appendix 15.3.3.6

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_pkconc2.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.36 Descriptive Statistics of Plasma Cotinine Concentrations (ng/mL) - FAS**

Timepoint	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)
Day 4	n	78	41	38
	BLOQ - n (%)	0	0	2 (5.3%)
	Geometric Mean (CV%)	206.54 (66.38%)	194.62 (48.59%)	4.23 (96.75%)
	95% CI	180.23, 236.68	168.29, 225.06	3.23, 5.53
	Median	230.50	198.00	4.11
	Q25, Q75	169.00, 296.00	134.00, 282.00	2.98, 6.38
	Min, Max	8.7, 502.0	88.9, 520.0	0.5, 21.1
	Mean (SD)	234.96 (99.269)	215.45 (99.659)	5.68 (4.731)

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: T₀ = Time of the first product use on Day 5.

Note: BLOQ values are imputed by LLOQ/2.

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: LLOQ = 1 ng/mL.

Appendix 15.3.3.6

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_pkconc2.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.36 Descriptive Statistics of Plasma Cotinine Concentrations (ng/mL) - FAS**

Timepoint	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)
Day 5 , 08:00-10:00 PM	n	0	0	39
	BLOQ - n (%)			6 (15.4%)
	Geometric Mean (CV%)			2.05 (97.55%)
	95% CI			1.56, 2.67
	Median			2.25
	Q25, Q75			1.36, 2.88
	Min, Max			0.5, 10.0
	Mean (SD)			2.78 (2.299)

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: T₀ = Time of the first product use on Day 5.

Note: BLOQ values are imputed by LLOQ/2.

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: LLOQ = 1 ng/mL.

Appendix 15.3.3.6

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_pkconc2.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.36 Descriptive Statistics of Plasma Cotinine Concentrations (ng/mL) - FAS**

Timepoint	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)
Day 5 , 15 min < T ₀	n	79	41	0
	BLOQ - n (%)	0	0	
	Geometric Mean (CV%)	192.15 (70.87%)	175.40 (50.59%)	
	95% CI	166.56, 221.66	150.86, 203.92	
	Median	221.00	175.00	
	Q25, Q75	159.00, 287.00	131.00, 241.00	
	Min, Max	6.8, 449.0	66.8, 534.0	
	Mean (SD)	221.85 (99.374)	195.36 (93.613)	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: T₀ = Time of the first product use on Day 5.

Note: BLOQ values are imputed by LLOQ/2.

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: LLOQ = 1 ng/mL.

Appendix 15.3.3.6

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_pkconc2.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.36 Descriptive Statistics of Plasma Cotinine Concentrations (ng/mL) - FAS**

Timepoint	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)
Day 5 , T ₀ + 2 h	n	79	41	0
	BLOQ - n (%)	0	0	
	Geometric Mean (CV%)	185.07 (70.36%)	167.44 (45.74%)	
	95% CI	160.56, 213.32	145.92, 192.14	
	Median	210.00	166.00	
	Q25, Q75	158.00, 278.00	131.00, 221.00	
	Min, Max	6.2, 469.0	70.3, 480.0	
	Mean (SD)	213.03 (94.420)	183.65 (82.808)	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: T₀ = Time of the first product use on Day 5.

Note: BLOQ values are imputed by LLOQ/2.

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: LLOQ = 1 ng/mL.

Appendix 15.3.3.6

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_pkconc2.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.36 Descriptive Statistics of Plasma Cotinine Concentrations (ng/mL) - FAS**

Timepoint	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)
Day 5, T ₀ + 4 h	n	79	41	0
	BLOQ - n (%)	0	0	
	Geometric Mean (CV%)	186.71 (73.15%)	170.39 (49.83%)	
	95% CI	161.24, 216.20	146.85, 197.70	
	Median	212.00	174.00	
	Q25, Q75	154.00, 284.00	122.00, 229.00	
	Min, Max	5.4, 444.0	69.9, 475.0	
	Mean (SD)	216.22 (96.444)	189.48 (89.718)	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: T₀ = Time of the first product use on Day 5.

Note: BLOQ values are imputed by LLOQ/2.

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: LLOQ = 1 ng/mL.

Appendix 15.3.3.6

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_pkconc2.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.36 Descriptive Statistics of Plasma Cotinine Concentrations (ng/mL) - FAS**

Timepoint	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)
Day 5 , T ₀ + 6 h	n	79	41	0
	BLOQ - n (%)	0	0	
	Geometric Mean (CV%)	209.11 (68.88%)	187.90 (50.36%)	
	95% CI	181.87, 240.44	161.72, 218.33	
	Median	242.00	192.00	
	Q25, Q75	173.00, 298.00	130.00, 259.00	
	Min, Max	8.5, 478.0	81.1, 559.0	
	Mean (SD)	239.85 (103.967)	209.74 (103.086)	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: T₀ = Time of the first product use on Day 5.

Note: BLOQ values are imputed by LLOQ/2.

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: LLOQ = 1 ng/mL.

Appendix 15.3.3.6

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_pkconc2.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.36 Descriptive Statistics of Plasma Cotinine Concentrations (ng/mL) - FAS**

Timepoint	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)
Day 5, T ₀ + 8 h	n	79	41	0
	BLOQ - n (%)	0	0	
	Geometric Mean (CV%)	208.88 (62.78%)	186.67 (48.99%)	
	95% CI	183.57, 237.67	161.25, 216.10	
	Median	233.00	185.00	
	Q25, Q75	176.00, 291.00	135.00, 253.00	
	Min, Max	18.4, 516.0	79.0, 537.0	
	Mean (SD)	237.56 (104.378)	207.36 (99.785)	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: T₀ = Time of the first product use on Day 5.

Note: BLOQ values are imputed by LLOQ/2.

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: LLOQ = 1 ng/mL.

Appendix 15.3.3.6

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_pkconc2.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.36 Descriptive Statistics of Plasma Cotinine Concentrations (ng/mL) - FAS**

Timepoint	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)
Day 5, T ₀ + 10 h	n	79	41	0
	BLOQ - n (%)	0	0	
	Geometric Mean (CV%)	214.30 (60.86%)	193.10 (49.09%)	
	95% CI	188.98, 243.01	166.76, 223.60	
	Median	242.00	194.00	
	Q25, Q75	181.00, 302.00	142.00, 255.00	
	Min, Max	24.9, 505.0	79.0, 569.0	
	Mean (SD)	242.94 (106.754)	214.62 (104.006)	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: T₀ = Time of the first product use on Day 5.

Note: BLOQ values are imputed by LLOQ/2.

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: LLOQ = 1 ng/mL.

Appendix 15.3.3.6

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_pkconc2.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.36 Descriptive Statistics of Plasma Cotinine Concentrations (ng/mL) - FAS**

Timepoint	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)
Day 5, T ₀ + 12 h	n	79	41	0
	BLOQ - n (%)	0	0	
	Geometric Mean (CV%)	222.28 (60.77%)	202.45 (48.99%)	
	95% CI	196.05, 252.03	174.88, 234.37	
	Median	247.00	204.00	
	Q25, Q75	185.00, 314.00	150.00, 272.00	
	Min, Max	25.4, 564.0	83.7, 606.0	
	Mean (SD)	251.98 (111.452)	225.01 (109.981)	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: T₀ = Time of the first product use on Day 5.

Note: BLOQ values are imputed by LLOQ/2.

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: LLOQ = 1 ng/mL.

Appendix 15.3.3.6

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_pkconc2.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.36 Descriptive Statistics of Plasma Cotinine Concentrations (ng/mL) - FAS**

Timepoint	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)
Day 5, T ₀ + 14 h	n	79	41	0
	BLOQ - n (%)	0	0	
	Geometric Mean (CV%)	224.82 (61.02%)	204.69 (48.06%)	
	95% CI	198.20, 255.02	177.26, 236.37	
	Median	244.00	213.00	
	Q25, Q75	187.00, 318.00	156.00, 272.00	
	Min, Max	23.7, 587.0	86.1, 570.0	
	Mean (SD)	254.74 (112.095)	226.56 (107.081)	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: T₀ = Time of the first product use on Day 5.

Note: BLOQ values are imputed by LLOQ/2.

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: LLOQ = 1 ng/mL.

Appendix 15.3.3.6

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_pkconc2.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.36 Descriptive Statistics of Plasma Cotinine Concentrations (ng/mL) - FAS**

Timepoint	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)
Day 5 , T ₀ + 16 h	n	79	41	0
	BLOQ - n (%)	0	0	
	Geometric Mean (CV%)	239.99 (61.76%)	219.73 (48.20%)	
	95% CI	211.30, 272.58	190.21, 253.83	
	Median	261.00	230.00	
	Q25, Q75	198.00, 335.00	169.00, 285.00	
	Min, Max	23.1, 646.0	92.4, 619.0	
	Mean (SD)	272.29 (120.460)	243.18 (114.756)	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: T₀ = Time of the first product use on Day 5.

Note: BLOQ values are imputed by LLOQ/2.

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: LLOQ = 1 ng/mL.

Appendix 15.3.3.6

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_pkconc2.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.36 Descriptive Statistics of Plasma Cotinine Concentrations (ng/mL) - FAS**

Timepoint	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)
% Change from Baseline at n Day 5, sample closest to 08:00 PM		79	41	39
	Mean (SD)	11.94 (35.217)	-0.31 (25.126)	-98.83 (0.815)
	Median	10.92	-5.20	-99.02
	Q25, Q75	-15.19, 27.20	-14.24, 7.94	-99.38, -98.56
	Min, Max	-61.5, 96.2	-34.2, 99.6	-99.8, -96.6
	95% CI	4.05, 19.84	-8.24, 7.63	-99.10, -98.56

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: T₀ = Time of the first product use on Day 5.

Note: BLOQ values are imputed by LLOQ/2.

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: LLOQ = 1 ng/mL.

Appendix 15.3.3.6

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_pkconc2.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.36 Descriptive Statistics of Plasma Cotinine Concentrations (ng/mL) - FAS**

Timepoint	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)
Day 6/Discharge , 08:00-10:00 AM	n	0	0	39
	BLOQ - n (%)			7 (17.9%)
	Geometric Mean (CV%)			1.68 (90.59%)
	95% CI			1.30, 2.16
	Median			1.64
	Q25, Q75			1.19, 2.39
	Min, Max			0.5, 9.1
	Mean (SD)			2.26 (1.984)

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: T₀ = Time of the first product use on Day 5.

Note: BLOQ values are imputed by LLOQ/2.

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: LLOQ = 1 ng/mL.

Appendix 15.3.3.6

Path: /cvm/projects/prj/development/000000106324/dev/tables/t_pkconc2.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.36 Descriptive Statistics of Plasma Cotinine Concentrations (ng/mL) - FAS**

Timepoint	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)
Day 6/Discharge , T ₀ + 20 h	n	79	41	0
	BLOQ - n (%)	0	0	
	Geometric Mean (CV%)	241.97 (63.88%)	221.14 (49.14%)	
	95% CI	212.25, 275.84	190.94, 256.11	
	Median	270.00	224.00	
	Q25, Q75	197.00, 344.00	170.00, 298.00	
	Min, Max	21.4, 608.0	90.2, 641.0	
	Mean (SD)	275.82 (121.876)	245.39 (116.833)	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: T₀ = Time of the first product use on Day 5.

Note: BLOQ values are imputed by LLOQ/2.

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: LLOQ = 1 ng/mL.

Appendix 15.3.3.6

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_pkconc2.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.36 Descriptive Statistics of Plasma Cotinine Concentrations (ng/mL) - FAS**

Timepoint	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)
Day 6/Discharge , T ₀ + 24 h	n	79	41	0
	BLOQ - n (%)	0	0	
	Geometric Mean (CV%)	225.27 (64.18%)	196.80 (47.41%)	
	95% CI	197.49, 256.94	170.72, 226.85	
	Median	250.00	205.00	
	Q25, Q75	179.00, 331.00	153.00, 256.00	
	Min, Max	20.8, 537.0	81.7, 575.0	
	Mean (SD)	257.22 (113.938)	216.87 (99.561)	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: T₀ = Time of the first product use on Day 5.

Note: BLOQ values are imputed by LLOQ/2.

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: LLOQ = 1 ng/mL.

Appendix 15.3.3.6

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_pkconc2.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.37 Descriptive Statistics of Plasma Nicotine Concentration Parameters on Day 5 - FAS**

Parameter	Statistic	THS 2.2 (N=80)	CC (N=41)
C _{peak} (ng/mL)	n	79	40
	Geometric Mean (CV%)	23.39 (65.82%)	20.87 (39.25%)
	95% CI	20.44, 26.76	18.49, 23.56
	Median	26.30	21.20
	Q25, Q75	19.20, 32.90	16.70, 27.70
	Min, Max	2.2, 73.0	7.7, 41.6
t _{peak} (h)	n	79	40
	Median	15.93	15.93
	Q25, Q75	12.00, 15.94	14.00, 15.94
	Min, Max	2.0, 16.2	4.0, 16.0
C _{avg} (ng/mL)	n	79	40
	Geometric Mean (CV%)	12.40 (70.89%)	11.08 (39.31%)
	95% CI	10.75, 14.31	9.81, 12.51
	Median	14.69	11.62
	Q25, Q75	10.15, 17.98	9.23, 14.17
	Min, Max	0.8, 29.9	4.6, 24.6

Note: CC = Conventional cigarettes; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Appendix 15.3.3.5

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_pkparam.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.38 Descriptive Statistics of Plasma Cotinine Concentration Parameters on Day 5 - FAS**

Parameter	Statistic	THS 2.2 (N=80)	CC (N=41)
C _{peak} (ng/mL)	n	79	41
	Geometric Mean (CV%)	259.22 (61.79%)	227.79 (48.35%)
	95% CI	228.21, 294.43	197.11, 263.25
	Median	282.00	230.00
	Q25, Q75	215.00, 370.00	177.00, 298.00
	Min, Max	25.4, 646.0	92.4, 641.0
t _{peak} (h)	n	79	41
	Median	16.03	16.00
	Q25, Q75	14.23, 20.00	15.93, 20.00
	Min, Max	0.0, 24.0	0.0, 24.0
C _{avg} (ng/mL)	n	79	41
	Geometric Mean (CV%)	219.58 (62.34%)	197.98 (47.94%)
	95% CI	193.13, 249.66	171.50, 228.55
	Median	248.04	201.19
	Q25, Q75	180.06, 310.17	147.98, 257.33
	Min, Max	17.8, 479.1	82.4, 570.8

Note: CC = Conventional cigarettes; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Appendix 15.3.3.7

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_pkparam2.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.39 Analysis of Plasma Nicotine Concentration Parameters on Day 5 - FAS**

Parameter	Statistic	THS 2.2 (N=80)	CC (N=41)	THS 2.2:CC Ratio (%)
C _{avg} (ng/mL)	n	79	40	
	Geometric LS Mean (CV%) ¹	12.44	11.01	112.91 (59.51)
	95% CI	11.00, 14.07	9.26, 13.09	91.36, 139.54
C _{peak} (ng/mL)	n	79	40	
	Geometric LS Mean (CV%) ¹	23.44	20.78	112.80 (56.42)
	95% CI	20.85, 26.36	17.62, 24.51	92.15, 138.07

Note: CC = Conventional cigarettes; THS = Tobacco Heating System.

Note: 1: Geometric LS Mean and 95% CI are the adjusted geometric least squares means based on an ANOVA model conducted on log-transformed values with study arm, sex and CC consumption reported at screening as fixed effect factors. Geometrical CV% of the ratio is estimated from the residual mean squares.

Note: 2: 95% CI are estimated only for the median difference based on the Hodges-Lehmann estimate.

Appendix 15.2.4.37

Path: /cvn/projects/prj/development/000000106324/dev/tables/tl_anlparm.sas

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Program Run: 27OCT14 cvn_ahedge Program Status: FINAL

**Table 15.2.4.39 Analysis of Plasma Nicotine Concentration Parameters on Day 5 - FAS**

Parameter	Statistic	THS 2.2 (N=80)	CC (N=41)	THS 2.2 - CC
t _{peak} (h)	n	79	40	
	Median	15.93	15.93	0.00
	95% CI ²			0.00, 0.00

Note: CC = Conventional cigarettes; THS = Tobacco Heating System.

Note: 1: Geometric LS Mean and 95% CI are the adjusted geometric least squares means based on an ANOVA model conducted on log-transformed values with study arm, sex and CC consumption reported at screening as fixed effect factors. Geometrical CV% of the ratio is estimated from the residual mean squares.

Note: 2: 95% CI are estimated only for the median difference based on the Hodges-Lehmann estimate.

Appendix 15.2.4.37

Path: /cvn/projects/prj/development/000000106324/dev/tables/tl_anlpkparm.sas

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Program Run: 27OCT14 cvn_ahedge Program Status: FINAL

**Table 15.2.4.40 Analysis of Plasma Cotine Concentration Parameters on Day 5 - FAS**

Parameter	Statistic	THS 2.2 (N=80)	CC (N=41)	THS 2.2:CC Ratio (%)
C _{avg} (ng/mL)	n	79	41	
	Geometric LS Mean (CV%) ¹	220.07	198.21	111.03 (56.37)
	95% CI	195.76, 247.40	168.47, 233.19	90.87, 135.65
C _{peak} (ng/mL)	n	79	41	
	Geometric LS Mean (CV%) ¹	259.79	228.23	113.83 (55.86)
	95% CI	231.30, 291.78	194.25, 268.16	93.31, 138.85

Note: CC = Conventional cigarettes; THS = Tobacco Heating System.

Note: 1: Geometric LS Mean and 95% CI are the adjusted geometric least squares means based on an ANOVA model conducted on log-transformed values with study arm, sex and CC consumption reported at screening as fixed effect factors. Geometrical CV% of the ratio is estimated from the residual mean squares.

Note: 2: 95% CI are estimated only for the median difference based on the Hodges-Lehmann estimate.

Appendix 15.2.4.38

Path: /cvn/projects/prj/development/000000106324/dev/tables/tl_anlparm.sas

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Program Run: 27OCT14 cvn_ahedge Program Status: FINAL

**Table 15.2.4.40 Analysis of Plasma Cotine Concentration Parameters on Day 5 - FAS**

Parameter	Statistic	THS 2.2 (N=80)	CC (N=41)	THS 2.2 - CC
t _{peak} (h)	n	79	41	
	Median	16.03	16.00	0.00
	95% CI ²			0.00, 0.07

Note: CC = Conventional cigarettes; THS = Tobacco Heating System.

Note: 1: Geometric LS Mean and 95% CI are the adjusted geometric least squares means based on an ANOVA model conducted on log-transformed values with study arm, sex and CC consumption reported at screening as fixed effect factors. Geometrical CV% of the ratio is estimated from the residual mean squares.

Note: 2: 95% CI are estimated only for the median difference based on the Hodges-Lehmann estimate.

Appendix 15.2.4.38

Path: /cvn/projects/prj/development/000000106324/dev/tables/tl_anlpkparm.sas

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Program Run: 27OCT14 cvn_ahedge Program Status: FINAL

**Table 15.2.4.41 Analysis of Change from Day 0 Plasma Nicotine and Cotinine Concentrations at 08:00 PM on Day 5 - FAS**

Variable (units)	Statistic	THS 2.2 (N=80)	CC (N=41)	THS - CC
Cotinine (ng/mL)	n	79	41	
	LS Mean	25.20	-5.70	30.91
	95% CI	10.86, 39.55	-25.61, 14.21	6.36, 55.45
Nicotine (ng/mL)	n	79	41	
	LS Mean	3.54	0.99	2.55
	95% CI	1.93, 5.14	-1.24, 3.21	-0.19, 5.29

Note: CC = Conventional cigarettes; THS = Tobacco Heating System.

Note: Adjusted least squares (LS) means and confidence intervals (CIs) from an ANCOVA model conducted on the change from Day 0 in Day 5 (result closest to 08:00 PM) values with log-transformed Day 0 value, study arm, sex and CC consumption reported at screening as fixed effect factors.

Appendix 15.4.4.41 and 15.3.3.4

Path: /cvn/projects/prj/development/000000106324/dev/tables/tl_anlpkconc.sas

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Program Run: 27OCT14 cvn_ahedge Program Status: FINAL

**Table 15.2.4.42 Descriptive Statistics of Ames Mutagenicity Test (YG1024+S9) (REV/24h) - FAS**

Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
		Raw value	% Change (*)	Raw value	% Change (*)
Baseline	n	64		28	
	Geometric Mean (CV%)	NC		NC	
	Geometric 95% CI	NC		NC	
	Median	19681.50		15775.00	
	Q25, Q75	9203.00, 31122.50		9240.00, 20180.00	
	Min, Max	0.0, 107250.0		0.0, 72216.0	
	Mean (SD)	23199.26 (21054.119)		16914.76 (14336.658)	
	95% CI	17940.09, 28458.43		11355.58, 22473.94	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: NC = Not calculated.

Appendix 15.3.5.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_ames.sas

Program Run: 05NOV15 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.42 Descriptive Statistics of Ames Mutagenicity Test (YG1024+S9) (REV/24h) - FAS**

Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
		Raw value	% Change (*)	Raw value	% Change (*)
Day 5	n	52	42	31	23
	Geometric Mean (CV%)	NC		NC	
	Geometric 95% CI	NC		NC	
	Median	8822.75	-56.93	21689.00	29.29
	Q25, Q75	5361.00, 15202.00	-74.50, -7.25	16641.00, 35560.50	-11.60, 98.60
	Min, Max	0.0, 39600.0	-100.0, 1707300.0	0.0, 63840.0	-64.5, 6111000.0
	Mean (SD)	10670.52 (8155.693)	52997.96 (273527.662)	26975.64 (15866.931)	370667.73 (1348543.061)
	95% CI	8399.95, 12941.08	-32239.27, 138235.18	21155.59, 32795.68	-212486.03, 953821.48

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: NC = Not calculated.

Appendix 15.3.5.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_ames.sas

Program Run: 05NOV15 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.42 Descriptive Statistics of Ames Mutagenicity Test (YG1024+S9) (REV/24h) - FAS**

Timepoint	Statistic	SA (N=39)	
		Raw value	% Change (*)
Baseline	n	23	
	Geometric Mean (CV%)	19556.81 (128.70%)	
	Geometric 95% CI	12754.86, 29986.13	
	Median	22540.00	
	Q25, Q75	7716.30, 35700.00	
	Min, Max	3060.0, 113620.0	
	Mean (SD)	30212.25 (29347.944)	
	95% CI	17521.24, 42903.26	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: NC = Not calculated.

Appendix 15.3.5.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_ames.sas

Program Run: 05NOV15 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.42 Descriptive Statistics of Ames Mutagenicity Test (YG1024+S9) (REV/24h) - FAS**

Timepoint	Statistic	SA (N=39)	
		Raw value	% Change (*)
Day 5	n	26	17
	Geometric Mean (CV%)	NC	
	Geometric 95% CI	NC	
	Median	7437.00	-62.44
	Q25, Q75	3813.00, 12221.00	-87.07, -11.93
	Min, Max	0.0, 65400.0	-100.0, 768.6
	Mean (SD)	10609.63 (12918.414)	-3.67 (203.176)
	95% CI	5391.76, 15827.49	-108.14, 100.80

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: NC = Not calculated.

Appendix 15.3.5.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_ames.sas

Program Run: 05NOV15 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.43 Descriptive Statistics of QSU-brief Questionnaire Factors and Total Score - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)	
			Raw value	% Change(*)	Raw value	% Change(*)	Raw value	% Change(*)
Factor 1 - Reward	Baseline	n	80		41		39	
		Mean (SD)	4.29 (1.903)		3.90 (1.863)		4.13 (1.643)	
		95% CI	3.86, 4.72		3.31, 4.50		3.59, 4.67	
		Median	4.00		3.40		4.00	
		Q25, Q75	2.70, 6.20		2.40, 5.00		2.80, 5.60	
		Min, Max	1.0, 7.0		1.0, 7.0		1.4, 7.0	
	Day 1	n	80	80	41	41	39	39
		Mean (SD)	3.76 (2.044)	-2.15 (56.101)	3.94 (1.745)	9.39 (51.441)	5.91 (1.366)	63.35 (72.613)
		95% CI	3.30, 4.22	-14.64, 10.34	3.39, 4.50	-6.85, 25.63	5.46, 6.36	39.81, 86.90
		Median	3.20	-8.70	3.80	0.00	6.40	45.83
		Q25, Q75	2.10, 5.60	-39.70, 8.54	2.60, 5.00	-15.63, 15.79	5.60, 7.00	6.25, 100.00
		Min, Max	1.0, 7.0	-85.7, 180.0	1.0, 7.0	-54.5, 257.1	2.0, 7.0	-33.3, 271.4

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: QSU-brief scores reported on a 7-point scale. Higher scores indicate greater intensity of urge.

Appendix 15.3.6.11

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_qsu.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.4.43 Descriptive Statistics of QSU-brief Questionnaire Factors and Total Score - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)	
			Raw value	% Change(*)	Raw value	% Change(*)	Raw value	% Change(*)
Factor 1 - Reward	Day 2	n	80	80	41	41	39	39
		Mean (SD)	3.89 (1.896)	5.34 (65.313)	3.98 (1.798)	10.90 (50.573)	5.77 (1.459)	57.73 (64.552)
		95% CI	3.46, 4.31	-9.20, 19.87	3.40, 4.55	-5.06, 26.87	5.30, 6.25	36.80, 78.66
		Median	3.80	-9.88	3.80	0.00	6.20	38.10
		Q25, Q75	2.20, 5.50	-35.58, 18.83	2.60, 5.40	-11.43, 17.40	5.20, 7.00	2.94, 107.15
		Min, Max	1.0, 7.0	-85.7, 312.5	1.0, 7.0	-54.5, 171.4	2.0, 7.0	-30.8, 191.7
	Day 3	n	79	79	41	41	39	39
		Mean (SD)	3.70 (1.935)	-3.68 (50.930)	3.99 (1.929)	11.78 (64.807)	5.69 (1.528)	56.61 (69.093)
		95% CI	3.27, 4.14	-15.09, 7.74	3.37, 4.60	-8.68, 32.24	5.19, 6.19	34.21, 79.01
		Median	3.80	-8.57	3.80	-4.00	6.40	40.00
		Q25, Q75	1.80, 5.00	-34.38, 14.29	2.40, 5.40	-19.05, 15.00	4.80, 7.00	3.22, 107.15
		Min, Max	1.0, 7.0	-85.7, 150.0	1.0, 7.0	-54.5, 328.6	1.8, 7.0	-42.3, 244.4

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: QSU-brief scores reported on a 7-point scale. Higher scores indicate greater intensity of urge.

Appendix 15.3.6.11

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_qsu.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.4.43 Descriptive Statistics of QSU-brief Questionnaire Factors and Total Score - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)	
			Raw value	% Change(*)	Raw value	% Change(*)	Raw value	% Change(*)
Factor 1 - Reward	Day 4	n	79	79	41	41	39	39
		Mean (SD)	3.59 (1.856)	-7.87 (50.349)	3.59 (1.976)	-5.52 (39.938)	5.14 (1.919)	38.41 (63.278)
		95% CI	3.17, 4.01	-19.15, 3.41	2.96, 4.21	-18.13, 7.09	4.51, 5.77	17.89, 58.93
		Median	3.40	-14.29	3.20	-5.71	5.80	33.33
		Q25, Q75	1.80, 5.00	-41.18, 0.00	2.00, 5.20	-25.00, 3.13	3.40, 7.00	-2.95, 78.95
		Min, Max	1.0, 7.0	-85.7, 262.5	1.0, 7.0	-66.7, 180.0	1.4, 7.0	-75.0, 191.7
	Day 5	n	79	79	41	41	39	39
		Mean (SD)	3.44 (1.927)	-6.89 (66.443)	3.65 (2.067)	0.69 (58.884)	5.08 (1.952)	37.69 (70.278)
		95% CI	3.00, 3.87	-21.78, 8.00	3.00, 4.31	-17.90, 19.28	4.44, 5.71	14.90, 60.48
		Median	3.00	-16.67	3.20	-13.33	5.60	25.00
		Q25, Q75	1.80, 5.00	-55.00, 4.17	2.00, 5.40	-30.44, 9.38	3.40, 7.00	-9.53, 75.00
		Min, Max	1.0, 7.0	-84.8, 337.5	1.0, 7.0	-66.7, 250.0	1.0, 7.0	-70.4, 244.4

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: QSU-brief scores reported on a 7-point scale. Higher scores indicate greater intensity of urge.

Appendix 15.3.6.11

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_qsu.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.4.43 Descriptive Statistics of QSU-brief Questionnaire Factors and Total Score - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)	
			Raw value	% Change(*)	Raw value	% Change(*)	Raw value	% Change(*)
Factor 2 - Relief	Baseline	n	80		41		39	
		Mean (SD)	3.08 (1.846)		3.08 (1.913)		2.86 (1.455)	
		95% CI	2.66, 3.50		2.47, 3.69		2.38, 3.33	
		Median	2.70		2.20		2.40	
		Q25, Q75	1.50, 4.40		1.60, 4.20		1.80, 3.80	
		Min, Max	1.0, 7.0		1.0, 7.0		1.0, 7.0	
	Day 1	n	80	80	41	41	39	39
		Mean (SD)	2.84 (1.847)	2.76 (51.364)	2.98 (1.748)	1.20 (26.052)	4.67 (1.702)	95.95 (116.975)
		95% CI	2.43, 3.26	-8.68, 14.19	2.42, 3.53	-7.02, 9.43	4.12, 5.23	58.03, 133.88
		Median	2.20	0.00	2.60	0.00	4.80	50.00
		Q25, Q75	1.20, 4.10	-25.83, 19.10	1.60, 4.00	-17.25, 13.34	3.60, 6.20	15.38, 130.00
		Min, Max	1.0, 7.0	-82.8, 180.0	1.0, 7.0	-46.7, 76.9	1.0, 7.0	-47.4, 520.0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: QSU-brief scores reported on a 7-point scale. Higher scores indicate greater intensity of urge.

Appendix 15.3.6.11

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_qsu.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.4.43 Descriptive Statistics of QSU-brief Questionnaire Factors and Total Score - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)	
			Raw value	% Change(*)	Raw value	% Change(*)	Raw value	% Change(*)
Factor 2 - Relief	Day 2	n	80	80	41	41	39	39
		Mean (SD)	2.65 (1.726)	1.51 (70.869)	2.84 (1.838)	-4.18 (31.600)	4.38 (1.874)	79.10 (112.279)
		95% CI	2.26, 3.04	-14.26, 17.29	2.26, 3.43	-14.16, 5.80	3.77, 5.00	42.70, 115.50
		Median	2.10	-5.06	2.20	-5.88	4.60	33.33
		Q25, Q75	1.20, 3.50	-36.04, 14.65	1.40, 4.00	-26.67, 10.00	2.80, 6.20	5.00, 126.67
		Min, Max	1.0, 7.0	-85.7, 420.0	1.0, 7.0	-54.5, 100.0	1.2, 7.0	-38.1, 500.0
	Day 3	n	79	79	41	41	39	39
		Mean (SD)	2.55 (1.701)	-8.33 (41.184)	2.83 (1.751)	-4.64 (25.712)	4.28 (1.804)	77.40 (113.863)
		95% CI	2.16, 2.94	-17.56, 0.89	2.27, 3.39	-12.76, 3.48	3.69, 4.87	40.49, 114.32
		Median	1.80	-3.45	2.20	0.00	4.40	46.67
		Q25, Q75	1.00, 3.40	-33.34, 7.70	1.40, 4.00	-20.00, 10.00	2.80, 6.00	0.00, 130.00
		Min, Max	1.0, 7.0	-85.7, 160.0	1.0, 7.0	-50.0, 90.9	1.0, 7.0	-42.9, 520.0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: QSU-brief scores reported on a 7-point scale. Higher scores indicate greater intensity of urge.

Appendix 15.3.6.11

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_qsu.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.4.43 Descriptive Statistics of QSU-brief Questionnaire Factors and Total Score - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)	
			Raw value	% Change(*)	Raw value	% Change(*)	Raw value	% Change(*)
Factor 2 - Relief	Day 4	n	79	79	41	41	39	39
		Mean (SD)	2.61 (1.767)	-6.36 (49.195)	2.82 (1.877)	-4.70 (38.176)	3.78 (1.981)	50.35 (91.316)
		95% CI	2.21, 3.01	-17.38, 4.66	2.22, 3.42	-16.76, 7.35	3.13, 4.43	20.74, 79.95
		Median	2.00	-9.09	2.00	-12.50	3.40	22.22
		Q25, Q75	1.00, 3.80	-36.37, 8.70	1.40, 4.00	-26.09, 0.00	2.00, 5.60	-4.17, 90.91
		Min, Max	1.0, 7.0	-82.8, 280.0	1.0, 7.0	-54.5, 125.0	1.0, 7.0	-68.4, 337.5
	Day 5	n	79	79	41	41	39	39
		Mean (SD)	2.43 (1.667)	-5.00 (74.436)	2.75 (1.849)	-3.29 (58.827)	3.72 (2.060)	46.16 (91.447)
		95% CI	2.05, 2.81	-21.68, 11.67	2.16, 3.33	-21.87, 15.28	3.05, 4.40	16.51, 75.81
		Median	1.80	-14.29	2.00	-13.64	3.20	20.00
		Q25, Q75	1.00, 3.40	-47.06, 0.00	1.20, 3.80	-33.34, 4.17	2.00, 5.60	-11.54, 94.12
		Min, Max	1.0, 7.0	-82.8, 420.0	1.0, 7.0	-68.6, 300.0	1.0, 7.0	-73.7, 337.5

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: QSU-brief scores reported on a 7-point scale. Higher scores indicate greater intensity of urge.

Appendix 15.3.6.11

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_qsu.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.4.43 Descriptive Statistics of QSU-brief Questionnaire Factors and Total Score - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)	
			Raw value	% Change(*)	Raw value	% Change(*)	Raw value	% Change(*)
Total score	Baseline	n	80		41		39	
		Mean (SD)	3.69 (1.807)		3.49 (1.858)		3.49 (1.478)	
		95% CI	3.28, 4.09		2.90, 4.08		3.01, 3.98	
		Median	3.15		2.90		3.20	
		Q25, Q75	2.20, 5.30		2.10, 4.70		2.10, 4.70	
		Min, Max	1.0, 7.0		1.0, 7.0		1.4, 7.0	
	Day 1	n	80	80	41	41	39	39
		Mean (SD)	3.30 (1.877)	-1.11 (52.547)	3.46 (1.676)	5.57 (36.135)	5.29 (1.448)	73.02 (79.062)
		95% CI	2.88, 3.73	-12.81, 10.59	2.92, 3.99	-5.84, 16.98	4.82, 5.76	47.38, 98.65
		Median	2.70	-7.06	3.30	0.00	5.50	51.28
		Q25, Q75	1.90, 5.00	-35.93, 14.15	2.00, 4.50	-15.79, 12.50	4.80, 6.50	14.03, 124.00
		Min, Max	1.0, 7.0	-84.4, 170.6	1.0, 7.0	-37.5, 153.8	1.6, 7.0	-37.0, 250.0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: QSU-brief scores reported on a 7-point scale. Higher scores indicate greater intensity of urge.

Appendix 15.3.6.11

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_qsu.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.4.43 Descriptive Statistics of QSU-brief Questionnaire Factors and Total Score - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)	
			Raw value	% Change(*)	Raw value	% Change(*)	Raw value	% Change(*)
Total score	Day 2	n	80	80	41	41	39	39
		Mean (SD)	3.27 (1.715)	2.75 (64.478)	3.41 (1.763)	4.13 (38.854)	5.08 (1.568)	63.83 (73.235)
		95% CI	2.88, 3.66	-11.61, 17.10	2.85, 3.97	-8.14, 16.40	4.57, 5.59	40.09, 87.57
		Median	2.90	-11.47	2.60	-1.79	5.40	42.31
		Q25, Q75	1.90, 4.45	-34.85, 18.51	2.10, 4.70	-15.79, 17.65	3.80, 6.60	5.17, 100.00
		Min, Max	1.0, 7.0	-85.7, 353.8	1.0, 7.0	-51.2, 123.5	1.7, 7.0	-34.0, 242.1
	Day 3	n	79	79	41	41	39	39
		Mean (SD)	3.13 (1.743)	-6.66 (43.801)	3.41 (1.751)	4.39 (42.490)	4.99 (1.585)	61.82 (75.745)
		95% CI	2.73, 3.52	-16.48, 3.16	2.85, 3.96	-9.02, 17.81	4.47, 5.51	37.26, 86.38
		Median	2.80	-7.55	2.80	-4.17	5.20	40.00
		Q25, Q75	1.70, 4.20	-37.50, 10.00	2.00, 4.60	-20.32, 16.22	4.00, 6.40	4.76, 95.46
		Min, Max	1.0, 7.0	-85.7, 117.6	1.0, 7.0	-48.8, 184.6	1.5, 7.0	-42.6, 247.4

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: QSU-brief scores reported on a 7-point scale. Higher scores indicate greater intensity of urge.

Appendix 15.3.6.11

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_qsu.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.4.43 Descriptive Statistics of QSU-brief Questionnaire Factors and Total Score - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)	
			Raw value	% Change(*)	Raw value	% Change(*)	Raw value	% Change(*)
Total score	Day 4	n	79	79	41	41	39	39
		Mean (SD)	3.10 (1.740)	-8.23 (47.798)	3.20 (1.879)	-5.47 (36.942)	4.46 (1.864)	41.33 (68.796)
		95% CI	2.71, 3.50	-18.94, 2.48	2.60, 3.80	-17.14, 6.19	3.85, 5.07	19.02, 63.63
		Median	2.70	-12.00	2.30	-9.52	4.80	37.14
		Q25, Q75	1.70, 4.20	-36.67, 8.11	1.80, 4.50	-23.08, 0.00	2.70, 6.30	0.00, 76.48
		Min, Max	1.0, 7.0	-84.4, 269.2	1.0, 7.0	-61.5, 152.6	1.2, 7.0	-72.5, 233.3
	Day 5	n	79	79	41	41	39	39
		Mean (SD)	2.93 (1.658)	-7.58 (65.083)	3.20 (1.907)	-1.25 (57.331)	4.40 (1.904)	40.09 (74.364)
		95% CI	2.56, 3.31	-22.16, 7.00	2.59, 3.81	-19.35, 16.85	3.78, 5.02	15.98, 64.20
		Median	2.80	-20.00	2.50	-11.63	4.80	27.27
		Q25, Q75	1.60, 3.90	-50.00, 5.41	1.60, 4.70	-31.25, 7.15	2.90, 6.20	-3.93, 73.69
		Min, Max	1.0, 7.0	-78.7, 369.2	1.0, 7.0	-61.5, 270.6	1.0, 7.0	-71.7, 247.1

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: QSU-brief scores reported on a 7-point scale. Higher scores indicate greater intensity of urge.

Appendix 15.3.6.11

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_qsu.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.4.44 Analysis of Change from Baseline QSU-brief Questionnaire Factors and Total Score - FAS**

Variable	Time point	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	THS 2.2-CC	THS 2.2-SA
Total Score	Overall	n	80	41	39		
		LS Mean	-0.52	-0.20	1.31	-0.31	-1.83
		95% CI	-0.77, -0.26	-0.56, 0.15	0.95, 1.68	-0.75, 0.12	-2.27, -1.39
	Day 1	n	80	41	39		
		LS Mean	-0.35	-0.08	1.76	-0.27	-2.11
		95% CI	-0.65, -0.05	-0.50, 0.34	1.33, 2.19	-0.78, 0.24	-2.63, -1.59
	Day 2	n	80	41	39		
		LS Mean	-0.38	-0.13	1.55	-0.26	-1.93
		95% CI	-0.69, -0.08	-0.55, 0.29	1.12, 1.98	-0.77, 0.26	-2.46, -1.41
	Day 3	n	79	41	39		
		LS Mean	-0.54	-0.13	1.46	-0.41	-2.00
		95% CI	-0.84, -0.24	-0.54, 0.28	1.03, 1.88	-0.92, 0.10	-2.52, -1.48

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Adjusted least squares (LS) means and confidence intervals (CIs) from an ANCOVA model with study arm, sex, CC consumption reported at screening, day and study arm*day fitted as fixed effect factors with baseline fitted as a covariate. Day fitted as a repeated factor. Where baseline is defined as the last assessment prior to Day 1 product use for THS 2.2 and CC subjects and prior to 06:29 AM on Day 1 for SA subjects.

Note: Comparison overall time points is the main comparison.

Note: QSU-brief scores reported on a 7-point scale. Higher scores indicate greater intensity of urge.

Appendix 15.4.4.44 and 15.3.6.11

Path: /cvn/projects/prj/development/000000106324/dev/tables/tlf_anlqsu.sas

Program Run: 27OCT14 cvn_ahedge Program Status: FINAL

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**Table 15.2.4.44 Analysis of Change from Baseline QSU-brief Questionnaire Factors and Total Score - FAS**

Variable	Time point	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	THS 2.2-CC	THS 2.2-SA
Total Score	Day 4	n	79	41	39		
		LS Mean	-0.57	-0.34	0.93	-0.23	-1.50
		95% CI	-0.88, -0.25	-0.77, 0.10	0.49, 1.37	-0.76, 0.30	-2.04, -0.96
	Day 5	n	79	41	39		
		LS Mean	-0.73	-0.34	0.87	-0.40	-1.60
		95% CI	-1.08, -0.39	-0.82, 0.14	0.38, 1.36	-0.99, 0.19	-2.21, -1.00
Factor 1 - Reward	Overall	n	80	41	39		
		LS Mean	-0.56	-0.20	1.38	-0.36	-1.94
		95% CI	-0.85, -0.28	-0.60, 0.20	0.97, 1.79	-0.85, 0.13	-2.44, -1.45
	Day 1	n	80	41	39		
		LS Mean	-0.47	-0.09	1.77	-0.38	-2.24
		95% CI	-0.81, -0.12	-0.57, 0.39	1.28, 2.26	-0.97, 0.21	-2.83, -1.64

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Adjusted least squares (LS) means and confidence intervals (CIs) from an ANCOVA model with study arm, sex, CC consumption reported at screening, day and study arm*day fitted as fixed effect factors with baseline fitted as a covariate. Day fitted as a repeated factor. Where baseline is defined as the last assessment prior to Day 1 product use for THS 2.2 and CC subjects and prior to 06:29 AM on Day 1 for SA subjects.

Note: Comparison overall time points is the main comparison.

Note: QSU-brief scores reported on a 7-point scale. Higher scores indicate greater intensity of urge.

Appendix 15.4.4.44 and 15.3.6.11

Path: /cvn/projects/prj/development/000000106324/dev/tables/tlf_anlqsu.sas

Program Run: 27OCT14 cvn_ahedge Program Status: FINAL

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**Table 15.2.4.44 Analysis of Change from Baseline QSU-brief Questionnaire Factors and Total Score - FAS**

Variable	Time point	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	THS 2.2-CC	THS 2.2-SA
Factor 1 - Reward	Day 2	n	80	41	39		
		LS Mean	-0.34	-0.06	1.64	-0.29	-1.98
		95% CI	-0.69, 0.00	-0.53, 0.42	1.15, 2.13	-0.88, 0.30	-2.58, -1.38
	Day 3	n	79	41	39		
		LS Mean	-0.55	-0.05	1.55	-0.50	-2.10
		95% CI	-0.90, -0.19	-0.53, 0.44	1.05, 2.05	-1.10, 0.10	-2.71, -1.49
	Day 4	n	79	41	39		
		LS Mean	-0.66	-0.45	1.00	-0.21	-1.66
		95% CI	-1.01, -0.30	-0.94, 0.04	0.50, 1.50	-0.81, 0.39	-2.27, -1.04
	Day 5	n	79	41	39		
		LS Mean	-0.81	-0.38	0.94	-0.43	-1.75
		95% CI	-1.21, -0.41	-0.94, 0.18	0.37, 1.51	-1.12, 0.25	-2.45, -1.05

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Adjusted least squares (LS) means and confidence intervals (CIs) from an ANCOVA model with study arm, sex, CC consumption reported at screening, day and study arm*day fitted as fixed effect factors with baseline fitted as a covariate. Day fitted as a repeated factor. Where baseline is defined as the last assessment prior to Day 1 product use for THS 2.2 and CC subjects and prior to 06:29 AM on Day 1 for SA subjects.

Note: Comparison overall time points is the main comparison.

Note: QSU-brief scores reported on a 7-point scale. Higher scores indicate greater intensity of urge.

Appendix 15.4.4.44 and 15.3.6.11

Path: /cvn/projects/prj/development/000000106324/dev/tables/tlf_anlqsu.sas

Program Run: 27OCT14 cvn_ahedge Program Status: FINAL

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**Table 15.2.4.44 Analysis of Change from Baseline QSU-brief Questionnaire Factors and Total Score - FAS**

Variable	Time point	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	THS 2.2-CC	THS 2.2-SA
Factor 2 - Relief	Overall	n	80	41	39		
		LS Mean	-0.46	-0.23	1.26	-0.23	-1.71
		95% CI	-0.71, -0.20	-0.58, 0.12	0.90, 1.62	-0.66, 0.20	-2.15, -1.28
	Day 1	n	80	41	39		
		LS Mean	-0.22	-0.09	1.76	-0.13	-1.98
		95% CI	-0.51, 0.07	-0.50, 0.31	1.35, 2.18	-0.63, 0.37	-2.49, -1.48
	Day 2	n	80	41	39		
		LS Mean	-0.42	-0.23	1.47	-0.19	-1.89
		95% CI	-0.72, -0.11	-0.65, 0.19	1.04, 1.90	-0.71, 0.33	-2.41, -1.36
	Day 3	n	79	41	39		
		LS Mean	-0.53	-0.24	1.37	-0.29	-1.90
		95% CI	-0.82, -0.24	-0.64, 0.16	0.96, 1.78	-0.78, 0.20	-2.40, -1.41

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Adjusted least squares (LS) means and confidence intervals (CIs) from an ANCOVA model with study arm, sex, CC consumption reported at screening, day and study arm*day fitted as fixed effect factors with baseline fitted as a covariate. Day fitted as a repeated factor. Where baseline is defined as the last assessment prior to Day 1 product use for THS 2.2 and CC subjects and prior to 06:29 AM on Day 1 for SA subjects.

Note: Comparison overall time points is the main comparison.

Note: QSU-brief scores reported on a 7-point scale. Higher scores indicate greater intensity of urge.

Appendix 15.4.4.44 and 15.3.6.11

Path: /cvn/projects/prj/development/000000106324/dev/tables/tlf_anlqsu.sas

Program Run: 27OCT14 cvn_ahedge Program Status: FINAL

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**Table 15.2.4.44 Analysis of Change from Baseline QSU-brief Questionnaire Factors and Total Score - FAS**

Variable	Time point	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	THS 2.2-CC	THS 2.2-SA
Factor 2 - Relief	Day 4	n	79	41	39		
		LS Mean	-0.47	-0.25	0.87	-0.22	-1.34
		95% CI	-0.77, -0.17	-0.67, 0.17	0.44, 1.30	-0.73, 0.30	-1.86, -0.81
	Day 5	n	79	41	39		
		LS Mean	-0.65	-0.32	0.81	-0.33	-1.46
		95% CI	-0.99, -0.31	-0.79, 0.15	0.33, 1.29	-0.91, 0.25	-2.05, -0.87

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Adjusted least squares (LS) means and confidence intervals (CIs) from an ANCOVA model with study arm, sex, CC consumption reported at screening, day and study arm*day fitted as fixed effect factors with baseline fitted as a covariate. Day fitted as a repeated factor. Where baseline is defined as the last assessment prior to Day 1 product use for THS 2.2 and CC subjects and prior to 06:29 AM on Day 1 for SA subjects.

Note: Comparison overall time points is the main comparison.

Note: QSU-brief scores reported on a 7-point scale. Higher scores indicate greater intensity of urge.

Appendix 15.4.4.44 and 15.3.6.11

Path: /cvn/projects/prj/development/000000106324/dev/tables/tlf_anlqsu.sas

Program Run: 27OCT14 cvn_ahedge Program Status: FINAL

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**Table 15.2.4.45 Descriptive Statistics of MNWS Questionnaire Total Scores - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)	
			Raw value	% Change(*)	Raw value	% Change(*)	Raw value	% Change(*)
Total Score 1	Baseline	n	80		41		39	
		Mean (SD)	6.2 (5.99)		5.5 (5.15)		6.5 (6.45)	
		95% CI	4.8, 7.6		3.8, 7.1		4.3, 8.6	
		Median	4.0		5.0		5.0	
		Q25, Q75	2.0, 8.0		2.0, 7.0		2.0, 8.0	
		Min, Max	0, 28		0, 24		0, 31	
	Day 2	n	80	80	41	41	39	39
		Mean (SD)	6.0 (5.93)	29.9 (160.81)	5.0 (5.49)	0.0 (64.88)	11.1 (7.60)	188.3 (369.64)
		95% CI	4.6, 7.4	-5.9, 65.8	3.3, 6.8	-20.5, 20.6	8.6, 13.6	68.5, 308.2
		Median	4.0	0.0	3.0	-4.8	10.0	77.8
		Q25, Q75	2.0, 9.5	-43.8, 53.6	1.0, 6.0	-33.4, 16.7	6.0, 16.0	0.0, 200.0
		Min, Max	0, 28	-100, 900	0, 24	-100, 200	0, 27	-100, 2100

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: MNWS scores reported on a 5-point scale (0-4). Higher scores indicate greater intensity on that scale.

Appendix 15.3.6.12

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_mnws.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.45 Descriptive Statistics of MNWS Questionnaire Total Scores - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)	
			Raw value	% Change(*)	Raw value	% Change(*)	Raw value	% Change(*)
Total Score 1	Day 3	n	80	80	41	41	39	39
		Mean (SD)	5.2 (5.66)	2.3 (80.84)	4.8 (4.84)	1.9 (59.12)	9.7 (6.82)	159.0 (284.92)
		95% CI	3.9, 6.5	-15.7, 20.3	3.3, 6.4	-16.8, 20.6	7.5, 12.0	66.6, 251.4
		Median	3.0	0.0	3.0	0.0	8.0	54.5
		Q25, Q75	1.0, 6.0	-62.5, 36.7	1.0, 7.0	-25.0, 14.3	5.0, 14.0	0.0, 250.0
		Min, Max	0, 30	-100, 250	0, 24	-100, 175	1, 31	-59, 1400
	Day 4	n	79	79	41	41	39	39
		Mean (SD)	4.8 (4.50)	10.2 (107.29)	4.7 (4.92)	8.4 (91.03)	8.8 (7.22)	94.1 (148.30)
		95% CI	3.8, 5.9	-13.8, 34.3	3.1, 6.3	-20.4, 37.2	6.4, 11.2	46.0, 142.2
		Median	3.0	-21.4	3.0	-10.0	6.0	42.9
		Q25, Q75	2.0, 8.0	-57.2, 25.0	1.0, 6.0	-50.0, 16.7	4.0, 13.0	-12.5, 200.0
		Min, Max	0, 22	-100, 400	0, 26	-100, 400	0, 33	-100, 600

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: MNWS scores reported on a 5-point scale (0-4). Higher scores indicate greater intensity on that scale.

Appendix 15.3.6.12

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_mnws.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.45 Descriptive Statistics of MNWS Questionnaire Total Scores - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)	
			Raw value	% Change(*)	Raw value	% Change(*)	Raw value	% Change(*)
Total Score 1	Day 5	n	79	79	41	41	39	39
		Mean (SD)	4.9 (4.70)	9.3 (109.97)	5.4 (5.41)	6.0 (72.38)	8.4 (6.30)	116.1 (298.36)
		95% CI	3.8, 6.0	-15.3, 34.0	3.7, 7.2	-16.9, 28.9	6.3, 10.5	19.3, 212.9
		Median	4.0	-18.8	5.0	0.0	7.0	28.6
		Q25, Q75	1.0, 7.0	-56.0, 50.0	1.0, 8.0	-33.4, 23.1	3.0, 13.0	-22.8, 100.0
		Min, Max	0, 19	-100, 500	0, 23	-100, 200	0, 26	-100, 1700
	Day 6/ Discharge	n	79	79	41	41	39	39
		Mean (SD)	5.0 (5.66)	-5.4 (96.20)	5.7 (6.43)	15.5 (97.94)	7.4 (6.12)	90.2 (356.86)
		95% CI	3.7, 6.3	-27.0, 16.2	3.6, 7.8	-15.5, 46.5	5.3, 9.4	-25.6, 205.9
		Median	3.0	-28.6	5.0	0.0	4.0	0.0
		Q25, Q75	1.0, 8.0	-66.7, 7.7	1.0, 6.0	-30.0, 40.0	3.0, 12.0	-33.4, 100.0
		Min, Max	0, 24	-100, 320	0, 24	-100, 500	0, 22	-100, 2200

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: MNWS scores reported on a 5-point scale (0-4). Higher scores indicate greater intensity on that scale.

Appendix 15.3.6.12

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_mnws.sas

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**Table 15.2.4.45 Descriptive Statistics of MNWS Questionnaire Total Scores - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)	
			Raw value	% Change(*)	Raw value	% Change(*)	Raw value	% Change(*)
Total Score 2	Baseline	n	80		41		39	
		Mean (SD)	8.3 (7.38)		7.4 (6.12)		8.7 (8.32)	
		95% CI	6.6, 10.0		5.4, 9.4		6.0, 11.5	
		Median	6.0		6.0		6.0	
		Q25, Q75	3.0, 10.5		3.0, 9.0		3.0, 12.0	
		Min, Max	0, 31		0, 27		0, 37	
	Day 2	n	80	80	41	41	39	39
		Mean (SD)	9.2 (8.52)	32.8 (105.19)	7.4 (7.60)	1.6 (49.39)	13.1 (9.08)	168.8 (428.92)
		95% CI	7.2, 11.1	9.4, 56.3	5.0, 9.9	-14.0, 17.3	10.1, 16.1	29.7, 307.9
		Median	7.0	0.0	5.0	0.0	12.0	75.0
		Q25, Q75	3.0, 13.5	-33.4, 80.4	2.0, 10.0	-28.6, 19.1	7.0, 19.0	5.5, 150.0
		Min, Max	0, 43	-100, 450	0, 36	-100, 133	0, 35	-100, 2500

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: MNWS scores reported on a 5-point scale (0-4). Higher scores indicate greater intensity on that scale.

Appendix 15.3.6.12

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_mnws.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.45 Descriptive Statistics of MNWS Questionnaire Total Scores - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)	
			Raw value	% Change(*)	Raw value	% Change(*)	Raw value	% Change(*)
Total Score 2	Day 3	n	80	80	41	41	39	39
		Mean (SD)	8.0 (7.60)	16.7 (85.98)	6.9 (6.20)	2.7 (50.66)	11.8 (8.01)	138.5 (311.80)
		95% CI	6.2, 9.7	-2.5, 35.9	4.8, 8.9	-13.3, 18.7	9.2, 14.5	37.3, 239.6
		Median	5.0	0.0	5.0	0.0	10.0	35.3
		Q25, Q75	3.0, 11.0	-43.1, 57.2	2.0, 9.0	-33.4, 28.6	5.0, 17.0	0.0, 150.0
		Min, Max	0, 33	-93, 300	0, 29	-71, 133	1, 36	-59, 1500
	Day 4	n	79	79	41	41	39	39
		Mean (SD)	7.6 (6.53)	22.9 (98.79)	6.7 (5.83)	11.0 (103.04)	10.9 (8.66)	78.7 (155.24)
		95% CI	6.1, 9.2	0.7, 45.1	4.8, 8.6	-21.6, 43.5	8.0, 13.7	28.4, 129.1
		Median	6.0	0.0	6.0	-11.1	7.0	29.4
		Q25, Q75	3.0, 10.0	-40.8, 66.7	2.0, 9.0	-45.9, 16.7	5.0, 18.0	-16.7, 100.0
		Min, Max	0, 34	-100, 400	0, 28	-100, 500	0, 38	-100, 700

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: MNWS scores reported on a 5-point scale (0-4). Higher scores indicate greater intensity on that scale.

Appendix 15.3.6.12

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_mnws.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.45 Descriptive Statistics of MNWS Questionnaire Total Scores - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)	
			Raw value	% Change(*)	Raw value	% Change(*)	Raw value	% Change(*)
Total Score 2	Day 5	n	79	79	41	41	39	39
		Mean (SD)	7.0 (6.55)	2.1 (80.39)	7.5 (6.62)	6.8 (75.55)	10.4 (8.22)	98.9 (302.41)
		95% CI	5.5, 8.5	-16.0, 20.2	5.4, 9.7	-17.1, 30.7	7.7, 13.1	0.8, 197.0
		Median	5.0	-20.0	7.0	0.0	8.0	17.6
		Q25, Q75	2.0, 10.0	-50.0, 33.4	2.0, 11.0	-33.4, 22.3	4.0, 17.0	-21.7, 100.0
		Min, Max	0, 33	-100, 300	0, 25	-100, 300	0, 32	-100, 1800
	Day 6/ Discharge	n	79	79	41	41	39	39
		Mean (SD)	7.3 (7.51)	2.5 (105.65)	7.9 (8.08)	10.5 (84.78)	9.0 (7.84)	79.0 (358.08)
		95% CI	5.6, 9.0	-21.2, 26.2	5.3, 10.5	-16.3, 37.3	6.4, 11.6	-37.2, 195.1
		Median	5.0	-16.7	6.0	0.0	6.0	0.0
		Q25, Q75	2.0, 12.0	-50.0, 18.2	2.0, 10.0	-33.4, 33.4	4.0, 14.0	-33.4, 100.0
		Min, Max	0, 37	-100, 600	0, 34	-100, 400	0, 29	-100, 2200

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: MNWS scores reported on a 5-point scale (0-4). Higher scores indicate greater intensity on that scale.

Appendix 15.3.6.12

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_mnws.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.46 Analysis of Change from Baseline MNWS Questionnaire Total Scores - FAS**

Variable	Time point	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	THS 2.2-CC	THS 2.2-SA
Total Score 1	Overall	n	80	41	39		
		LS Mean	-1.15	-0.13	2.54	-1.01	-3.69
		95% CI	-2.07, -0.22	-1.44, 1.17	1.29, 3.79	-2.61, 0.58	-5.24, -2.13
	Day 1	n	11	5	11		
		LS Mean	-1.74	2.13	1.55	-3.87	-3.29
		95% CI	-4.38, 0.91	-1.75, 6.01	-1.29, 4.40	-8.56, 0.82	-7.17, 0.59
	Day 2	n	80	41	39		
		LS Mean	-0.17	-0.68	4.74	0.51	-4.91
		95% CI	-1.26, 0.93	-2.21, 0.85	3.18, 6.31	-1.37, 2.39	-6.82, -3.00
	Day 3	n	80	41	39		
		LS Mean	-1.01	-0.90	3.41	-0.11	-4.41
		95% CI	-2.01, 0.00	-2.31, 0.51	1.97, 4.85	-1.83, 1.62	-6.17, -2.66

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Adjusted least squares (LS) means and confidence intervals (CIs) from an ANCOVA model with study arm, sex, CC consumption reported at screening, day and study arm*day fitted as fixed effect factors with baseline fitted as a covariate. Day fitted as a repeated factor. Where baseline is defined as the last assessment prior to Day 1 product use for THS 2.2 and CC subjects and prior to 06:29 AM on Day 1 for SA subjects.

Note: Comparison overall time points is the main comparison.

Note: Comparison overall time points is the main comparison.

Note: MNWS scores reported on a 5-point scale (0-4). Higher scores indicate greater intensity on that scale.

Appendix 15.4.4.46 and 15.3.6.12

Path: /cvn/projects/prj/development/000000106324/dev/tables/tlf_anlmnws.sas

Program Run: 27OCT14 cvn_ahedge Program Status: FINAL

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**Table 15.2.4.46 Analysis of Change from Baseline MNWS Questionnaire Total Scores - FAS**

Variable	Time point	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	THS 2.2-CC	THS 2.2-SA
Total Score 1	Day 4	n	79	41	39		
		LS Mean	-1.41	-1.00	2.49	-0.41	-3.89
		95% CI	-2.34, -0.48	-2.30, 0.30	1.15, 3.82	-2.01, 1.19	-5.52, -2.27
	Day 5	n	79	41	39		
		LS Mean	-1.33	-0.31	2.02	-1.02	-3.36
		95% CI	-2.25, -0.41	-1.59, 0.97	0.71, 3.33	-2.59, 0.56	-4.96, -1.76
	Day 6	n	79	41	39		
		LS Mean	-1.24	-0.05	1.02	-1.19	-2.26
		95% CI	-2.26, -0.22	-1.47, 1.38	-0.43, 2.48	-2.95, 0.56	-4.04, -0.49

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Adjusted least squares (LS) means and confidence intervals (CIs) from an ANCOVA model with study arm, sex, CC consumption reported at screening, day and study arm*day fitted as fixed effect factors with baseline fitted as a covariate. Day fitted as a repeated factor. Where baseline is defined as the last assessment prior to Day 1 product use for THS 2.2 and CC subjects and prior to 06:29 AM on Day 1 for SA subjects.

Note: Comparison overall time points is the main comparison.

Note: Comparison overall time points is the main comparison.

Note: MNWS scores reported on a 5-point scale (0-4). Higher scores indicate greater intensity on that scale.

Appendix 15.4.4.46 and 15.3.6.12

Path: /cvn/projects/prj/development/000000106324/dev/tables/tlf_anlmnws.sas

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**Table 15.2.4.46 Analysis of Change from Baseline MNWS Questionnaire Total Scores - FAS**

Variable	Time point	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	THS 2.2-CC	THS 2.2-SA
Total Score 2	Overall	n	80	41	39		
		LS Mean	-0.52	-0.04	2.25	-0.48	-2.76
		95% CI	-1.66, 0.63	-1.65, 1.57	0.67, 3.82	-2.45, 1.50	-4.71, -0.82
	Day 1	n	11	5	11		
		LS Mean	-0.39	1.78	1.16	-2.17	-1.55
		95% CI	-3.20, 2.42	-2.34, 5.90	-1.91, 4.24	-7.15, 2.82	-5.71, 2.62
	Day 2	n	80	41	39		
		LS Mean	0.87	-0.25	4.54	1.13	-3.67
		95% CI	-0.53, 2.28	-2.21, 1.71	2.53, 6.55	-1.28, 3.53	-6.12, -1.22
	Day 3	n	80	41	39		
		LS Mean	-0.36	-0.84	3.23	0.47	-3.60
		95% CI	-1.61, 0.88	-2.58, 0.91	1.45, 5.02	-1.67, 2.62	-5.78, -1.42

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Adjusted least squares (LS) means and confidence intervals (CIs) from an ANCOVA model with study arm, sex, CC consumption reported at screening, day and study arm*day fitted as fixed effect factors with baseline fitted as a covariate. Day fitted as a repeated factor. Where baseline is defined as the last assessment prior to Day 1 product use for THS 2.2 and CC subjects and prior to 06:29 AM on Day 1 for SA subjects.

Note: Comparison overall time points is the main comparison.

Note: Comparison overall time points is the main comparison.

Note: MNWS scores reported on a 5-point scale (0-4). Higher scores indicate greater intensity on that scale.

Appendix 15.4.4.46 and 15.3.6.12

Path: /cvn/projects/prj/development/000000106324/dev/tables/tlf_anlmnws.sas

Program Run: 27OCT14 cvn_ahedge Program Status: FINAL

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**Table 15.2.4.46 Analysis of Change from Baseline MNWS Questionnaire Total Scores - FAS**

Variable	Time point	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	THS 2.2-CC	THS 2.2-SA
Total Score 2	Day 4	n	79	41	39		
		LS Mean	-0.73	-1.01	2.29	0.27	-3.02
		95% CI	-1.93, 0.46	-2.68, 0.66	0.57, 4.00	-1.78, 2.33	-5.11, -0.93
	Day 5	n	79	41	39		
		LS Mean	-1.41	-0.16	1.82	-1.25	-3.23
		95% CI	-2.59, -0.23	-1.80, 1.49	0.14, 3.51	-3.28, 0.77	-5.29, -1.17
	Day 6	n	79	41	39		
		LS Mean	-1.08	0.24	0.44	-1.31	-1.52
		95% CI	-2.31, 0.16	-1.48, 1.95	-1.32, 2.20	-3.43, 0.80	-3.67, 0.63

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Adjusted least squares (LS) means and confidence intervals (CIs) from an ANCOVA model with study arm, sex, CC consumption reported at screening, day and study arm*day fitted as fixed effect factors with baseline fitted as a covariate. Day fitted as a repeated factor. Where baseline is defined as the last assessment prior to Day 1 product use for THS 2.2 and CC subjects and prior to 06:29 AM on Day 1 for SA subjects.

Note: Comparison overall time points is the main comparison.

Note: Comparison overall time points is the main comparison.

Note: MNWS scores reported on a 5-point scale (0-4). Higher scores indicate greater intensity on that scale.

Appendix 15.4.4.46 and 15.3.6.12

Path: /cvn/projects/prj/development/000000106324/dev/tables/tlf_anlmnws.sas

Program Run: 27OCT14 cvn_ahedge Program Status: FINAL

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**Table 15.2.4.47 Descriptive Statistics of MCEQ Questionnaire Subscales - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	% Change(*)	Raw value	% Change(*)
Aversion Subscale	Baseline	n	80		41	
		Mean (SD)	1.2 (0.48)		1.5 (0.97)	
		95% CI	1.1, 1.4		1.1, 1.8	
		Median	1.0		1.0	
		Q25, Q75	1.0, 1.0		1.0, 1.5	
		Min, Max	1, 4		1, 5	
	Day 1	n	79	79	41	41
		Mean (SD)	1.7 (1.08)	40.2 (78.78)	1.3 (0.65)	0.3 (47.48)
		95% CI	1.4, 2.0	22.5, 57.9	1.0, 1.5	-14.7, 15.4
		Median	1.0	0.0	1.0	0.0
		Q25, Q75	1.0, 2.0	0.0, 50.0	1.0, 1.0	0.0, 0.0
		Min, Max	1, 6	-40, 350	1, 4	-78, 250

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.13

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_mceq.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.47 Descriptive Statistics of MCEQ Questionnaire Subscales - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	% Change(*)	Raw value	% Change(*)
Aversion Subscale	Day 2	n	79	79	41	41
		Mean (SD)	1.5 (0.91)	23.1 (65.46)	1.3 (0.75)	2.0 (44.62)
		95% CI	1.2, 1.7	8.4, 37.8	1.0, 1.6	-12.2, 16.1
		Median	1.0	0.0	1.0	0.0
		Q25, Q75	1.0, 1.5	0.0, 0.0	1.0, 1.5	0.0, 0.0
		Min, Max	1, 7	-50, 350	1, 5	-78, 200
	Day 3	n	78	78	41	41
		Mean (SD)	1.3 (0.59)	12.3 (42.23)	1.2 (0.48)	-4.6 (36.72)
		95% CI	1.1, 1.5	2.7, 21.9	1.0, 1.4	-16.2, 7.1
		Median	1.0	0.0	1.0	0.0
		Q25, Q75	1.0, 1.5	0.0, 0.0	1.0, 1.0	0.0, 0.0
		Min, Max	1, 4	-50, 150	1, 3	-78, 150

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.13

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_mceq.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.47 Descriptive Statistics of MCEQ Questionnaire Subscales - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	% Change(*)	Raw value	% Change(*)
Aversion Subscale	Day 4	n	78	78	41	41
		Mean (SD)	1.3 (0.71)	5.2 (36.66)	1.2 (0.60)	-5.8 (30.48)
		95% CI	1.1, 1.5	-3.1, 13.6	1.0, 1.5	-15.5, 3.9
		Median	1.0	0.0	1.0	0.0
		Q25, Q75	1.0, 1.0	0.0, 0.0	1.0, 1.0	0.0, 0.0
		Min, Max	1, 6	-50, 175	1, 4	-78, 100
	Day 5	n	78	78	41	41
		Mean (SD)	1.4 (0.95)	23.3 (76.30)	1.3 (0.64)	9.8 (66.84)
		95% CI	1.2, 1.7	6.0, 40.5	1.1, 1.6	-11.4, 30.9
		Median	1.0	0.0	1.0	0.0
		Q25, Q75	1.0, 1.5	0.0, 0.0	1.0, 1.5	0.0, 0.0
		Min, Max	1, 6	-60, 350	1, 4	-78, 300

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.13

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_mceq.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.47 Descriptive Statistics of MCEQ Questionnaire Subscales - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	% Change(*)	Raw value	% Change(*)
Craving Reduction Subscale	Baseline	n	80		41	
		Mean (SD)	4.9 (1.39)		5.0 (1.28)	
		95% CI	4.6, 5.3		4.6, 5.5	
		Median	5.0		5.0	
		Q25, Q75	4.0, 6.0		4.0, 6.0	
		Min, Max	1, 7		1, 7	
	Day 1	n	79	79	41	41
		Mean (SD)	3.3 (1.52)	-30.4 (27.48)	4.7 (1.69)	-4.7 (33.51)
		95% CI	2.9, 3.7	-36.6, -24.2	4.1, 5.2	-15.3, 5.9
		Median	3.0	-28.6	5.0	0.0
		Q25, Q75	2.0, 5.0	-50.0, 0.0	4.0, 6.0	-16.7, 16.7
		Min, Max	1, 7	-86, 25	1, 7	-80, 100

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.13

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_mceq.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.47 Descriptive Statistics of MCEQ Questionnaire Subscales - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	% Change(*)	Raw value	% Change(*)
Craving Reduction Subscale	Day 2	n	79	79	41	41
		Mean (SD)	3.6 (1.62)	-19.6 (57.44)	4.6 (1.65)	-2.4 (45.10)
		95% CI	3.2, 4.0	-32.5, -6.7	4.0, 5.2	-16.7, 11.9
		Median	4.0	-20.0	5.0	0.0
		Q25, Q75	2.0, 5.0	-50.0, 0.0	4.0, 6.0	-20.0, 20.0
		Min, Max	1, 7	-86, 400	1, 7	-86, 200
	Day 3	n	78	78	41	41
		Mean (SD)	3.4 (1.46)	-28.5 (28.72)	4.5 (1.69)	-9.9 (27.80)
		95% CI	3.0, 3.7	-35.0, -22.0	3.9, 5.0	-18.7, -1.0
		Median	3.5	-25.0	5.0	0.0
		Q25, Q75	2.0, 4.0	-50.0, 0.0	3.0, 6.0	-25.0, 0.0
		Min, Max	1, 7	-86, 50	1, 7	-86, 67

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.13

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_mceq.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.47 Descriptive Statistics of MCEQ Questionnaire Subscales - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	% Change(*)	Raw value	% Change(*)
Craving Reduction Subscale	Day 4	n	78	78	41	41
		Mean (SD)	3.3 (1.60)	-29.1 (42.37)	4.4 (1.78)	-8.9 (36.20)
		95% CI	2.9, 3.7	-38.7, -19.5	3.8, 5.0	-20.4, 2.6
		Median	3.0	-28.6	5.0	0.0
		Q25, Q75	2.0, 5.0	-60.0, 0.0	4.0, 5.0	-20.0, 0.0
		Min, Max	1, 7	-86, 200	1, 7	-86, 100
	Day 5	n	78	78	41	41
		Mean (SD)	3.4 (1.51)	-27.5 (31.94)	4.8 (1.70)	5.1 (72.47)
		95% CI	3.0, 3.8	-34.8, -20.3	4.2, 5.3	-17.8, 28.1
		Median	4.0	-25.0	5.0	0.0
		Q25, Q75	2.0, 5.0	-50.0, 0.0	4.0, 6.0	-20.0, 20.0
		Min, Max	1, 7	-86, 100	1, 7	-86, 400

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.13

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_mceq.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.47 Descriptive Statistics of MCEQ Questionnaire Subscales - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	% Change(*)	Raw value	% Change(*)
Enjoyment of Respiratory Tract Sensation Subscale	Baseline	n	80		41	
		Mean (SD)	3.7 (1.40)		3.4 (2.02)	
		95% CI	3.3, 4.0		2.7, 4.1	
		Median	4.0		4.0	
		Q25, Q75	3.0, 5.0		1.0, 5.0	
		Min, Max	1, 7		1, 7	
	Day 1	n	79	79	41	41
		Mean (SD)	2.2 (1.31)	-32.8 (44.03)	3.2 (1.91)	5.7 (61.59)
		95% CI	1.9, 2.5	-42.7, -22.8	2.5, 3.8	-13.8, 25.2
		Median	2.0	-40.0	3.0	0.0
		Q25, Q75	1.0, 3.0	-66.7, 0.0	1.0, 4.0	-16.7, 0.0
		Min, Max	1, 5	-83, 200	1, 7	-86, 300

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.13

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_mceq.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.47 Descriptive Statistics of MCEQ Questionnaire Subscales - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	% Change(*)	Raw value	% Change(*)
Enjoyment of Respiratory Tract Sensation Subscale	Day 2	n	79	79	41	41
		Mean (SD)	2.6 (1.43)	-24.7 (40.22)	3.3 (1.82)	19.8 (86.69)
		95% CI	2.2, 3.0	-33.7, -15.6	2.7, 3.9	-7.6, 47.2
		Median	3.0	-20.0	4.0	0.0
		Q25, Q75	1.0, 4.0	-50.0, 0.0	2.0, 4.0	-14.3, 0.0
		Min, Max	1, 6	-83, 200	1, 7	-86, 300
	Day 3	n	78	78	41	41
		Mean (SD)	2.3 (1.28)	-33.3 (33.13)	3.2 (1.74)	17.8 (91.47)
		95% CI	2.0, 2.6	-40.8, -25.8	2.6, 3.8	-11.2, 46.7
		Median	2.0	-33.3	4.0	0.0
		Q25, Q75	1.0, 4.0	-66.7, 0.0	1.0, 4.0	-20.0, 0.0
		Min, Max	1, 5	-83, 100	1, 7	-75, 300

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.13

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_mceq.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.47 Descriptive Statistics of MCEQ Questionnaire Subscales - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	% Change(*)	Raw value	% Change(*)
Enjoyment of Respiratory Tract Sensation Subscale	Day 4	n	78	78	41	41
		Mean (SD)	2.5 (1.41)	-25.4 (46.78)	3.1 (1.71)	11.4 (79.71)
		95% CI	2.1, 2.9	-36.0, -14.8	2.5, 3.7	-13.8, 36.6
		Median	2.0	-33.3	3.0	0.0
		Q25, Q75	1.0, 4.0	-60.0, 0.0	2.0, 4.0	-20.0, 0.0
		Min, Max	1, 6	-83, 200	1, 7	-60, 300
	Day 5	n	78	78	41	41
		Mean (SD)	2.5 (1.43)	-27.7 (37.72)	3.5 (1.86)	26.9 (97.54)
		95% CI	2.1, 2.9	-36.2, -19.1	2.9, 4.1	-3.9, 57.7
		Median	2.0	-25.0	4.0	0.0
		Q25, Q75	1.0, 4.0	-60.0, 0.0	2.0, 4.0	-20.0, 0.0
		Min, Max	1, 6	-83, 100	1, 7	-50, 400

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.13

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_mceq.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.47 Descriptive Statistics of MCEQ Questionnaire Subscales - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	% Change(*)	Raw value	% Change(*)
Psychological Reward Subscale	Baseline	n	80		41	
		Mean (SD)	3.6 (1.49)		3.9 (1.54)	
		95% CI	3.2, 4.0		3.3, 4.4	
		Median	3.8		4.0	
		Q25, Q75	2.6, 4.6		3.2, 4.6	
		Min, Max	1, 7		1, 7	
	Day 1	n	79	79	41	41
		Mean (SD)	2.5 (1.41)	-27.4 (26.50)	3.5 (1.58)	-7.4 (21.04)
		95% CI	2.2, 2.9	-33.4, -21.4	3.0, 4.1	-14.1, -0.7
		Median	2.2	-25.0	3.6	-5.6
		Q25, Q75	1.4, 3.6	-45.8, -9.5	2.2, 4.6	-21.5, 6.7
		Min, Max	1, 7	-77, 60	1, 7	-48, 39

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.13

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_mceq.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.47 Descriptive Statistics of MCEQ Questionnaire Subscales - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	% Change(*)	Raw value	% Change(*)
Psychological Reward Subscale	Day 2	n	79	79	41	41
		Mean (SD)	2.7 (1.56)	-25.3 (26.39)	3.6 (1.55)	-4.1 (28.68)
		95% CI	2.3, 3.0	-31.2, -19.3	3.1, 4.1	-13.2, 5.0
		Median	2.4	-19.0	3.6	-5.0
		Q25, Q75	1.2, 3.8	-43.8, -4.7	2.4, 5.0	-22.9, 13.4
		Min, Max	1, 7	-84, 23	1, 6	-53, 83
	Day 3	n	78	78	41	41
		Mean (SD)	2.5 (1.47)	-27.6 (29.29)	3.3 (1.47)	-12.6 (26.92)
		95% CI	2.2, 2.9	-34.3, -21.0	2.8, 3.8	-21.1, -4.0
		Median	2.2	-25.6	3.0	-11.4
		Q25, Q75	1.4, 3.8	-50.0, -5.5	2.2, 4.2	-31.9, 0.0
		Min, Max	1, 7	-81, 86	1, 6	-57, 67

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.13

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_mceq.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.47 Descriptive Statistics of MCEQ Questionnaire Subscales - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	% Change(*)	Raw value	% Change(*)
Psychological Reward Subscale	Day 4	n	78	78	41	41
		Mean (SD)	2.5 (1.39)	-28.7 (35.27)	3.3 (1.51)	-13.5 (23.38)
		95% CI	2.1, 2.8	-36.7, -20.7	2.7, 3.8	-20.9, -6.1
		Median	2.0	-31.3	3.2	-11.4
		Q25, Q75	1.2, 3.4	-50.0, -8.6	2.2, 4.2	-29.1, 0.0
		Min, Max	1, 6	-77, 186	1, 7	-69, 38
	Day 5	n	78	78	41	41
		Mean (SD)	2.4 (1.36)	-30.9 (27.10)	3.4 (1.56)	-9.2 (26.51)
		95% CI	2.0, 2.8	-37.1, -24.8	2.9, 4.0	-17.6, -0.8
		Median	2.0	-33.9	3.4	-11.5
		Q25, Q75	1.2, 3.6	-50.0, -4.7	2.2, 4.6	-22.9, 0.0
		Min, Max	1, 6	-77, 19	1, 7	-65, 83

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.13

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_mceq.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.47 Descriptive Statistics of MCEQ Questionnaire Subscales - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	% Change(*)	Raw value	% Change(*)
Smoking Satisfaction Subscale	Baseline	n	80		41	
		Mean (SD)	4.6 (1.06)		4.7 (1.26)	
		95% CI	4.3, 4.9		4.3, 5.2	
		Median	4.9		4.7	
		Q25, Q75	4.0, 5.3		4.0, 5.7	
		Min, Max	2, 7		2, 7	
	Day 1	n	79	79	41	41
		Mean (SD)	2.8 (1.37)	-37.1 (30.78)	4.4 (1.50)	-6.9 (18.23)
		95% CI	2.5, 3.2	-44.1, -30.2	3.9, 5.0	-12.7, -1.1
		Median	2.7	-42.5	4.3	-4.3
		Q25, Q75	1.7, 4.0	-66.0, -14.8	3.3, 5.3	-16.7, 6.0
		Min, Max	1, 6	-85, 43	1, 7	-61, 40

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.13

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_mceq.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.47 Descriptive Statistics of MCEQ Questionnaire Subscales - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	% Change(*)	Raw value	% Change(*)
Smoking Satisfaction Subscale	Day 2	n	79	79	41	41
		Mean (SD)	3.3 (1.34)	-28.4 (26.65)	4.6 (1.61)	-4.4 (22.93)
		95% CI	2.9, 3.6	-34.4, -22.4	4.0, 5.1	-11.7, 2.9
		Median	3.3	-23.5	5.0	0.0
		Q25, Q75	2.0, 4.3	-46.0, -10.0	3.3, 5.7	-15.0, 7.5
		Min, Max	1, 6	-85, 43	1, 7	-70, 43
	Day 3	n	78	78	41	41
		Mean (SD)	3.1 (1.36)	-31.5 (27.14)	4.4 (1.77)	-8.6 (25.99)
		95% CI	2.8, 3.5	-37.7, -25.4	3.8, 5.0	-16.9, -0.4
		Median	3.0	-29.1	4.7	-5.0
		Q25, Q75	2.0, 4.0	-46.6, -14.8	3.0, 6.0	-20.0, 6.4
		Min, Max	1, 6	-85, 57	1, 7	-73, 43

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.13

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_mceq.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.47 Descriptive Statistics of MCEQ Questionnaire Subscales - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	% Change(*)	Raw value	% Change(*)
Smoking Satisfaction Subscale	Day 4	n	78	78	41	41
		Mean (SD)	3.2 (1.33)	-30.0 (25.95)	4.4 (1.71)	-8.5 (25.05)
		95% CI	2.8, 3.5	-35.9, -24.1	3.8, 5.0	-16.4, -0.5
		Median	3.3	-26.0	4.3	-4.3
		Q25, Q75	2.3, 4.3	-50.0, -7.5	3.0, 5.7	-27.1, 5.3
		Min, Max	1, 6	-80, 33	1, 7	-70, 54
	Day 5	n	78	78	41	41
		Mean (SD)	3.2 (1.33)	-29.5 (29.14)	4.5 (1.73)	-6.6 (24.13)
		95% CI	2.8, 3.5	-36.1, -22.9	3.9, 5.1	-14.3, 1.1
		Median	3.3	-28.1	5.0	0.0
		Q25, Q75	2.0, 4.0	-50.0, -6.9	3.7, 6.0	-17.6, 10.6
		Min, Max	1, 6	-83, 57	1, 7	-65, 40

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.13

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_mceq.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.48 Analysis of Change from Baseline in MCEQ Questionnaire Subscales - FAS**

Subscales	Time point	Statistic	THS 2.2 (N=80)	CC (N=41)	THS 2.2-CC
Aversion Subscale	Overall	n	79	41	
		LS Mean	0.15	-0.10	0.25
		95% CI	0.02, 0.28	-0.27, 0.07	0.04, 0.46
	Day 1	n	79	41	
		LS Mean	0.40	-0.08	0.47
		95% CI	0.20, 0.60	-0.35, 0.20	0.13, 0.82
	Day 2	n	79	41	
		LS Mean	0.18	-0.05	0.24
		95% CI	0.00, 0.37	-0.30, 0.20	-0.07, 0.54
	Day 3	n	78	41	
		LS Mean	0.03	-0.17	0.20
		95% CI	-0.09, 0.15	-0.34, -0.01	0.00, 0.41

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Adjusted least squares (LS) means and confidence intervals (CIs) from an ANCOVA model with study arm, sex, CC consumption reported at screening, day and study arm*day fitted as fixed effect factors with baseline fitted as a covariate. Day fitted as a repeated factor. Where baseline is defined as the last assessment prior to Day 1 product use for THS 2.2 and CC subjects and prior to 06:29 AM on Day 1 for SA subjects.

Note: Comparison overall time points is the main comparison.

Note: MCEQ subscales reported on a 7-point scale (1-7). Higher scores indicate greater intensity on that scale.

Appendix 15.4.4.48 and 15.3.6.13

Path: /cvn/projects/prj/development/000000106324/dev/tables/tlf_anlmceq.sas

Program Run: 27OCT14 cvn_ahedge Program Status: FINAL

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**Table 15.2.4.48 Analysis of Change from Baseline in MCEQ Questionnaire Subscales - FAS**

Subscales	Time point	Statistic	THS 2.2 (N=80)	CC (N=41)	THS 2.2-CC
Aversion Subscale	Day 4	n	78	41	
		LS Mean	-0.02	-0.15	0.13
		95% CI	-0.15, 0.11	-0.33, 0.04	-0.10, 0.36
	Day 5	n	78	41	
		LS Mean	0.15	-0.05	0.20
		95% CI	-0.04, 0.35	-0.31, 0.21	-0.12, 0.53
Craving Reduction Subscale	Overall	n	79	41	
		LS Mean	-1.53	-0.41	-1.12
		95% CI	-1.81, -1.26	-0.79, -0.04	-1.58, -0.66
	Day 1	n	79	41	
		LS Mean	-1.62	-0.33	-1.29
		95% CI	-1.94, -1.30	-0.78, 0.11	-1.84, -0.73

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Adjusted least squares (LS) means and confidence intervals (CIs) from an ANCOVA model with study arm, sex, CC consumption reported at screening, day and study arm*day fitted as fixed effect factors with baseline fitted as a covariate. Day fitted as a repeated factor. Where baseline is defined as the last assessment prior to Day 1 product use for THS 2.2 and CC subjects and prior to 06:29 AM on Day 1 for SA subjects.

Note: Comparison overall time points is the main comparison.

Note: MCEQ subscales reported on a 7-point scale (1-7). Higher scores indicate greater intensity on that scale.

Appendix 15.4.4.48 and 15.3.6.13

Path: /cvn/projects/prj/development/000000106324/dev/tables/tlf_anlmceq.sas

Program Run: 27OCT14 cvn_ahedge Program Status: FINAL

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**Table 15.2.4.48 Analysis of Change from Baseline in MCEQ Questionnaire Subscales - FAS**

Subscales	Time point	Statistic	THS 2.2 (N=80)	CC (N=41)	THS 2.2-CC
Craving Reduction Subscale	Day 2	n	79	41	
		LS Mean	-1.32	-0.38	-0.93
		95% CI	-1.66, -0.97	-0.86, 0.10	-1.53, -0.34
	Day 3	n	78	41	
		LS Mean	-1.56	-0.53	-1.03
		95% CI	-1.87, -1.25	-0.96, -0.10	-1.56, -0.50
	Day 4	n	78	41	
		LS Mean	-1.67	-0.58	-1.09
		95% CI	-2.01, -1.32	-1.06, -0.10	-1.68, -0.50
	Day 5	n	78	41	
		LS Mean	-1.51	-0.24	-1.27
		95% CI	-1.86, -1.16	-0.72, 0.24	-1.87, -0.68

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Adjusted least squares (LS) means and confidence intervals (CIs) from an ANCOVA model with study arm, sex, CC consumption reported at screening, day and study arm*day fitted as fixed effect factors with baseline fitted as a covariate. Day fitted as a repeated factor. Where baseline is defined as the last assessment prior to Day 1 product use for THS 2.2 and CC subjects and prior to 06:29 AM on Day 1 for SA subjects.

Note: Comparison overall time points is the main comparison.

Note: MCEQ subscales reported on a 7-point scale (1-7). Higher scores indicate greater intensity on that scale.

Appendix 15.4.4.48 and 15.3.6.13

Path: /cvn/projects/prj/development/000000106324/dev/tables/tlf_anlmceq.sas

Program Run: 27OCT14 cvn_ahedge Program Status: FINAL

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**Table 15.2.4.48 Analysis of Change from Baseline in MCEQ Questionnaire Subscales - FAS**

Subscales	Time point	Statistic	THS 2.2 (N=80)	CC (N=41)	THS 2.2-CC
Enjoyment of Respiratory Tract Sensation Subscale	Overall	n	79	41	
		LS Mean	-1.23	-0.23	-1.00
		95% CI	-1.44, -1.01	-0.52, 0.06	-1.36, -0.64
	Day 1	n	79	41	
		LS Mean	-1.43	-0.31	-1.13
		95% CI	-1.72, -1.14	-0.71, 0.10	-1.62, -0.63
	Day 2	n	79	41	
		LS Mean	-1.04	-0.18	-0.86
		95% CI	-1.32, -0.76	-0.57, 0.21	-1.34, -0.38
	Day 3	n	78	41	
		LS Mean	-1.35	-0.28	-1.07
		95% CI	-1.60, -1.09	-0.63, 0.07	-1.50, -0.64

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Adjusted least squares (LS) means and confidence intervals (CIs) from an ANCOVA model with study arm, sex, CC consumption reported at screening, day and study arm*day fitted as fixed effect factors with baseline fitted as a covariate. Day fitted as a repeated factor. Where baseline is defined as the last assessment prior to Day 1 product use for THS 2.2 and CC subjects and prior to 06:29 AM on Day 1 for SA subjects.

Note: Comparison overall time points is the main comparison.

Note: MCEQ subscales reported on a 7-point scale (1-7). Higher scores indicate greater intensity on that scale.

Appendix 15.4.4.48 and 15.3.6.13

Path: /cvn/projects/prj/development/000000106324/dev/tables/tlf_anlmceq.sas

Program Run: 27OCT14 cvn_ahedge Program Status: FINAL

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**Table 15.2.4.48 Analysis of Change from Baseline in MCEQ Questionnaire Subscales - FAS**

Subscales	Time point	Statistic	THS 2.2 (N=80)	CC (N=41)	THS 2.2-CC
Enjoyment of Respiratory Tract Sensation Subscale	Day 4	n	78	41	
		LS Mean	-1.16	-0.38	-0.78
		95% CI	-1.44, -0.87	-0.77, 0.01	-1.26, -0.30
	Day 5	n	78	41	
		LS Mean	-1.15	0.01	-1.17
		95% CI	-1.43, -0.87	-0.37, 0.40	-1.64, -0.69
Psychological Reward Subscale	Overall	n	79	41	
		LS Mean	-1.13	-0.41	-0.72
		95% CI	-1.33, -0.93	-0.68, -0.13	-1.06, -0.39
	Day 1	n	79	41	
		LS Mean	-1.10	-0.29	-0.80
		95% CI	-1.31, -0.88	-0.59, 0.00	-1.17, -0.44

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Adjusted least squares (LS) means and confidence intervals (CIs) from an ANCOVA model with study arm, sex, CC consumption reported at screening, day and study arm*day fitted as fixed effect factors with baseline fitted as a covariate. Day fitted as a repeated factor. Where baseline is defined as the last assessment prior to Day 1 product use for THS 2.2 and CC subjects and prior to 06:29 AM on Day 1 for SA subjects.

Note: Comparison overall time points is the main comparison.

Note: MCEQ subscales reported on a 7-point scale (1-7). Higher scores indicate greater intensity on that scale.

Appendix 15.4.4.48 and 15.3.6.13

Path: /cvn/projects/prj/development/000000106324/dev/tables/tlf_anlmceq.sas

Program Run: 27OCT14 cvn_ahedge Program Status: FINAL

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**Table 15.2.4.48 Analysis of Change from Baseline in MCEQ Questionnaire Subscales - FAS**

Subscales	Time point	Statistic	THS 2.2 (N=80)	CC (N=41)	THS 2.2-CC
Psychological Reward Subscale	Day 2	n	79	41	
		LS Mean	-0.98	-0.24	-0.75
		95% CI	-1.22, -0.74	-0.57, 0.09	-1.15, -0.34
	Day 3	n	78	41	
		LS Mean	-1.12	-0.56	-0.56
		95% CI	-1.35, -0.89	-0.88, -0.24	-0.95, -0.17
	Day 4	n	78	41	
		LS Mean	-1.20	-0.55	-0.64
		95% CI	-1.43, -0.97	-0.87, -0.23	-1.04, -0.25
	Day 5	n	78	41	
		LS Mean	-1.25	-0.38	-0.87
		95% CI	-1.49, -1.01	-0.71, -0.06	-1.27, -0.47

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Adjusted least squares (LS) means and confidence intervals (CIs) from an ANCOVA model with study arm, sex, CC consumption reported at screening, day and study arm*day fitted as fixed effect factors with baseline fitted as a covariate. Day fitted as a repeated factor. Where baseline is defined as the last assessment prior to Day 1 product use for THS 2.2 and CC subjects and prior to 06:29 AM on Day 1 for SA subjects.

Note: Comparison overall time points is the main comparison.

Note: MCEQ subscales reported on a 7-point scale (1-7). Higher scores indicate greater intensity on that scale.

Appendix 15.4.4.48 and 15.3.6.13

Path: /cvn/projects/prj/development/000000106324/dev/tables/tlf_anlmceq.sas

Program Run: 27OCT14 cvn_ahedge Program Status: FINAL

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**Table 15.2.4.48 Analysis of Change from Baseline in MCEQ Questionnaire Subscales - FAS**

Subscales	Time point	Statistic	THS 2.2 (N=80)	CC (N=41)	THS 2.2-CC
Smoking Satisfaction Subscale	Overall	n	79	41	
		LS Mean	-1.51	-0.25	-1.26
		95% CI	-1.75, -1.27	-0.59, 0.09	-1.68, -0.85
	Day 1	n	79	41	
		LS Mean	-1.81	-0.27	-1.54
		95% CI	-2.09, -1.53	-0.65, 0.11	-2.01, -1.07
	Day 2	n	79	41	
		LS Mean	-1.37	-0.15	-1.22
		95% CI	-1.64, -1.10	-0.52, 0.22	-1.67, -0.76
	Day 3	n	78	41	
		LS Mean	-1.50	-0.32	-1.18
		95% CI	-1.78, -1.21	-0.71, 0.08	-1.67, -0.70

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Adjusted least squares (LS) means and confidence intervals (CIs) from an ANCOVA model with study arm, sex, CC consumption reported at screening, day and study arm*day fitted as fixed effect factors with baseline fitted as a covariate. Day fitted as a repeated factor. Where baseline is defined as the last assessment prior to Day 1 product use for THS 2.2 and CC subjects and prior to 06:29 AM on Day 1 for SA subjects.

Note: Comparison overall time points is the main comparison.

Note: MCEQ subscales reported on a 7-point scale (1-7). Higher scores indicate greater intensity on that scale.

Appendix 15.4.4.48 and 15.3.6.13

Path: /cvn/projects/prj/development/000000106324/dev/tables/tlf_anlmceq.sas

Program Run: 27OCT14 cvn_ahedge Program Status: FINAL

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**Table 15.2.4.48 Analysis of Change from Baseline in MCEQ Questionnaire Subscales - FAS**

Subscales	Time point	Statistic	THS 2.2 (N=80)	CC (N=41)	THS 2.2-CC
Smoking Satisfaction Subscale	Day 4	n	78	41	
		LS Mean	-1.43	-0.31	-1.12
		95% CI	-1.70, -1.16	-0.69, 0.07	-1.58, -0.65
	Day 5	n	78	41	
		LS Mean	-1.44	-0.19	-1.25
		95% CI	-1.73, -1.16	-0.59, 0.20	-1.73, -0.77

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Adjusted least squares (LS) means and confidence intervals (CIs) from an ANCOVA model with study arm, sex, CC consumption reported at screening, day and study arm*day fitted as fixed effect factors with baseline fitted as a covariate. Day fitted as a repeated factor. Where baseline is defined as the last assessment prior to Day 1 product use for THS 2.2 and CC subjects and prior to 06:29 AM on Day 1 for SA subjects.

Note: Comparison overall time points is the main comparison.

Note: MCEQ subscales reported on a 7-point scale (1-7). Higher scores indicate greater intensity on that scale.

Appendix 15.4.4.48 and 15.3.6.13

Path: /cvn/projects/prj/development/000000106324/dev/tables/tlf_anlmceq.sas

Program Run: 27OCT14 cvn_ahedge Program Status: FINAL

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**Table 15.2.4.49 Descriptive Statistics of CYP1A2 Activity (%) - FAS**

Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)	
		Raw value	%Change (*)	Raw value	%Change (*)	Raw value	%Change (*)
Baseline	n	80		40		38	
	BLOQ - n (%)	0		0		0	
	Mean (SD)	112.426 (36.0013)		110.260 (29.9344)		113.141 (44.8515)	
	95% CI	104.414, 120.438		100.686, 119.834		98.398, 127.884	
	Median	111.142		112.561		104.343	
	Q25, Q75	86.418, 136.004		82.416, 126.177		79.747, 136.017	
	Min, Max	32.10, 204.80		49.54, 177.59		40.02, 204.40	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.4.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_cyp1.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.4.49 Descriptive Statistics of CYP1A2 Activity (%) - FAS**

Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)	
		Raw value	%Change (*)	Raw value	%Change (*)	Raw value	%Change (*)
Day 5	n	79	79	41	40	39	38
	BLOQ - n (%)	0	0	0	0	0	0
	Mean (SD)	91.710 (29.1658)	-16.509 (17.9780)	123.009 (34.6639)	12.714 (15.6626)	94.450 (36.5886)	-14.860 (15.7985)
	95% CI	85.177, 98.243	-20.536, -12.481	112.067, 133.951	7.705, 17.724	82.589, 106.311	-20.053, -9.667
	Median	90.420	-18.491	121.947	11.200	86.408	-16.514
	Q25, Q75	75.601, 109.646	-27.788, -9.091	98.853, 147.928	1.583, 23.897	69.731, 109.195	-23.946, -4.641
	Min, Max	29.37, 186.18	-48.87, 54.20	49.98, 186.55	-20.96, 48.70	30.63, 190.61	-45.71, 26.84

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.4.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_cyp1.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.4.50 Analysis of CYP1A2 Activity (%) on Day 5 - FAS**

Parameter	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	THS - CC	THS - SA
Absolute	n	79	41	39		
	LS Mean (SE)	91.35	124.95	93.34	-33.60 (3.537)	-1.99 (3.597)
	95% CI	87.30, 95.41	119.26, 130.65	87.50, 99.18	-40.59, -26.61	-9.10, 5.12
Change from baseline	n	79	40	38		
	LS Mean (SE)	-20.81	12.79	-18.83	-33.60 (3.537)	-1.99 (3.597)
	95% CI	-24.87, -16.76	7.09, 18.49	-24.67, -12.98	-40.59, -26.61	-9.10, 5.12

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Adjusted least squares (LS) means and confidence intervals (CIs) from an ANOVA model with baseline value, study arm, sex and CC consumption reported at screening as fixed effect factors.

Baseline is defined as the last assessment prior to Day 1 product use for THS 2.2 and CC subjects and prior to 06:29 AM on Day 1 for SA subjects.

Appendix 15.4.4.50 and 15.3.4.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/tl_anlcyp.sas

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Program Run: 27OCT14 cvn_ahedge Program Status: FINAL

**Table 15.2.4.51 Descriptive Statistics of CYP2A6 Activity (%) - FAS**

Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)	
		Raw value	%Change (*)	Raw value	%Change (*)	Raw value	%Change (*)
Baseline	n	80		41		39	
	BLOQ - n (%)	0		0		0	
	Mean (SD)	41.556 (15.3968)		44.313 (15.2036)		40.874 (15.7486)	
	95% CI	38.129, 44.983		39.514, 49.112		35.769, 45.980	
	Median	40.501		40.794		38.310	
	Q25, Q75	30.930, 52.040		31.460, 55.663		29.095, 50.091	
	Min, Max	6.68, 73.00		16.00, 73.57		14.65, 79.21	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.6.15

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_cyp2.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.4.51 Descriptive Statistics of CYP2A6 Activity (%) - FAS**

Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)	
		Raw value	%Change (*)	Raw value	%Change (*)	Raw value	%Change (*)
Day 6/ Discharge	n	79	79	41	41	30	30
	BLOQ - n (%)	0	0	0	0	0	0
	Mean (SD)	26.794 (8.7966)	-32.377 (14.8052)	27.927 (8.6240)	-33.581 (20.6722)	98.652 (46.3961)	166.984 (107.3340)
	95% CI	24.823, 28.765	-35.693, -29.060	25.205, 30.650	-40.106, -27.055	81.327, 115.977	126.904, 207.064
	Median	27.583	-34.700	27.125	-39.634	92.381	150.270
	Q25, Q75	20.694, 32.152	-41.879, -23.237	20.647, 33.221	-45.197, -27.756	64.165, 127.313	86.588, 222.396
	Min, Max	7.86, 49.57	-66.98, 17.72	13.49, 51.77	-61.84, 39.05	24.57, 191.30	23.67, 456.52

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.6.15

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_cyp2.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.52 Analysis of CYP2A6 Activity (%) on Day 5 - FAS**

Parameter	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	THS - CC	THS - SA
Absolute	n	79	41	30		
	LS Mean (SE)	26.63	26.15	102.13	0.48 (3.671)	-75.50 (4.125)
	95% CI	22.40, 30.86	20.25, 32.06	95.16, 109.10	-6.78, 7.74	-83.66, -67.34
Change from baseline	n	79	41	30		
	LS Mean (SE)	-14.86	-15.34	60.64	0.48 (3.671)	-75.50 (4.125)
	95% CI	-19.09, -10.63	-21.24, -9.43	53.68, 67.61	-6.78, 7.74	-83.66, -67.34

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Adjusted least squares (LS) means and confidence intervals (CIs) from an ANOVA model with baseline value, study arm, sex and CC consumption reported at screening as fixed effect factors.

Baseline is defined as the last assessment prior to Day 1 product use for THS 2.2 and CC subjects and prior to 06:29 AM on Day 1 for SA subjects.

Appendix 15.4.4.52 and 15.3.6.11

Path: /cvn/projects/prj/development/000000106324/dev/tables/tl_anlcyp.sas

Program Run: 27OCT14 cvn_ahedge Program Status: FINAL

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**Table 15.2.4.53 Analysis of Urinary 8-epi-PGF2 Alpha on Day 5 - FAS**

Variable	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	THS : CC Ratio (%)	THS : SA Ratio (%)
Urinary Quantity over n 24 hours (ng)		79	41	39		
	Geometric LS Mean (SE) ¹	330.82	359.73	391.76	91.96 (0.054)	84.45 (0.058)
	95% CI	310.90, 352.02	330.03, 392.10	355.93, 431.18	82.67, 102.30	75.31, 94.68
Concentration Adjusted for Creatinine (pg/mg creat)	n	79	41	39		
	Geometric LS Mean (SE) ¹	230.45	244.13	249.42	94.40 (0.041)	92.40 (0.044)
	95% CI	220.01, 241.39	228.88, 260.39	232.20, 267.91	87.16, 102.23	84.84, 100.62

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Adjusted geometric least squares (LS) means and confidence intervals (CIs) from an ANCOVA model conducted on log-transformed Day 5 values with log-transformed baseline value, study arm, sex and CC consumption reported at screening as fixed effect factors.

Baseline is defined as the last assessment prior to Day 1 product use for THS 2.2 and CC subjects and prior to 06:29 AM on Day 1 for SA subjects.

Note: 1 = Standard errors presented from analysis performed on the log_e scale.

Note: Collection over 24 hours starting on the Day stated in the title.

Appendix 15.4.4.53 and 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/tl_anlsecondbio.sas

Program Run: 27OCT14 cvn_ahedge Program Status: FINAL

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**Table 15.2.4.54 Descriptive Statistics of Urinary 8-epi-PGF2 α - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Quantity Excreted over 24 hours (ng)	Baseline	n	80		41	
		BLOQ - n (%)	0		0	
		Geometric Mean (CV%)	335.13 (35.88%)		312.98 (37.03%)	
		Geometric 95% CI	310.16, 362.12		279.49, 350.47	
		Median	331.90		302.75	
		Q25, Q75	263.34, 419.70		249.10, 424.80	
		Min, Max	152.1, 864.0		148.9, 553.7	
		Mean (SD)	356.13 (131.285)		332.41 (113.585)	
		95% CI				

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.3

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_8epi.sas

Program Run: 23FEB15 cvn_jhardman Program Status: FINAL

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**Table 15.2.4.54 Descriptive Statistics of Urinary 8-epi-PGF2 α - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Quantity Excreted over 24 hours (ng) Day 5	n		79	79	41	41
	BLOQ - n (%)		0		0	
	Geometric Mean (CV%)		338.99 (49.48%)		349.57 (38.44%)	
	Geometric 95% CI		305.26, 376.45		310.92, 393.02	
	Median		356.00	5.08	360.89	12.88
	Q25, Q75		274.35, 433.13	-14.66, 27.19	268.80, 460.38	3.23, 23.19
	Min, Max		62.1, 997.0	-65.7, 89.7	160.0, 625.0	-48.1, 121.4
	Mean (SD)		373.37 (162.007)	5.09 (28.072)	372.43 (128.055)	16.07 (33.045)
	95% CI			-1.20, 11.38		5.64, 26.51

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.3

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_8epi.sas

Program Run: 23FEB15 cvn_jhardman Program Status: FINAL

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**Table 15.2.4.54 Descriptive Statistics of Urinary 8-epi-PGF2 α - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Baseline	n	80		41	
		BLOQ - n (%)	0		0	
		Geometric Mean (CV%)	209.01 (34.14%)		189.93 (31.86%)	
		Geometric 95% CI	194.12, 225.04		172.17, 209.52	
		Median	201.80		185.98	
		Q25, Q75	174.02, 255.78		159.45, 218.19	
		Min, Max	92.0, 564.2		79.2, 378.4	
		Mean (SD)	221.35 (82.458)		199.15 (64.157)	
		95% CI				

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.3

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_8epi.sas

Program Run: 23FEB15 cvn_jhardman Program Status: FINAL

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**Table 15.2.4.54 Descriptive Statistics of Urinary 8-epi-PGF2 α - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Day 5	n	79	79	41	41
		BLOQ - n (%)	0		0	
		Geometric Mean (CV%)	236.11 (33.59%)		232.76 (28.97%)	
		Geometric 95% CI	219.44, 254.05		212.81, 254.58	
		Median	237.29	12.35	238.36	23.50
		Q25, Q75	183.99, 300.51	-0.53, 25.54	190.03, 265.21	4.47, 36.67
		Min, Max	121.2, 540.4	-32.6, 112.3	126.9, 452.3	-15.9, 96.3
		Mean (SD)	249.09 (84.603)	15.38 (26.501)	242.20 (71.140)	24.71 (24.169)
		95% CI		9.44, 21.32		17.08, 32.35

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.3

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_8epi.sas

Program Run: 23FEB15 cvn_jhardman Program Status: FINAL

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**Table 15.2.4.54 Descriptive Statistics of Urinary 8-epi-PGF2 α - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Quantity Excreted over 24 hours (ng)	Baseline	n	33	
		BLOQ - n (%)	0	
		Geometric Mean (CV%)	315.85 (36.46%)	
		Geometric 95% CI	278.66, 358.01	
		Median	353.43	
		Q25, Q75	273.70, 396.15	
		Min, Max	121.2, 538.0	
		Mean (SD)	333.60 (103.035)	
		95% CI		

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.3

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_8epi.sas

Program Run: 23FEB15 cvn_jhardman Program Status: FINAL

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**Table 15.2.4.54 Descriptive Statistics of Urinary 8-epi-PGF2 α - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Quantity Excreted over 24 hours (ng)	Day 5	n	39	33
		BLOQ - n (%)	0	
		Geometric Mean (CV%)	379.54 (44.96%)	
		Geometric 95% CI	330.26, 436.17	
		Median	366.13	17.97
		Q25, Q75	276.02, 553.13	-0.46, 53.11
		Min, Max	144.0, 1162.2	-26.1, 136.5
		Mean (SD)	415.36 (188.736)	25.86 (37.901)
		95% CI		12.42, 39.31

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.3

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_8epi.sas

Program Run: 23FEB15 cvn_jhardman Program Status: FINAL

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**Table 15.2.4.54 Descriptive Statistics of Urinary 8-epi-PGF2 α - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Baseline	n	33	
		BLOQ - n (%)	0	
		Geometric Mean (CV%)	202.59 (38.73%)	
		Geometric 95% CI	177.44, 231.31	
		Median	193.55	
		Q25, Q75	157.22, 270.96	
		Min, Max	94.7, 434.0	
		Mean (SD)	216.82 (82.380)	
		95% CI		

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.3

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_8epi.sas

Program Run: 23FEB15 cvn_jhardman Program Status: FINAL

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**Table 15.2.4.54 Descriptive Statistics of Urinary 8-epi-PGF2 α - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Day 5	n	39	33
		BLOQ - n (%)	0	
		Geometric Mean (CV%)	241.95 (39.81%)	
		Geometric 95% CI	213.66, 273.98	
		Median	249.35	25.25
		Q25, Q75	185.55, 306.71	5.75, 46.44
		Min, Max	105.6, 573.1	-39.6, 141.4
		Mean (SD)	259.76 (100.875)	27.76 (35.884)
		95% CI		15.03, 40.49

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.3

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_8epi.sas

Program Run: 23FEB15 cvn_jhardman Program Status: FINAL

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**Table 15.2.4.55 Analysis of Urinary 11-DTX-B2 on Day 5 - FAS**

Variable	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	THS : CC Ratio (%)	THS : SA Ratio (%)
Urinary Quantity over n 24 hours (ng)		79	41	39		
	Geometric LS Mean (SE) ¹	727.62	895.44	717.31	81.26 (0.056)	101.44 (0.060)
	95% CI	682.91, 775.26	819.50, 978.42	650.33, 791.19	72.83, 90.66	90.19, 114.09
Concentration Adjusted for Creatinine (pg/mg creat)	n	79	41	39		
	Geometric LS Mean (SE) ¹	506.02	605.09	459.73	83.63 (0.040)	110.07 (0.043)
	95% CI	483.59, 529.50	567.83, 644.79	428.66, 493.05	77.31, 90.46	101.22, 119.70

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Adjusted geometric least squares (LS) means and confidence intervals (CIs) from an ANCOVA model conducted on log-transformed Day 5 values with log-transformed baseline value, study arm, sex and CC consumption reported at screening as fixed effect factors.

Baseline is defined as the last assessment prior to Day 1 product use for THS 2.2 and CC subjects and prior to 06:29 AM on Day 1 for SA subjects.

Note: 1 = Standard errors presented from analysis performed on the log_e scale.

Note: Collection over 24 hours starting on the Day stated in the title.

Appendix 15.4.4.55 and 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/tl_anlsecondbio.sas

Program Run: 27OCT14 cvn_ahedge Program Status: FINAL

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**Table 15.2.4.56 Descriptive Statistics of Urinary 11-DTX-B2 - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Baseline	n	80		40	
		BLOQ - n (%)	0		0	
		Geometric Mean (CV%)	536.12 (41.23%)		467.88 (50.53%)	
		Geometric 95% CI	490.87, 585.54		401.70, 544.95	
		Median	502.69		460.34	
		Q25, Q75	415.31, 721.87		354.14, 652.10	
		Min, Max	160.9, 1167.8		110.9, 1075.0	
		Mean (SD)	578.31 (228.866)		516.09 (218.169)	
		95% CI				

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.3

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_11dtx.sas

Program Run: 23FEB15 cvn_jhardman Program Status: FINAL

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**Table 15.2.4.56 Descriptive Statistics of Urinary 11-DTX-B2 - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Day 5	n	79	79	41	40
		BLOQ - n (%)	0	0	0	0
		Geometric Mean (CV%)	525.88 (31.14%)		565.18 (48.12%)	
		Geometric 95% CI	491.24, 562.97		489.36, 652.75	
		Median	534.62	-4.66	556.70	22.48
		Q25, Q75	404.67, 662.99		426.39, 873.65	
		Min, Max	244.9, 1044.6		187.3, 1400.9	
		Mean (SD)	550.37 (169.237)		621.86 (267.628)	
		95% CI			14.84, 44.90	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.3

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_11dtx.sas

Program Run: 23FEB15 cvn_jhardman Program Status: FINAL

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**Table 15.2.4.56 Descriptive Statistics of Urinary 11-DTX-B2 - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Quantity Excreted over 24 hours (ng)	Baseline	n	80		40	
		BLOQ - n (%)	0		0	
		Geometric Mean (CV%)	859.64 (35.54%)		764.90 (40.98%)	
		Geometric 95% CI	796.14, 928.22		674.36, 867.60	
		Median	863.20		812.89	
		Q25, Q75	710.81, 1069.22		664.40, 949.79	
		Min, Max	353.7, 2016.5		224.7, 2454.9	
		Mean (SD)	911.66 (323.260)		823.70 (347.875)	
		95% CI				

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.3

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_11dtx.sas

Program Run: 23FEB15 cvn_jhardman Program Status: FINAL

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**Table 15.2.4.56 Descriptive Statistics of Urinary 11-DTX-B2 - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)	
			Raw value	%Change (*)	Raw value	%Change (*)
Quantity Excreted over 24 hours (ng)	Day 5	n	79	79	41	40
		BLOQ - n (%)	0	0	0	0
		Geometric Mean (CV%)	755.03 (35.97%)		848.79 (38.86%)	
		Geometric 95% CI	698.28, 816.39		754.04, 955.45	
		Median	778.50	-14.01	873.60	3.16
		Q25, Q75	610.60, 943.20	-25.28, 4.78	747.30, 1028.50	-9.63, 35.03
		Min, Max	272.7, 1521.5	-66.2, 205.2	314.2, 2215.6	-34.0, 142.8
		Mean (SD)	798.77 (262.389)	-7.19 (36.216)	907.44 (343.764)	20.98 (47.156)
		95% CI		-15.31, 0.93		5.90, 36.07

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.3

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_11dtx.sas

Program Run: 23FEB15 cvn_jhardman Program Status: FINAL

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**Table 15.2.4.56 Descriptive Statistics of Urinary 11-DTX-B2 - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Baseline	n	33	
		BLOQ - n (%)	0	
		Geometric Mean (CV%)	468.61 (49.68%)	
		Geometric 95% CI	396.73, 553.52	
		Median	451.06	
		Q25, Q75	380.35, 616.75	
		Min, Max	118.3, 1317.3	
		Mean (SD)	518.13 (238.765)	
		95% CI		

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.3

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_11dtx.sas

Program Run: 23FEB15 cvn_jhardman Program Status: FINAL

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**Table 15.2.4.56 Descriptive Statistics of Urinary 11-DTX-B2 - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Concentration Adjusted for Creatinine (pg/mg creat)	Day 5	n	39	33
		BLOQ - n (%)	0	0
		Geometric Mean (CV%)	437.64 (42.44%)	
		Geometric 95% CI	383.55, 499.35	
		Median	428.34	-2.60
		Q25, Q75	356.73, 559.41	-15.79, 9.68
		Min, Max	151.6, 970.3	-41.8, 58.9
		Mean (SD)	473.64 (194.049)	-2.65 (23.932)
		95% CI		-11.14, 5.84

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.3

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_11dtx.sas

Program Run: 23FEB15 cvn_jhardman Program Status: FINAL

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**Table 15.2.4.56 Descriptive Statistics of Urinary 11-DTX-B2 - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Quantity Excreted over 24 hours (ng)	Baseline	n	33	
		BLOQ - n (%)	0	
		Geometric Mean (CV%)	730.60 (48.73%)	
		Geometric 95% CI	620.30, 860.51	
		Median	857.50	
		Q25, Q75	499.70, 1026.60	
		Min, Max	258.4, 1459.5	
		Mean (SD)	803.79 (335.322)	
		95% CI		

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.3

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_11dtx.sas

Program Run: 23FEB15 cvn_jhardman Program Status: FINAL

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**Table 15.2.4.56 Descriptive Statistics of Urinary 11-DTX-B2 - FAS**

Variable	Timepoint	Statistic	SA (N=39)	
			Raw value	%Change (*)
Quantity Excreted over 24 hours (ng)	Day 5	n	39	33
		BLOQ - n (%)	0	0
		Geometric Mean (CV%)	686.52 (40.51%)	
		Geometric 95% CI	605.03, 778.98	
		Median	691.56	-8.26
		Q25, Q75	549.60, 940.50	-22.75, 16.82
		Min, Max	303.4, 1477.4	-60.9, 81.3
		Mean (SD)	737.58 (280.911)	-2.42 (31.649)
		95% CI		-13.65, 8.80

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: Collection over 24 hours starting on the Day stated in the table.

Note: No BLOQ values recorded for this parameter.

Appendix 15.3.3.3

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_11dtx.sas

Program Run: 23FEB15 cvn_jhardman Program Status: FINAL

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**Table 15.2.4.57 Statistical Analysis of Relationship Between NEQ and Risk Markers on Day 5 - FAS**

Relationship	Randomization Arm	Slope	P-Value
Nicotine Equivalents (mg/g creat) vs. Prostaglandin F2 Alpha (pg/mg creat)	THS 2.2	2.84	0.127
	CC	6.19	0.014
	SA	507.12	0.038
	THS 2.2 vs. CC	-3.34	0.314
Nicotine Equivalents (mg) vs. Prostaglandin F2 Alpha (ng)	THS 2.2	9.68	<0.001
	CC	9.09	<0.001
	SA	795.44	<0.001
	THS 2.2 vs. CC	0.58	0.848

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Model for each arm separately was: Biomarker = NEQ for each arm separately. P-value for test of $R^2=0$

Model for THS 2.2 vs CC was: Biomarker = NEQ + randomization arm + NEQ*randomization arm. P-value for significance of NEQ*randomization arm. Slope estimate is the difference in slopes (THS 2.2 - CC)

Appendix 15.4.4.57

Path: /cvn/projects/prj/development/000000106324/dev/tables/tlf_anlneqrisk.sas

(Page 1 of 2)

Program Run: 31OCT14 cvn_ahedge Program Status: FINAL

**Table 15.2.4.57 Statistical Analysis of Relationship Between NEQ and Risk Markers on Day 5 - FAS**

Relationship	Randomization Arm	Slope	P-Value
Nicotine Equivalents (mg/g creat) vs. 11-Dehydro-Thromboxane B2 (pg/mg creat)	THS 2.2	12.13	<0.001
	CC	27.52	0.003
	SA	-721.77	0.130
	THS 2.2 vs. CC	-15.39	0.058
Nicotine Equivalents (mg) vs. 11-Dehydro-Thromboxane B2 (ng)	THS 2.2	16.95	<0.001
	CC	18.69	0.004
	SA	-85.58	0.789
	THS 2.2 vs. CC	-1.73	0.771

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Model for each arm separately was: Biomarker = NEQ for each arm separately. P-value for test of $R^2=0$

Model for THS 2.2 vs CC was: Biomarker = NEQ + randomization arm + NEQ*randomization arm. P-value for significance of NEQ*randomization arm. Slope estimate is the difference in slopes (THS 2.2 - CC)

Appendix 15.4.4.57

Path: /cvn/projects/prj/development/000000106324/dev/tables/tlf_anlneqrisk.sas

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Program Run: 31OCT14 cvn_ahedge Program Status: FINAL

**Table 15.2.4.58 Descriptive Statistics of HST Parameters per Cigarette - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)
			Raw Value	%Change (*)	Raw Value	%Change (*)	Raw Value
Total number of puffs (average over visit)	Baseline	n	56		27		26
		Mean (SD)	15.10 (3.711)		17.16 (4.563)		14.89 (3.561)
		95% CI	14.10, 16.10		15.35, 18.97		13.44, 16.33
		Median	15.00		16.18		14.34
		Q25, Q75	12.33, 17.10		14.12, 20.40		12.41, 16.91
		Min, Max	8.6, 24.8		10.5, 27.4		9.0, 26.7

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.7.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hst.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.58 Descriptive Statistics of HST Parameters per Cigarette - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)
			Raw Value	%Change (*)	Raw Value	%Change (*)	Raw Value
Total number of puffs (average over visit)	Day 1	n	79	55	27	27	
		Mean (SD)	16.20 (3.521)	13.94 (27.627)	17.21 (4.694)	1.33 (19.290)	
		95% CI	15.41, 16.99	6.46, 21.41	15.35, 19.07	-6.30, 8.97	
		Median	15.42	10.60	17.71	-4.55	
		Q25, Q75	13.90, 17.90	-7.65, 30.49	13.28, 20.00	-8.04, 7.30	
		Min, Max	10.7, 28.9	-34.3, 75.1	10.3, 27.5	-29.4, 77.8	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.7.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hst.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.58 Descriptive Statistics of HST Parameters per Cigarette - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)
			Raw Value	%Change (*)	Raw Value	%Change (*)	Raw Value
Total number of puffs (average over visit)	Day 4	n	78	55	27	27	
		Mean (SD)	15.78 (2.796)	10.96 (25.516)	17.76 (4.686)	4.29 (14.461)	
		95% CI	15.14, 16.41	4.06, 17.87	15.90, 19.62	-1.43, 10.02	
		Median	15.15	6.03	17.00	0.75	
		Q25, Q75	13.94, 17.00	-3.30, 29.86	14.00, 21.25	-3.49, 9.87	
		Min, Max	9.1, 23.4	-36.9, 80.2	10.9, 27.2	-14.3, 61.9	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.7.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hst.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.58 Descriptive Statistics of HST Parameters per Cigarette - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)
			Raw Value	%Change (*)	Raw Value	%Change (*)	Raw Value
Total puff volume (mL) (average over visit)	Baseline	n	56		27		26
		Mean (SD)	774.64 (167.735)		899.34 (340.992)		774.21 (148.569)
		95% CI	729.72, 819.57		764.45, 1034.24		714.20, 834.23
		Median	761.16		817.39		796.34
		Q25, Q75	654.21, 888.67		703.73, 940.98		671.94, 868.29
		Min, Max	410.0, 1196.4		479.0, 2174.6		536.1, 1166.8

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.7.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hst.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.58 Descriptive Statistics of HST Parameters per Cigarette - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)
			Raw Value	%Change (*)	Raw Value	%Change (*)	Raw Value
Total puff volume (mL) (average over visit)	Day 1	n	79	55	27	27	
		Mean (SD)	792.83 (301.144)	8.84 (36.329)	819.29 (228.514)	-6.03 (17.912)	
		95% CI	725.37, 860.28	-0.99, 18.66	728.89, 909.69	-13.12, 1.06	
		Median	784.72	7.36	853.40	-4.29	
		Q25, Q75	585.02, 984.67	-19.83, 39.49	647.66, 985.78	-11.62, 4.80	
		Min, Max	226.8, 1417.5	-55.9, 84.4	324.7, 1339.8	-60.8, 25.2	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.7.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hst.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.58 Descriptive Statistics of HST Parameters per Cigarette - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)
			Raw Value	%Change (*)	Raw Value	%Change (*)	Raw Value
Total puff volume (mL) (average over visit)	Day 4	n	78	55	27	27	
		Mean (SD)	810.55 (257.816)	10.29 (34.850)	845.81 (223.989)	-3.16 (14.094)	
		95% CI	752.42, 868.68	0.87, 19.72	757.20, 934.42	-8.74, 2.42	
		Median	834.52	3.60	833.51	1.71	
		Q25, Q75	654.66, 970.78	-15.29, 39.67	681.15, 953.18	-5.54, 5.06	
		Min, Max	160.3, 1566.9	-61.7, 96.5	353.6, 1408.7	-56.2, 11.6	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.7.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hst.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.58 Descriptive Statistics of HST Parameters per Cigarette - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)
			Raw Value	%Change (*)	Raw Value	%Change (*)	Raw Value
Average puff volume (mL) (average over visit)	Baseline	n	56		27		26
		Mean (SD)	53.46 (13.333)		53.64 (17.282)		53.38 (10.430)
		95% CI	49.89, 57.04		46.80, 60.48		49.17, 57.60
		Median	54.39		53.95		53.64
		Q25, Q75	44.03, 61.82		43.52, 61.62		45.20, 58.92
		Min, Max	23.5, 87.2		22.7, 104.7		31.6, 83.3

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.7.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hst.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.58 Descriptive Statistics of HST Parameters per Cigarette - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)
			Raw Value	%Change (*)	Raw Value	%Change (*)	Raw Value
Average puff volume (mL) (average over visit)	Day 1	n	79	55	27	27	
		Mean (SD)	50.62 (19.501)	-3.88 (24.550)	49.27 (13.947)	-6.07 (13.797)	
		95% CI	46.25, 54.99	-10.52, 2.77	43.75, 54.80	-11.53, -0.61	
		Median	53.95	-1.01	49.16	-2.92	
		Q25, Q75	35.60, 63.32	-20.72, 14.56	40.98, 57.34	-8.10, -0.17	
		Min, Max	9.9, 105.7	-66.5, 67.4	22.3, 80.6	-57.0, 14.9	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.7.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hst.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.58 Descriptive Statistics of HST Parameters per Cigarette - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)
			Raw Value	%Change (*)	Raw Value	%Change (*)	Raw Value
Average puff volume (mL) (average over visit)	Day 4	n	78	55	27	27	
		Mean (SD)	52.91 (18.467)	-0.20 (21.236)	49.41 (12.998)	-5.72 (13.392)	
		95% CI	48.74, 57.08	-5.94, 5.55	44.26, 54.55	-11.03, -0.42	
		Median	52.82	0.70	51.45	-3.93	
		Q25, Q75	43.08, 63.05	-10.91, 13.48	44.24, 56.29	-8.65, 1.65	
		Min, Max	9.9, 133.3	-53.7, 53.0	22.6, 77.6	-49.3, 17.0	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.7.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hst.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.58 Descriptive Statistics of HST Parameters per Cigarette - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)
			Raw Value	%Change (*)	Raw Value	%Change (*)	Raw Value
Average puff duration (s) (average over visit)	Baseline	n	56		27		26
		Mean (SD)	1.63 (0.439)		1.55 (0.411)		1.63 (0.440)
		95% CI	1.51, 1.75		1.39, 1.72		1.45, 1.81
		Median	1.64		1.48		1.50
		Q25, Q75	1.32, 1.91		1.21, 1.98		1.32, 1.94
		Min, Max	0.7, 2.7		0.9, 2.4		0.9, 2.8

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.7.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hst.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.4.58 Descriptive Statistics of HST Parameters per Cigarette - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)
			Raw Value	%Change (*)	Raw Value	%Change (*)	Raw Value
Average puff duration (s) Day 1 (average over visit)	n		79	55	27	27	
		Mean (SD)	2.02 (0.705)	28.00 (28.865)	1.57 (0.450)	1.54 (13.788)	
		95% CI	1.85, 2.18	20.19, 35.81	1.39, 1.75	-3.92, 7.00	
		Median	2.00	30.03	1.50	0.18	
		Q25, Q75	1.46, 2.46	7.32, 46.91	1.23, 1.88	-7.07, 6.80	
		Min, Max	0.6, 3.8	-31.9, 127.8	0.9, 2.6	-26.2, 33.4	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.7.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hst.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.58 Descriptive Statistics of HST Parameters per Cigarette - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)
			Raw Value	%Change (*)	Raw Value	%Change (*)	Raw Value
Average puff duration (s) Day 4 (average over visit)	n		78	55	27	27	
		Mean (SD)	2.12 (0.819)	34.19 (32.318)	1.55 (0.451)	0.13 (14.529)	
		95% CI	1.93, 2.31	25.45, 42.93	1.37, 1.74	-5.62, 5.88	
		Median	2.03	31.79	1.52	1.88	
		Q25, Q75	1.48, 2.65	10.07, 53.55	1.07, 2.02	-7.95, 5.60	
		Min, Max	0.6, 4.7	-27.1, 143.6	0.9, 2.4	-26.9, 44.0	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.7.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hst.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.4.58 Descriptive Statistics of HST Parameters per Cigarette - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)
			Raw Value	%Change (*)	Raw Value	%Change (*)	Raw Value
Total puff duration (s) (average over visit)	Baseline	n	56		27		26
		Mean (SD)	23.84 (6.200)		25.92 (6.747)		23.53 (5.719)
		95% CI	22.17, 25.50		23.25, 28.59		21.21, 25.84
		Median	24.74		25.00		23.74
		Q25, Q75	18.77, 28.24		21.21, 29.86		18.79, 26.58
		Min, Max	12.5, 41.1		13.8, 47.3		15.7, 38.9

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.7.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hst.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.58 Descriptive Statistics of HST Parameters per Cigarette - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)
			Raw Value	%Change (*)	Raw Value	%Change (*)	Raw Value
Total puff duration (s) (average over visit)	Day 1	n	79	55	27	27	
		Mean (SD)	32.08 (11.438)	44.99 (43.796)	26.23 (7.562)	1.93 (20.989)	
		95% CI	29.52, 34.65	33.14, 56.83	23.24, 29.23	-6.38, 10.24	
		Median	30.65	41.50	27.82	0.01	
		Q25, Q75	23.94, 39.23	12.47, 72.50	19.59, 30.79	-11.75, 14.03	
		Min, Max	9.4, 59.5	-41.5, 148.5	12.2, 41.9	-31.4, 58.2	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.7.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hst.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.4.58 Descriptive Statistics of HST Parameters per Cigarette - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)
			Raw Value	%Change (*)	Raw Value	%Change (*)	Raw Value
Total puff duration (s) (average over visit)	Day 4	n	78	55	27	27	
		Mean (SD)	33.25 (13.553)	50.42 (61.077)	26.66 (7.718)	3.00 (17.063)	
		95% CI	30.18, 36.31	33.90, 66.94	23.60, 29.72	-3.75, 9.76	
		Median	30.74	36.55	24.86	0.59	
		Q25, Q75	24.37, 41.20	15.66, 80.42	20.72, 33.30	-8.03, 14.78	
		Min, Max	9.6, 68.3	-49.2, 341.6	11.7, 43.8	-29.0, 50.1	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.7.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hst.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.58 Descriptive Statistics of HST Parameters per Cigarette - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)
			Raw Value	%Change (*)	Raw Value	%Change (*)	Raw Value
Average flow (mL/s) (average over visit)	Baseline	n	56		27		26
		Mean (SD)	34.96 (7.932)		36.18 (9.211)		35.17 (8.891)
		95% CI	32.83, 37.09		32.53, 39.83		31.57, 38.76
		Median	33.88		35.39		34.55
		Q25, Q75	30.36, 38.11		30.62, 44.15		28.76, 41.53
		Min, Max	16.1, 57.1		19.3, 62.1		20.2, 51.5

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.7.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hst.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.4.58 Descriptive Statistics of HST Parameters per Cigarette - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)
			Raw Value	%Change (*)	Raw Value	%Change (*)	Raw Value
Average flow (mL/s) (average over visit)	Day 1	n	79	55	27	27	
		Mean (SD)	26.30 (6.914)	-23.66 (13.698)	33.29 (8.468)	-7.16 (12.085)	
		95% CI	24.75, 27.86	-27.36, -19.95	29.94, 36.65	-11.94, -2.37	
		Median	25.40	-22.55	33.97	-4.66	
		Q25, Q75	22.51, 31.10	-31.18, -14.04	28.28, 36.49	-12.80, -0.44	
		Min, Max	9.5, 41.9	-59.8, -1.0	16.9, 49.9	-54.8, 8.0	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.7.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hst.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.4.58 Descriptive Statistics of HST Parameters per Cigarette - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)
			Raw Value	%Change (*)	Raw Value	%Change (*)	Raw Value
Average flow (mL/s) (average over visit)	Day 4	n	78	55	27	27	
		Mean (SD)	27.15 (7.237)	-22.99 (15.597)	33.99 (8.005)	-4.57 (13.621)	
		95% CI	25.51, 28.79	-27.22, -18.77	30.82, 37.16	-9.96, 0.82	
		Median	26.32	-21.81	35.15	-1.80	
		Q25, Q75	21.94, 32.44	-31.87, -12.83	27.58, 37.80	-10.46, 2.78	
		Min, Max	15.2, 45.9	-62.3, 12.1	19.7, 50.9	-42.6, 13.7	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.7.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hst.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.4.58 Descriptive Statistics of HST Parameters per Cigarette - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)
			Raw Value	%Change (*)	Raw Value	%Change (*)	Raw Value
Average peak flow (mL/s) (average over visit)	Baseline	n	56		27		26
		Mean (SD)	55.28 (13.900)		52.42 (12.610)		51.70 (13.558)
		95% CI	51.55, 59.01		47.42, 57.41		46.22, 57.18
		Median	54.04		52.47		52.30
		Q25, Q75	45.23, 62.81		45.17, 62.48		39.56, 63.61
		Min, Max	23.5, 89.6		26.3, 79.6		27.6, 75.1

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.7.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hst.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.4.58 Descriptive Statistics of HST Parameters per Cigarette - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)
			Raw Value	%Change (*)	Raw Value	%Change (*)	Raw Value
Average peak flow (mL/s) (average over visit)	Day 1	n	79	55	27	27	
		Mean (SD)	41.14 (12.253)	-23.57 (13.989)	47.92 (11.701)	-7.90 (11.370)	
		95% CI	38.39, 43.89	-27.35, -19.78	43.29, 52.56	-12.40, -3.40	
		Median	41.03	-22.27	46.31	-5.57	
		Q25, Q75	33.20, 48.22	-33.49, -13.37	41.37, 56.75	-13.51, -0.77	
		Min, Max	14.2, 81.1	-59.4, 6.3	24.0, 66.7	-50.6, 7.7	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.7.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hst.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.4.58 Descriptive Statistics of HST Parameters per Cigarette - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)
			Raw Value	%Change (*)	Raw Value	%Change (*)	Raw Value
Average peak flow (mL/s) (average over visit)	Day 4	n	78	55	27	27	
		Mean (SD)	41.89 (11.565)	-24.01 (15.686)	49.13 (11.895)	-5.33 (12.714)	
		95% CI	39.28, 44.50	-28.25, -19.76	44.42, 53.84	-10.36, -0.29	
		Median	39.60	-22.83	48.48	-4.41	
		Q25, Q75	34.38, 49.12	-34.53, -11.31	42.14, 58.33	-8.90, 1.56	
		Min, Max	21.9, 71.2	-55.5, 14.7	24.2, 70.5	-39.2, 12.7	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.7.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hst.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.58 Descriptive Statistics of HST Parameters per Cigarette - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)
			Raw Value	%Change (*)	Raw Value	%Change (*)	Raw Value
Total inter puff interval (s) (average over visit)	Baseline	n	56		27		26
		Mean (SD)	245.30 (52.982)		226.27 (56.980)		225.72 (44.128)
		95% CI	231.11, 259.50		203.73, 248.82		207.90, 243.55
		Median	238.54		228.42		218.35
		Q25, Q75	212.60, 285.24		188.35, 265.39		195.84, 255.62
		Min, Max	128.5, 362.9		115.0, 338.2		145.0, 312.8

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.7.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hst.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.4.58 Descriptive Statistics of HST Parameters per Cigarette - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)
			Raw Value	%Change (*)	Raw Value	%Change (*)	Raw Value
Total inter puff interval (s) (average over visit)	Day 1	n	79	55	27	27	
		Mean (SD)	177.62 (49.826)	-25.79 (18.944)	227.35 (58.145)	3.21 (27.612)	
		95% CI	166.45, 188.78	-30.92, -20.67	204.34, 250.36	-7.72, 14.14	
		Median	176.73	-26.50	238.83	1.90	
		Q25, Q75	141.01, 201.66	-39.33, -13.02	179.78, 271.74	-12.81, 8.55	
		Min, Max	72.5, 290.5	-69.5, 31.9	109.9, 318.4	-48.4, 106.6	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.7.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hst.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.58 Descriptive Statistics of HST Parameters per Cigarette - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)
			Raw Value	%Change (*)	Raw Value	%Change (*)	Raw Value
Total inter puff interval (s) (average over visit)	Day 4	n	78	55	27	27	
		Mean (SD)	167.51 (51.782)	-30.30 (18.273)	222.46 (57.143)	1.52 (30.880)	
		95% CI	155.83, 179.19	-35.25, -25.36	199.86, 245.07	-10.70, 13.74	
		Median	160.36	-32.19	231.14	-1.79	
		Q25, Q75	128.84, 206.91	-43.08, -20.95	175.07, 264.72	-16.34, 13.35	
		Min, Max	85.4, 291.9	-66.8, 17.3	92.2, 329.9	-40.6, 122.0	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.7.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hst.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.58 Descriptive Statistics of HST Parameters per Cigarette - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)
			Raw Value	%Change (*)	Raw Value	%Change (*)	Raw Value
Average inter puff interval (s) (average over visit)	Baseline	n	56		27		26
		Mean (SD)	17.44 (5.670)		14.03 (4.548)		16.04 (4.196)
		95% CI	15.91, 18.96		12.23, 15.83		14.34, 17.74
		Median	16.53		13.16		16.44
		Q25, Q75	14.05, 20.38		11.05, 17.72		12.42, 18.47
		Min, Max	7.0, 35.8		4.7, 23.1		8.6, 24.3

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.7.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hst.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.4.58 Descriptive Statistics of HST Parameters per Cigarette - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)
			Raw Value	%Change (*)	Raw Value	%Change (*)	Raw Value
Average inter puff interval (s) (average over visit)	Day 1	n	79	55	27	27	
		Mean (SD)	11.57 (4.449)	-32.35 (20.908)	14.59 (6.497)	3.84 (26.314)	
		95% CI	10.57, 12.57	-38.01, -26.69	12.02, 17.17	-6.58, 14.25	
		Median	10.36	-34.13	12.78	-1.69	
		Q25, Q75	8.92, 13.63	-51.10, -20.07	10.05, 19.36	-14.78, 21.60	
		Min, Max	5.0, 26.4	-65.3, 18.2	5.3, 30.5	-59.3, 53.6	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.7.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hst.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.4.58 Descriptive Statistics of HST Parameters per Cigarette - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)
			Raw Value	%Change (*)	Raw Value	%Change (*)	Raw Value
Average inter puff interval (s) (average over visit)	Day 4	n	78	55	27	27	
		Mean (SD)	11.16 (4.735)	-35.62 (17.339)	13.81 (5.582)	-1.28 (25.970)	
		95% CI	10.09, 12.23	-40.31, -30.93	11.60, 16.02	-11.55, 9.00	
		Median	10.19	-34.01	13.00	-4.99	
		Q25, Q75	8.37, 12.61	-48.92, -27.35	9.63, 19.21	-20.22, 17.94	
		Min, Max	5.0, 31.0	-67.2, 11.8	4.9, 23.4	-49.6, 57.7	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.7.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hst.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.4.58 Descriptive Statistics of HST Parameters per Cigarette - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)
			Raw Value	%Change (*)	Raw Value	%Change (*)	Raw Value
Total smoking duration (s) (average over visit)	Baseline	n	56		27		26
		Mean (SD)	269.14 (52.732)		252.19 (56.340)		249.25 (41.132)
		95% CI	255.01, 283.27		229.90, 274.49		232.63, 265.87
		Median	261.81		253.42		241.39
		Q25, Q75	238.72, 306.03		215.44, 285.53		222.42, 274.48
		Min, Max	154.5, 382.5		145.0, 364.9		181.4, 337.4

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.7.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hst.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.4.58 Descriptive Statistics of HST Parameters per Cigarette - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)
			Raw Value	%Change (*)	Raw Value	%Change (*)	Raw Value
Total smoking duration (s) (average over visit)	Day 1	n	79	55	27	27	
		Mean (SD)	209.70 (47.256)	-19.42 (18.195)	253.58 (56.145)	2.63 (23.092)	
		95% CI	199.11, 220.29	-24.35, -14.50	231.37, 275.80	-6.51, 11.77	
		Median	211.64	-19.12	257.41	0.29	
		Q25, Q75	171.56, 235.96	-33.50, -9.75	209.38, 306.90	-10.47, 8.29	
		Min, Max	109.4, 314.4	-64.5, 38.2	140.4, 347.3	-40.8, 90.3	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.7.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hst.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.4.58 Descriptive Statistics of HST Parameters per Cigarette - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)
			Raw Value	%Change (*)	Raw Value	%Change (*)	Raw Value
Total smoking duration (s) (average over visit)	Day 4	n	78	55	27	27	
		Mean (SD)	200.75 (52.523)	-23.12 (18.979)	249.13 (55.246)	1.32 (25.970)	
		95% CI	188.91, 212.60	-28.25, -17.98	227.27, 270.99	-8.96, 11.59	
		Median	194.28	-26.95	256.75	-1.80	
		Q25, Q75	158.83, 246.36	-33.69, -11.19	207.46, 285.58	-12.27, 9.69	
		Min, Max	106.3, 323.4	-57.9, 25.1	125.7, 362.6	-36.3, 101.9	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.7.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hst.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.4.58 Descriptive Statistics of HST Parameters per Cigarette - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)
			Raw Value	%Change (*)	Raw Value	%Change (*)	Raw Value
Total work (mJ) (average Baseline over visit)		n	56		27		26
		Mean (SD)	2130.08 (793.226)		2248.07 (914.749)		2045.53 (817.913)
		95% CI	1917.65, 2342.52		1886.21, 2609.94		1715.17, 2375.90
		Median	2007.93		2160.31		1976.09
		Q25, Q75	1570.45, 2679.62		1715.20, 2660.32		1335.91, 2597.29
		Min, Max	588.7, 4187.9		579.5, 4920.3		787.0, 4067.6

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.7.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hst.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.4.58 Descriptive Statistics of HST Parameters per Cigarette - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)
			Raw Value	%Change (*)	Raw Value	%Change (*)	Raw Value
Total work (mJ) (average Day 1 over visit)		n	79	55	27	27	
		Mean (SD)	2105.79 (1145.728)	7.47 (48.092)	2067.75 (917.941)	-9.00 (15.658)	
		95% CI	1849.15, 2362.42	-5.53, 20.48	1704.62, 2430.88	-15.20, -2.80	
		Median	1922.50	-3.71	1966.41	-7.42	
		Q25, Q75	1187.36, 2710.44	-29.69, 40.71	1563.40, 2552.69	-13.33, -0.86	
		Min, Max	196.4, 5721.8	-79.1, 140.1	369.5, 4614.1	-53.7, 26.8	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.7.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hst.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.4.58 Descriptive Statistics of HST Parameters per Cigarette - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)
			Raw Value	%Change (*)	Raw Value	%Change (*)	Raw Value
Total work (mJ) (average Day 4 over visit)		n	78	55	27	27	
		Mean (SD)	2066.36 (1005.551)	1.76 (43.450)	2161.59 (1064.959)	-4.37 (17.048)	
		95% CI	1839.64, 2293.08	-10.00, 13.51	1740.30, 2582.87	-11.12, 2.38	
		Median	1814.76	-5.69	1952.93	-7.93	
		Q25, Q75	1487.46, 2635.42	-29.62, 26.92	1607.75, 2446.20	-14.64, 6.91	
		Min, Max	340.0, 5461.4	-65.6, 131.1	481.4, 6156.5	-47.5, 31.2	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.7.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hst.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.4.58 Descriptive Statistics of HST Parameters per Cigarette - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)
			Raw Value	%Change (*)	Raw Value	%Change (*)	Raw Value
Average work (mJ) (average over visit)	Baseline	n	56		27		26
		Mean (SD)	148.71 (60.732)		137.42 (58.704)		141.74 (58.976)
		95% CI	132.44, 164.98		114.20, 160.65		117.91, 165.56
		Median	141.01		130.48		130.06
		Q25, Q75	101.88, 195.55		94.78, 197.77		101.60, 183.81
		Min, Max	23.9, 308.9		40.0, 233.7		52.1, 277.5

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.7.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hst.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.58 Descriptive Statistics of HST Parameters per Cigarette - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)
			Raw Value	%Change (*)	Raw Value	%Change (*)	Raw Value
Average work (mJ) (average over visit)	Day 1	n	79	55	27	27	
		Mean (SD)	135.27 (74.469)	-5.42 (38.147)	126.25 (58.199)	-8.62 (14.908)	
		95% CI	118.59, 151.96	-15.74, 4.90	103.22, 149.28	-14.52, -2.71	
		Median	128.98	-7.88	111.16	-2.12	
		Q25, Q75	86.40, 173.74	-31.18, 16.06	81.46, 183.16	-16.20, 0.92	
		Min, Max	9.9, 358.5	-86.0, 120.1	35.0, 235.4	-49.3, 9.6	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.7.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hst.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.4.58 Descriptive Statistics of HST Parameters per Cigarette - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)
			Raw Value	%Change (*)	Raw Value	%Change (*)	Raw Value
Average work (mJ) (average over visit)	Day 4	n	78	55	27	27	
		Mean (SD)	137.64 (74.690)	-6.40 (35.773)	127.15 (57.847)	-6.90 (17.621)	
		95% CI	120.79, 154.48	-16.07, 3.28	104.26, 150.04	-13.88, 0.07	
		Median	122.95	-13.34	119.93	-5.84	
		Q25, Q75	93.42, 173.98	-33.04, 15.47	79.06, 175.55	-18.42, 4.92	
		Min, Max	20.9, 477.2	-65.7, 98.1	39.4, 231.1	-39.4, 37.3	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.7.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hst.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.4.58 Descriptive Statistics of HST Parameters per Cigarette - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)
			Raw Value	%Change (*)	Raw Value	%Change (*)	Raw Value
Average pressure drop (mmWg) (average over visit)	Baseline	n	56		27		26
		Mean (SD)	230.01 (64.847)		227.24 (70.203)		229.21 (73.092)
		95% CI	212.64, 247.38		199.47, 255.02		199.68, 258.74
		Median	222.99		221.28		228.25
		Q25, Q75	189.74, 264.56		182.21, 266.58		203.91, 264.15
		Min, Max	88.6, 383.9		95.6, 385.1		100.4, 415.8

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.7.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hst.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.4.58 Descriptive Statistics of HST Parameters per Cigarette - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)
			Raw Value	%Change (*)	Raw Value	%Change (*)	Raw Value
Average pressure drop (mmWg) (average over visit)	Day 1	n	79	55	27	27	
		Mean (SD)	223.55 (80.360)	-2.02 (25.409)	221.61 (75.044)	-2.74 (9.887)	
		95% CI	205.54, 241.55	-8.89, 4.85	191.91, 251.30	-6.65, 1.18	
		Median	213.22	-5.04	215.05	0.55	
		Q25, Q75	171.62, 274.80	-19.43, 20.43	172.40, 272.01	-8.56, 5.66	
		Min, Max	61.6, 469.0	-59.4, 48.8	98.1, 406.8	-27.0, 8.3	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.7.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hst.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.4.58 Descriptive Statistics of HST Parameters per Cigarette - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)
			Raw Value	%Change (*)	Raw Value	%Change (*)	Raw Value
Average pressure drop (mmWg) (average over visit)	Day 4	n	78	55	27	27	
		Mean (SD)	222.22 (80.821)	-5.39 (25.114)	225.16 (78.437)	-0.83 (13.280)	
		95% CI	204.00, 240.45	-12.18, 1.41	194.13, 256.19	-6.09, 4.43	
		Median	203.90	-7.48	229.26	-0.52	
		Q25, Q75	162.66, 275.92	-24.92, 9.05	157.32, 263.32	-9.89, 9.86	
		Min, Max	91.9, 479.6	-64.5, 50.5	108.8, 394.2	-23.8, 23.9	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.7.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hst.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.4.58 Descriptive Statistics of HST Parameters per Cigarette - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)
			Raw Value	%Change (*)	Raw Value	%Change (*)	Raw Value
Average peak pressure drop (mmWg) (average over visit)	Baseline	n	56		27		26
		Mean (SD)	395.98 (127.048)		355.22 (108.476)		361.19 (117.428)
		95% CI	361.95, 430.01		312.31, 398.14		313.76, 408.63
		Median	384.29		352.85		353.03
		Q25, Q75	306.43, 478.19		283.41, 462.45		277.26, 451.08
		Min, Max	135.7, 702.0		161.3, 538.3		141.0, 650.9

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.7.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hst.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.58 Descriptive Statistics of HST Parameters per Cigarette - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)
			Raw Value	%Change (*)	Raw Value	%Change (*)	Raw Value
Average peak pressure drop (mmWg) (average over visit)	Day 1	n	79	55	27	27	
		Mean (SD)	380.34 (151.781)	-1.35 (26.532)	341.43 (114.576)	-4.13 (10.839)	
		95% CI	346.34, 414.35	-8.53, 5.82	296.10, 386.75	-8.42, 0.16	
		Median	355.99	-2.92	321.62	-0.62	
		Q25, Q75	283.18, 447.02	-20.29, 19.33	265.30, 441.56	-11.94, 3.60	
		Min, Max	87.3, 827.2	-60.0, 55.3	145.6, 549.9	-35.6, 10.0	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.7.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hst.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.4.58 Descriptive Statistics of HST Parameters per Cigarette - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)
			Raw Value	%Change (*)	Raw Value	%Change (*)	Raw Value
Average peak pressure drop (mmWg) (average over visit)	Day 4	n	78	55	27	27	
		Mean (SD)	374.23 (147.130)	-5.76 (26.835)	348.91 (122.333)	-2.01 (13.420)	
		95% CI	341.05, 407.41	-13.02, 1.50	300.51, 397.31	-7.32, 3.30	
		Median	343.52	-10.24	336.33	0.48	
		Q25, Q75	267.27, 456.96	-25.42, 7.31	256.72, 433.51	-12.04, 8.68	
		Min, Max	136.7, 855.0	-64.5, 50.4	144.6, 584.4	-24.8, 22.2	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.7.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hst.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.4.58 Descriptive Statistics of HST Parameters per Cigarette - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)
			Raw Value	%Change (*)	Raw Value	%Change (*)	Raw Value
Smoking intensity (mL/s) (average over visit)	Baseline	n	56		27		26
		Mean (SD)	3.12 (1.090)		3.97 (2.196)		3.25 (0.745)
		95% CI	2.82, 3.41		3.10, 4.85		2.94, 3.56
		Median	2.91		3.27		3.32
		Q25, Q75	2.35, 3.79		2.60, 4.48		2.62, 3.70
		Min, Max	1.3, 6.3		1.7, 10.5		1.9, 5.0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.7.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hst.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.4.58 Descriptive Statistics of HST Parameters per Cigarette - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)
			Raw Value	%Change (*)	Raw Value	%Change (*)	Raw Value
Smoking intensity (mL/s) Day 1 (average over visit)		n	79	55	27	27	
		Mean (SD)	4.23 (1.849)	43.10 (54.100)	3.75 (1.889)	-0.56 (33.729)	
		95% CI	3.81, 4.65	28.47, 57.73	3.00, 4.51	-13.91, 12.79	
		Median	4.23	42.63	3.12	-5.27	
		Q25, Q75	2.79, 5.86	7.80, 72.39	2.63, 4.20	-17.76, 13.69	
		Min, Max	0.7, 7.3	-50.6, 231.9	1.1, 8.6	-65.3, 112.9	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.7.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hst.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.58 Descriptive Statistics of HST Parameters per Cigarette - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)
			Raw Value	%Change (*)	Raw Value	%Change (*)	Raw Value
Smoking intensity (mL/s) Day 4 (average over visit)		n	78	55	27	27	
		Mean (SD)	4.57 (1.788)	52.41 (49.762)	3.94 (2.099)	5.60 (41.004)	
		95% CI	4.16, 4.98	38.95, 65.87	3.11, 4.78	-10.62, 21.83	
		Median	4.54	46.03	3.34	5.37	
		Q25, Q75	3.46, 5.80	17.32, 77.74	2.46, 4.42	-20.84, 18.15	
		Min, Max	0.8, 9.1	-49.6, 203.2	1.6, 9.6	-64.3, 169.3	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.7.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hst.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.4.58 Descriptive Statistics of HST Parameters per Cigarette - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)
			Raw Value	%Change (*)	Raw Value	%Change (*)	Raw Value
Puffing time index (%) (average over visit)	Baseline	n	56		27		26
		Mean (SD)	9.53 (3.325)		11.10 (3.992)		10.13 (3.722)
		95% CI	8.64, 10.43		9.51, 12.68		8.63, 11.64
		Median	9.68		10.44		9.71
		Q25, Q75	7.01, 11.43		7.51, 13.68		7.18, 12.03
		Min, Max	4.4, 19.3		5.4, 21.1		5.7, 20.3

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.7.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hst.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.4.58 Descriptive Statistics of HST Parameters per Cigarette - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)
			Raw Value	%Change (*)	Raw Value	%Change (*)	Raw Value
Puffing time index (%) (average over visit)	Day 1	n	79	55	27	27	
		Mean (SD)	16.78 (6.955)	88.63 (64.725)	11.85 (5.268)	8.60 (36.111)	
		95% CI	15.21, 18.34	71.13, 106.14	9.77, 13.94	-5.69, 22.89	
		Median	15.96	82.19	11.30	3.10	
		Q25, Q75	11.52, 20.82	39.19, 126.11	7.45, 14.75	-11.60, 15.31	
		Min, Max	3.8, 34.3	-26.7, 280.7	5.1, 27.1	-38.7, 118.4	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.7.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hst.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.58 Descriptive Statistics of HST Parameters per Cigarette - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)
			Raw Value	%Change (*)	Raw Value	%Change (*)	Raw Value
Puffing time index (%) (average over visit)	Day 4	n	78	55	27	27	
		Mean (SD)	18.12 (7.582)	102.63 (66.290)	12.42 (6.339)	12.49 (43.166)	
		95% CI	16.40, 19.83	84.70, 120.55	9.90, 14.93	-4.59, 29.57	
		Median	17.75	86.77	9.35	2.72	
		Q25, Q75	12.96, 22.20	56.67, 150.80	7.82, 16.70	-16.20, 28.10	
		Min, Max	3.5, 43.5	-33.3, 312.8	4.6, 29.0	-37.6, 179.0	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.7.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hst.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.58 Descriptive Statistics of HST Parameters per Cigarette - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)
			Raw Value	%Change (*)	Raw Value	%Change (*)	Raw Value
Puff frequency (puffs/min) (average over visit)	Baseline	n	56		27		26
		Mean (SD)	3.57 (1.065)		4.44 (1.729)		3.72 (0.909)
		95% CI	3.28, 3.86		3.75, 5.12		3.35, 4.09
		Median	3.49		4.20		3.46
		Q25, Q75	2.84, 4.12		3.22, 5.06		3.15, 4.59
		Min, Max	1.7, 7.0		2.4, 10.6		2.4, 5.7

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.7.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hst.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.4.58 Descriptive Statistics of HST Parameters per Cigarette - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)
			Raw Value	%Change (*)	Raw Value	%Change (*)	Raw Value
Puff frequency (puffs/min) (average over visit)	Day 1	n	79	55	27	27	
		Mean (SD)	5.04 (1.451)	47.49 (38.599)	4.60 (1.745)	5.10 (27.099)	
		95% CI	4.71, 5.37	37.05, 57.93	3.90, 5.29	-5.62, 15.83	
		Median	5.05	44.54	4.50	5.00	
		Q25, Q75	4.05, 5.94	18.28, 73.54	3.04, 5.55	-12.23, 13.90	
		Min, Max	2.2, 9.2	-18.5, 130.8	2.0, 9.1	-31.2, 109.4	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.7.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hst.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.4.58 Descriptive Statistics of HST Parameters per Cigarette - FAS**

Variable	Timepoint	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)
			Raw Value	%Change (*)	Raw Value	%Change (*)	Raw Value
Puff frequency (puffs/min) (average over visit)	Day 4	n	78	55	27	27	
		Mean (SD)	5.22 (1.419)	51.45 (36.433)	4.85 (2.142)	9.68 (28.206)	
		95% CI	4.90, 5.55	41.59, 61.30	4.00, 5.70	-1.48, 20.84	
		Median	5.21	44.52	4.20	6.81	
		Q25, Q75	4.39, 6.22	32.83, 60.47	3.28, 6.10	-9.10, 23.83	
		Min, Max	1.9, 8.4	-10.7, 146.3	2.5, 11.2	-30.7, 115.7	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: * % change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.7.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hst.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.4.59 Analysis of HST Parameters (Averaged over all Cigarettes per Day) - FAS**

Variable (units)	Timepoint	Statistic	THS 2.2 (N=80)	CC (N=41)	THS : CC Ratio (%)
Total number of puffs	Day 1	n	55	27	
		Geometric LS Mean (CV%)	16.53	16.04	103.06 (18.49)
		95% CI	15.71, 17.40	14.88, 17.29	94.26, 112.70
	Day 4	n	55	27	
		Geometric LS Mean (CV%)	16.06	16.53	97.14 (17.08)
		95% CI	15.31, 16.84	15.42, 17.72	89.42, 105.53

Note: CC = Conventional cigarettes; THS = Tobacco Heating System.

Note: Adjusted geometric least squares (LS) means and confidence intervals (CIs) from an ANCOVA model conducted on log-transformed values with log-transformed baseline value, study arm, sex and CC consumption reported at screening as fixed effect factors. Geometrical CV% of the ratio is estimated from the residual mean squares.

Baseline is defined as the last assessment prior to Day 1 product use

Appendix 15.4.4.59

Path: /cvn/projects/prj/development/000000106324/dev/tables/tl_anlhst.sas

Program Run: 27OCT14 cvn_ahedge Program Status: FINAL

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**Table 15.2.4.59 Analysis of HST Parameters (Averaged over all Cigarettes per Day) - FAS**

Variable (units)	Timepoint	Statistic	THS 2.2 (N=80)	CC (N=41)	THS : CC Ratio (%)
Total puff volume (mL)	Day 1	n	55	27	
		Geometric LS Mean (CV%)	766.65	699.95	109.53 (30.67)
		95% CI	705.12, 833.55	620.39, 789.71	94.73, 126.64
	Day 4	n	55	27	
		Geometric LS Mean (CV%)	791.70	752.16	105.26 (27.10)
		95% CI	735.20, 852.54	675.90, 837.03	92.59, 119.65

Note: CC = Conventional cigarettes; THS = Tobacco Heating System.

Note: Adjusted geometric least squares (LS) means and confidence intervals (CIs) from an ANCOVA model conducted on log-transformed values with log-transformed baseline value, study arm, sex and CC consumption reported at screening as fixed effect factors. Geometrical CV% of the ratio is estimated from the residual mean squares.

Baseline is defined as the last assessment prior to Day 1 product use

Appendix 15.4.4.59

Path: /cvn/projects/prj/development/000000106324/dev/tables/tl_anlhst.sas

Program Run: 27OCT14 cvn_ahedge Program Status: FINAL

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**Table 15.2.4.59 Analysis of HST Parameters (Averaged over all Cigarettes per Day) - FAS**

Variable (units)	Timepoint	Statistic	THS 2.2 (N=80)	CC (N=41)	THS : CC Ratio (%)
Average puff volume (mL)	Day 1	n	55	27	
		Geometric LS Mean (CV%)	46.29	45.59	101.55 (24.54)
		95% CI	43.30, 49.50	41.42, 50.18	90.61, 113.80
	Day 4	n	55	27	
		Geometric LS Mean (CV%)	49.44	46.87	105.50 (21.22)
		95% CI	46.66, 52.40	43.12, 50.94	95.55, 116.48

Note: CC = Conventional cigarettes; THS = Tobacco Heating System.

Note: Adjusted geometric least squares (LS) means and confidence intervals (CIs) from an ANCOVA model conducted on log-transformed values with log-transformed baseline value, study arm, sex and CC consumption reported at screening as fixed effect factors. Geometrical CV% of the ratio is estimated from the residual mean squares.

Baseline is defined as the last assessment prior to Day 1 product use

Appendix 15.4.4.59

Path: /cvn/projects/prj/development/000000106324/dev/tables/tl_anlhst.sas

Program Run: 27OCT14 cvn_ahedge Program Status: FINAL

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**Table 15.2.4.59 Analysis of HST Parameters (Averaged over all Cigarettes per Day) - FAS**

Variable (units)	Timepoint	Statistic	THS 2.2 (N=80)	CC (N=41)	THS : CC Ratio (%)
Average puff duration (s)	Day 1	n	55	27	
		Geometric LS Mean (CV%)	1.89	1.50	126.07 (19.04)
		95% CI	1.78, 1.99	1.38, 1.62	115.33, 137.81
	Day 4	n	55	27	
		Geometric LS Mean (CV%)	1.98	1.49	132.25 (20.38)
		95% CI	1.86, 2.09	1.37, 1.62	120.24, 145.46

Note: CC = Conventional cigarettes; THS = Tobacco Heating System.

Note: Adjusted geometric least squares (LS) means and confidence intervals (CIs) from an ANCOVA model conducted on log-transformed values with log-transformed baseline value, study arm, sex and CC consumption reported at screening as fixed effect factors. Geometrical CV% of the ratio is estimated from the residual mean squares.

Baseline is defined as the last assessment prior to Day 1 product use

Appendix 15.4.4.59

Path: /cvn/projects/prj/development/000000106324/dev/tables/tl_anlhst.sas

Program Run: 27OCT14 cvn_ahedge Program Status: FINAL

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**Table 15.2.4.59 Analysis of HST Parameters (Averaged over all Cigarettes per Day) - FAS**

Variable (units)	Timepoint	Statistic	THS 2.2 (N=80)	CC (N=41)	THS : CC Ratio (%)
Total puff duration (s)	Day 1	n	55	27	
		Geometric LS Mean (CV%)	31.55	23.19	136.09 (26.96)
		95% CI	29.32, 33.96	20.84, 25.79	119.89, 154.47
	Day 4	n	55	27	
		Geometric LS Mean (CV%)	32.21	23.45	137.33 (29.94)
		95% CI	29.70, 34.94	20.84, 26.39	119.35, 158.03

Note: CC = Conventional cigarettes; THS = Tobacco Heating System.

Note: Adjusted geometric least squares (LS) means and confidence intervals (CIs) from an ANCOVA model conducted on log-transformed values with log-transformed baseline value, study arm, sex and CC consumption reported at screening as fixed effect factors. Geometrical CV% of the ratio is estimated from the residual mean squares.

Baseline is defined as the last assessment prior to Day 1 product use

Appendix 15.4.4.59

Path: /cvn/projects/prj/development/000000106324/dev/tables/tl_anlhst.sas

Program Run: 27OCT14 cvn_ahedge Program Status: FINAL

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**Table 15.2.4.59 Analysis of HST Parameters (Averaged over all Cigarettes per Day) - FAS**

Variable (units)	Timepoint	Statistic	THS 2.2 (N=80)	CC (N=41)	THS : CC Ratio (%)
Average flow (mL/s)	Day 1	n	55	27	
		Geometric LS Mean (CV%)	25.48	31.22	81.64 (18.77)
		95% CI	24.20, 26.84	29.00, 33.61	74.78, 89.12
	Day 4	n	55	27	
		Geometric LS Mean (CV%)	25.98	32.49	79.95 (19.91)
		95% CI	24.59, 27.44	30.05, 35.13	72.86, 87.73

Note: CC = Conventional cigarettes; THS = Tobacco Heating System.

Note: Adjusted geometric least squares (LS) means and confidence intervals (CIs) from an ANCOVA model conducted on log-transformed values with log-transformed baseline value, study arm, sex and CC consumption reported at screening as fixed effect factors. Geometrical CV% of the ratio is estimated from the residual mean squares.

Baseline is defined as the last assessment prior to Day 1 product use

Appendix 15.4.4.59

Path: /cvn/projects/prj/development/000000106324/dev/tables/tl_anlhst.sas

Program Run: 27OCT14 cvn_ahedge Program Status: FINAL

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**Table 15.2.4.59 Analysis of HST Parameters (Averaged over all Cigarettes per Day) - FAS**

Variable (units)	Timepoint	Statistic	THS 2.2 (N=80)	CC (N=41)	THS : CC Ratio (%)
Average Peak flow (mL/s)	Day 1	n	55	27	
		Geometric LS Mean (CV%)	39.14	47.37	82.61 (18.51)
		95% CI	37.20, 41.18	44.02, 50.98	75.72, 90.12
	Day 4	n	55	27	
		Geometric LS Mean (CV%)	39.45	49.05	80.44 (19.73)
		95% CI	37.37, 41.65	45.35, 53.04	73.28, 88.29

Note: CC = Conventional cigarettes; THS = Tobacco Heating System.

Note: Adjusted geometric least squares (LS) means and confidence intervals (CIs) from an ANCOVA model conducted on log-transformed values with log-transformed baseline value, study arm, sex and CC consumption reported at screening as fixed effect factors. Geometrical CV% of the ratio is estimated from the residual mean squares.

Baseline is defined as the last assessment prior to Day 1 product use

Appendix 15.4.4.59

Path: /cvn/projects/prj/development/000000106324/dev/tables/tl_anlhst.sas

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**Table 15.2.4.59 Analysis of HST Parameters (Averaged over all Cigarettes per Day) - FAS**

Variable (units)	Timepoint	Statistic	THS 2.2 (N=80)	CC (N=41)	THS : CC Ratio (%)
Total inter puff interval (s)	Day 1	n	55	27	
		Geometric LS Mean (CV%)	169.83	230.55	73.66 (25.63)
		95% CI	158.28, 182.22	208.57, 254.84	65.32, 83.07
	Day 4	n	55	27	
		Geometric LS Mean (CV%)	159.35	225.22	70.75 (26.00)
		95% CI	148.37, 171.14	203.47, 249.30	62.63, 79.93

Note: CC = Conventional cigarettes; THS = Tobacco Heating System.

Note: Adjusted geometric least squares (LS) means and confidence intervals (CIs) from an ANCOVA model conducted on log-transformed values with log-transformed baseline value, study arm, sex and CC consumption reported at screening as fixed effect factors. Geometrical CV% of the ratio is estimated from the residual mean squares.

Baseline is defined as the last assessment prior to Day 1 product use

Appendix 15.4.4.59

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Program Run: 27OCT14 cvn_ahedge Program Status: FINAL

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**Table 15.2.4.59 Analysis of HST Parameters (Averaged over all Cigarettes per Day) - FAS**

Variable (units)	Timepoint	Statistic	THS 2.2 (N=80)	CC (N=41)	THS : CC Ratio (%)
Average inter puff interval (s)	Day 1	n	55	27	
		Geometric LS Mean (CV%)	10.26	15.03	68.24 (29.37)
		95% CI	9.46, 11.13	13.36, 16.91	59.20, 78.68
	Day 4	n	55	27	
		Geometric LS Mean (CV%)	9.85	14.52	67.83 (26.88)
		95% CI	9.14, 10.61	13.03, 16.17	59.51, 77.31

Note: CC = Conventional cigarettes; THS = Tobacco Heating System.

Note: Adjusted geometric least squares (LS) means and confidence intervals (CIs) from an ANCOVA model conducted on log-transformed values with log-transformed baseline value, study arm, sex and CC consumption reported at screening as fixed effect factors. Geometrical CV% of the ratio is estimated from the residual mean squares.

Baseline is defined as the last assessment prior to Day 1 product use

Appendix 15.4.4.59

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**Table 15.2.4.59 Analysis of HST Parameters (Averaged over all Cigarettes per Day) - FAS**

Variable (units)	Timepoint	Statistic	THS 2.2 (N=80)	CC (N=41)	THS : CC Ratio (%)
Total smoking duration (s)	Day 1	n	55	27	
		Geometric LS Mean (CV%)	205.27	255.35	80.39 (21.10)
		95% CI	193.66, 217.58	235.07, 277.39	72.80, 88.77
	Day 4	n	55	27	
		Geometric LS Mean (CV%)	194.84	250.94	77.64 (22.79)
		95% CI	182.99, 207.47	229.52, 274.37	69.77, 86.41

Note: CC = Conventional cigarettes; THS = Tobacco Heating System.

Note: Adjusted geometric least squares (LS) means and confidence intervals (CIs) from an ANCOVA model conducted on log-transformed values with log-transformed baseline value, study arm, sex and CC consumption reported at screening as fixed effect factors. Geometrical CV% of the ratio is estimated from the residual mean squares.

Baseline is defined as the last assessment prior to Day 1 product use

Appendix 15.4.4.59

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**Table 15.2.4.59 Analysis of HST Parameters (Averaged over all Cigarettes per Day) - FAS**

Variable (units)	Timepoint	Statistic	THS 2.2 (N=80)	CC (N=41)	THS : CC Ratio (%)
Total work (mJ)	Day 1	n	55	27	
		Geometric LS Mean (CV%)	1838.31	1694.92	108.46 (44.74)
		95% CI	1632.95, 2069.50	1431.28, 2007.12	88.69, 132.64
	Day 4	n	55	27	
		Geometric LS Mean (CV%)	1845.35	1850.25	99.74 (36.40)
		95% CI	1673.75, 2034.53	1609.06, 2127.59	84.47, 117.75

Note: CC = Conventional cigarettes; THS = Tobacco Heating System.

Note: Adjusted geometric least squares (LS) means and confidence intervals (CIs) from an ANCOVA model conducted on log-transformed values with log-transformed baseline value, study arm, sex and CC consumption reported at screening as fixed effect factors. Geometrical CV% of the ratio is estimated from the residual mean squares.

Baseline is defined as the last assessment prior to Day 1 product use

Appendix 15.4.4.59

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**Table 15.2.4.59 Analysis of HST Parameters (Averaged over all Cigarettes per Day) - FAS**

Variable (units)	Timepoint	Statistic	THS 2.2 (N=80)	CC (N=41)	THS : CC Ratio (%)
Average Work (mJ)	Day 1	n	55	27	
		Geometric LS Mean (CV%)	109.38	113.66	96.23 (40.89)
		95% CI	98.12, 121.93	97.09, 133.05	79.86, 115.95
	Day 4	n	55	27	
		Geometric LS Mean (CV%)	115.07	118.93	96.76 (35.16)
		95% CI	104.72, 126.45	103.66, 136.45	82.22, 113.86

Note: CC = Conventional cigarettes; THS = Tobacco Heating System.

Note: Adjusted geometric least squares (LS) means and confidence intervals (CIs) from an ANCOVA model conducted on log-transformed values with log-transformed baseline value, study arm, sex and CC consumption reported at screening as fixed effect factors. Geometrical CV% of the ratio is estimated from the residual mean squares.

Baseline is defined as the last assessment prior to Day 1 product use

Appendix 15.4.4.59

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**Table 15.2.4.59 Analysis of HST Parameters (Averaged over all Cigarettes per Day) - FAS**

Variable (units)	Timepoint	Statistic	THS 2.2 (N=80)	CC (N=41)	THS : CC Ratio (%)
Average pressure drop (mmWg)	Day 1	n	55	27	
		Geometric LS Mean (CV%)	205.60	208.63	98.55 (24.33)
		95% CI	192.41, 219.70	189.63, 229.52	88.00, 110.36
	Day 4	n	55	27	
		Geometric LS Mean (CV%)	202.90	216.45	93.74 (24.24)
		95% CI	189.93, 216.74	196.78, 238.09	83.72, 104.95

Note: CC = Conventional cigarettes; THS = Tobacco Heating System.

Note: Adjusted geometric least squares (LS) means and confidence intervals (CIs) from an ANCOVA model conducted on log-transformed values with log-transformed baseline value, study arm, sex and CC consumption reported at screening as fixed effect factors. Geometrical CV% of the ratio is estimated from the residual mean squares.

Baseline is defined as the last assessment prior to Day 1 product use

Appendix 15.4.4.59

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**Table 15.2.4.59 Analysis of HST Parameters (Averaged over all Cigarettes per Day) - FAS**

Variable (units)	Timepoint	Statistic	THS 2.2 (N=80)	CC (N=41)	THS : CC Ratio (%)
Average Peak pressure drop (mmWg)	Day 1	n	55	27	
		Geometric LS Mean (CV%)	341.98	339.74	100.66 (25.16)
		95% CI	319.36, 366.21	307.41, 375.46	89.40, 113.34
	Day 4	n	55	27	
		Geometric LS Mean (CV%)	333.20	353.72	94.20 (25.55)
		95% CI	310.81, 357.21	319.44, 391.67	83.45, 106.34

Note: CC = Conventional cigarettes; THS = Tobacco Heating System.

Note: Adjusted geometric least squares (LS) means and confidence intervals (CIs) from an ANCOVA model conducted on log-transformed values with log-transformed baseline value, study arm, sex and CC consumption reported at screening as fixed effect factors. Geometrical CV% of the ratio is estimated from the residual mean squares.

Baseline is defined as the last assessment prior to Day 1 product use

Appendix 15.4.4.59

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**Table 15.2.4.59 Analysis of HST Parameters (Averaged over all Cigarettes per Day) - FAS**

Variable (units)	Timepoint	Statistic	THS 2.2 (N=80)	CC (N=41)	THS : CC Ratio (%)
Smoking Intensity (mL/s)	Day 1	n	55	27	
		Geometric LS Mean (CV%)	3.96	2.87	138.25 (37.18)
		95% CI	3.58, 4.39	2.48, 3.32	116.16, 164.55
	Day 4	n	55	27	
		Geometric LS Mean (CV%)	4.36	3.12	139.82 (33.78)
		95% CI	3.97, 4.78	2.73, 3.56	119.31, 163.84

Note: CC = Conventional cigarettes; THS = Tobacco Heating System.

Note: Adjusted geometric least squares (LS) means and confidence intervals (CIs) from an ANCOVA model conducted on log-transformed values with log-transformed baseline value, study arm, sex and CC consumption reported at screening as fixed effect factors. Geometrical CV% of the ratio is estimated from the residual mean squares.

Baseline is defined as the last assessment prior to Day 1 product use

Appendix 15.4.4.59

Path: /cvn/projects/prj/development/000000106324/dev/tables/tl_anlhst.sas

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**Table 15.2.4.59 Analysis of HST Parameters (Averaged over all Cigarettes per Day) - FAS**

Variable (units)	Timepoint	Statistic	THS 2.2 (N=80)	CC (N=41)	THS : CC Ratio (%)
Puffing Time Index (%)	Day 1	n	55	27	
		Geometric LS Mean (CV%)	16.07	9.66	166.45 (32.05)
		95% CI	14.73, 17.54	8.52, 10.95	143.21, 193.45
	Day 4	n	55	27	
		Geometric LS Mean (CV%)	17.54	9.92	176.78 (32.18)
		95% CI	16.07, 19.15	8.75, 11.26	151.99, 205.62

Note: CC = Conventional cigarettes; THS = Tobacco Heating System.

Note: Adjusted geometric least squares (LS) means and confidence intervals (CIs) from an ANCOVA model conducted on log-transformed values with log-transformed baseline value, study arm, sex and CC consumption reported at screening as fixed effect factors. Geometrical CV% of the ratio is estimated from the residual mean squares.

Baseline is defined as the last assessment prior to Day 1 product use

Appendix 15.4.4.59

Path: /cvn/projects/prj/development/000000106324/dev/tables/tl_anlhst.sas

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**Table 15.2.4.59 Analysis of HST Parameters (Averaged over all Cigarettes per Day) - FAS**

Variable (units)	Timepoint	Statistic	THS 2.2 (N=80)	CC (N=41)	THS : CC Ratio (%)
Puff Frequency (puffs/min)	Day 1	n	55	27	
		Geometric LS Mean (CV%)	5.08	3.87	131.32 (23.76)
		95% CI	4.76, 5.44	3.51, 4.26	116.97, 147.43
	Day 4	n	55	27	
		Geometric LS Mean (CV%)	5.29	4.00	132.33 (23.04)
		95% CI	4.96, 5.64	3.64, 4.39	118.25, 148.09

Note: CC = Conventional cigarettes; THS = Tobacco Heating System.

Note: Adjusted geometric least squares (LS) means and confidence intervals (CIs) from an ANCOVA model conducted on log-transformed values with log-transformed baseline value, study arm, sex and CC consumption reported at screening as fixed effect factors. Geometrical CV% of the ratio is estimated from the residual mean squares.

Baseline is defined as the last assessment prior to Day 1 product use

Appendix 15.4.4.59

Path: /cvn/projects/prj/development/000000106324/dev/tables/tl_anlhst.sas

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**Table 15.2.4.60 Descriptive Statistics of HST Questionnaire Data - FAS**

Timepoint	Questionnaire items	THS 2.2 (N=80)		CC (N=41)	
		n	(%)	n	(%)
Day 0	The smoking of the CC/Products differs with the device				
	Strongly agree	16	(20.0%)	3	(7.3%)
	Agree	24	(30.0%)	11	(26.8%)
	Neither agree nor disagree	5	(6.3%)	5	(12.2%)
	Disagree	12	(15.0%)	8	(19.5%)
	Strongly disagree	1	(1.3%)	0	
	You enjoy smoking with the device as much as without it				
	Strongly agree	7	(8.8%)	3	(7.3%)
	Agree	14	(17.5%)	8	(19.5%)
	Neither agree nor disagree	8	(10.0%)	5	(12.2%)
	Disagree	18	(22.5%)	9	(22.0%)
	Strongly disagree	11	(13.8%)	2	(4.9%)

Note: CC = Conventional cigarettes; THS = Tobacco Heating System.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.7.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hstqu.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.60 Descriptive Statistics of HST Questionnaire Data - FAS**

Timepoint	Questionnaire items	THS 2.2 (N=80)		CC (N=41)	
		n	(%)	n	(%)
Day 0	The taste of the CC /products is different with the device				
	Strongly agree	4	(5.0%)	2	(4.9%)
	Agree	12	(15.0%)	4	(9.8%)
	Neither agree nor disagree	8	(10.0%)	4	(9.8%)
	Disagree	30	(37.5%)	16	(39.0%)
	Strongly disagree	4	(5.0%)	1	(2.4%)
	The device is easy to use				
	Strongly agree	25	(31.3%)	12	(29.3%)
	Agree	25	(31.3%)	12	(29.3%)
	Neither agree nor disagree	3	(3.8%)	2	(4.9%)
	Disagree	3	(3.8%)	1	(2.4%)
	Strongly disagree	2	(2.5%)	0	

Note: CC = Conventional cigarettes; THS = Tobacco Heating System.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.7.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hstqu.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.60 Descriptive Statistics of HST Questionnaire Data - FAS**

Timepoint	Questionnaire items	THS 2.2 (N=80)		CC (N=41)	
		n	(%)	n	(%)
Day 0	Your smoking is disturbed by the device				
	Strongly agree	8	(10.0%)	2	(4.9%)
	Agree	11	(13.8%)	3	(7.3%)
	Neither agree nor disagree	12	(15.0%)	6	(14.6%)
	Disagree	21	(26.3%)	14	(34.1%)
	Strongly disagree	6	(7.5%)	2	(4.9%)
Day 4	The smoking of the CC/Products differs with the device				
	Strongly agree	30	(37.5%)	3	(7.3%)
	Agree	21	(26.3%)	8	(19.5%)
	Neither agree nor disagree	13	(16.3%)	6	(14.6%)
	Disagree	15	(18.8%)	9	(22.0%)
	Strongly disagree	0		1	(2.4%)

Note: CC = Conventional cigarettes; THS = Tobacco Heating System.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.7.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hstqu.sas

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**Table 15.2.4.60 Descriptive Statistics of HST Questionnaire Data - FAS**

Timepoint	Questionnaire items	THS 2.2 (N=80)		CC (N=41)	
		n	(%)	n	(%)
Day 4	You enjoy smoking with the device as much as without it				
	Strongly agree	1	(1.3%)	3	(7.3%)
	Agree	22	(27.5%)	11	(26.8%)
	Neither agree nor disagree	16	(20.0%)	7	(17.1%)
	Disagree	22	(27.5%)	4	(9.8%)
	Strongly disagree	18	(22.5%)	2	(4.9%)
	The taste of the CC /products is different with the device				
	Strongly agree	7	(8.8%)	0	
	Agree	10	(12.5%)	4	(9.8%)
	Neither agree nor disagree	20	(25.0%)	3	(7.3%)
	Disagree	36	(45.0%)	15	(36.6%)
	Strongly disagree	6	(7.5%)	5	(12.2%)

Note: CC = Conventional cigarettes; THS = Tobacco Heating System.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.7.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_hstqu.sas

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**Table 15.2.4.60 Descriptive Statistics of HST Questionnaire Data - FAS**

Timepoint	Questionnaire items	THS 2.2 (N=80)		CC (N=41)	
		n	(%)	n	(%)
Day 4	The device is easy to use				
	Strongly agree	28	(35.0%)	11	(26.8%)
	Agree	37	(46.3%)	13	(31.7%)
	Neither agree nor disagree	9	(11.3%)	2	(4.9%)
	Disagree	3	(3.8%)	1	(2.4%)
	Strongly disagree	2	(2.5%)	0	
	Your smoking is disturbed by the device				
	Strongly agree	16	(20.0%)	2	(4.9%)
	Agree	23	(28.8%)	6	(14.6%)
	Neither agree nor disagree	17	(21.3%)	2	(4.9%)
	Disagree	19	(23.8%)	14	(34.1%)
	Strongly disagree	4	(5.0%)	3	(7.3%)

Note: CC = Conventional cigarettes; THS = Tobacco Heating System.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.7.2

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.61 Descriptive Statistics of Visual Inspection of the THS 2.2 Tobacco Plugs Data - FAS**

Timepoint	Evaluation	THS 2.2 (N=80) [n of THS 2.2 products used = 1191]	
		n	(%)
Day 1	0	1142	(98.9%)
	1	13	(1.1%)
	2	0	
	Missing	36	

Note: THS = Tobacco Heating System.

Note: Percentages are based on the number of non-missing inspections.

Note: 0 = No overheating, 1 = White spot(s) inside the tobacco plug, 2 = Ashes inside the tobacco plug and burnt paper.

Note: Missing values assessed as the difference between the number of sticks used and the number of sticks for which visual inspection of the plug has been performed.

Appendix 15.3.6.16

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_plug.sas

Program Run: 03NOV14 cvn_jhenshaw Program Status: FINAL

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**Table 15.2.4.61 Descriptive Statistics of Visual Inspection of the THS 2.2 Tobacco Plugs Data - FAS**

Timepoint	Evaluation	THS 2.2 (N=80) [n of THS 2.2 products used = 1384]	
		n	(%)
Day 2	0	1343	(98.2%)
	1	24	(1.8%)
	2	0	
	Missing	17	

Note: THS = Tobacco Heating System.

Note: Percentages are based on the number of non-missing inspections.

Note: 0 = No overheating, 1 = White spot(s) inside the tobacco plug, 2 = Ashes inside the tobacco plug and burnt paper.

Note: Missing values assessed as the difference between the number of sticks used and the number of sticks for which visual inspection of the plug has been performed.

Appendix 15.3.6.16

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_plug.sas

Program Run: 03NOV14 cvn_jhenshaw Program Status: FINAL

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**Table 15.2.4.61 Descriptive Statistics of Visual Inspection of the THS 2.2 Tobacco Plugs Data - FAS**

Timepoint	Evaluation	THS 2.2 (N=80) [n of THS 2.2 products used = 1457]	
		n	(%)
Day 3	0	1408	(98.3%)
	1	25	(1.7%)
	2	0	
	Missing	24	

Note: THS = Tobacco Heating System.

Note: Percentages are based on the number of non-missing inspections.

Note: 0 = No overheating, 1 = White spot(s) inside the tobacco plug, 2 = Ashes inside the tobacco plug and burnt paper.

Note: Missing values assessed as the difference between the number of sticks used and the number of sticks for which visual inspection of the plug has been performed.

Appendix 15.3.6.16

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_plug.sas

Program Run: 03NOV14 cvn_jhenshaw Program Status: FINAL

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**Table 15.2.4.61 Descriptive Statistics of Visual Inspection of the THS 2.2 Tobacco Plugs Data - FAS**

Timepoint	Evaluation	THS 2.2 (N=80) [n of THS 2.2 products used = 1459]	
		n	(%)
Day 4	0	1375	(97.9%)
	1	29	(2.1%)
	2	0	
	Missing	55	

Note: THS = Tobacco Heating System.

Note: Percentages are based on the number of non-missing inspections.

Note: 0 = No overheating, 1 = White spot(s) inside the tobacco plug, 2 = Ashes inside the tobacco plug and burnt paper.

Note: Missing values assessed as the difference between the number of sticks used and the number of sticks for which visual inspection of the plug has been performed.

Appendix 15.3.6.16

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_plug.sas

Program Run: 03NOV14 cvn_jhenshaw Program Status: FINAL

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**Table 15.2.4.61 Descriptive Statistics of Visual Inspection of the THS 2.2 Tobacco Plugs Data - FAS**

Timepoint	Evaluation	THS 2.2 (N=80) [n of THS 2.2 products used = 1639]	
		n	(%)
Day 5	0	1569	(98.4%)
	1	25	(1.6%)
	2	0	
	Missing	45	

Note: THS = Tobacco Heating System.

Note: Percentages are based on the number of non-missing inspections.

Note: 0 = No overheating, 1 = White spot(s) inside the tobacco plug, 2 = Ashes inside the tobacco plug and burnt paper.

Note: Missing values assessed as the difference between the number of sticks used and the number of sticks for which visual inspection of the plug has been performed.

Appendix 15.3.6.16

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_plug.sas

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Program Run: 03NOV14 cvn_jhenshaw Program Status: FINAL

**Table 15.2.4.62 Descriptive Statistics of Filter Analysis from the THS 2.2 Products - FAS**

Variable (units)	Timepoint	Statistic	THS 2.2 (N=80)
Analysis Full Filter Nicotine Amount per filter (mg/filter) (total over visit)	Day 1	n (THS 2.2 product smoked)	1191
		n (filters analysed)	1180
		n (subjects)	80
		Geometric Mean (CV%)	19.31262 (35.41%)
		95% CI	17.89056, 20.84772
		Median	19.50620
		Q25, Q75	15.96934, 25.24561
		Min, Max	6.6159, 50.9089
	Day 2	n (THS 2.2 product smoked)	1384
		n (filters analysed)	1376
		n (subjects)	80
		Geometric Mean (CV%)	23.63034 (37.22%)
		95% CI	21.81012, 25.60246
		Median	23.80405
		Q25, Q75	19.75705, 29.40395
		Min, Max	9.8982, 57.5031

Note: THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: Cohorts 1 (subjects 1 - 64) and 2 (subjects 66 - 123) had the full filter analyzed, the remaining subjects had the mouthpiece analyzed.

Appendix 15.3.6.17

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_filter.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.62 Descriptive Statistics of Filter Analysis from the THS 2.2 Products - FAS**

Variable (units)	Timepoint	Statistic	THS 2.2 (N=80)
Analysis Full Filter Nicotine Amount per filter (mg/filter) (total over visit)	Day 3	n (THS 2.2 product smoked)	1457
		n (filters analysed)	1453
		n (subjects)	80
		Geometric Mean (CV%)	25.72810 (37.45%)
		95% CI	23.73548, 27.88800
		Median	26.10085
		Q25, Q75	20.97110, 32.31660
		Min, Max	7.0956, 61.9997
	Day 4	n (THS 2.2 product smoked)	1459
		n (filters analysed)	1452
		n (subjects)	79
		Geometric Mean (CV%)	23.92398 (38.13%)
		95% CI	22.02924, 25.98170
		Median	23.53760
		Q25, Q75	18.42160, 30.52830
		Min, Max	7.8204, 61.9359

Note: THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: Cohorts 1 (subjects 1 - 64) and 2 (subjects 66 - 123) had the full filter analyzed, the remaining subjects had the mouthpiece analyzed.

Appendix 15.3.6.17

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_filter.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.62 Descriptive Statistics of Filter Analysis from the THS 2.2 Products - FAS**

Variable (units)	Timepoint	Statistic	THS 2.2 (N=80)
Analysis Full Filter Nicotine Amount per filter (mg/filter) Day 5 (total over visit)		n (THS 2.2 product smoked)	1639
		n (filters analysed)	1638
		n (subjects)	79
		Geometric Mean (CV%)	30.08588 (36.44%)
		95% CI	27.79825, 32.56177
		Median	30.55960
		Q25, Q75	24.38550, 37.09030
		Min, Max	11.4820, 71.7236
Analysis Full Filter Nicotine Amount per mL (mg/mL) Day 1 (total over visit)		n (THS 2.2 product smoked)	1191
		n (filters analysed)	1180
		n (subjects)	80
		Geometric Mean (CV%)	1.93127 (35.42%)
		95% CI	1.78905, 2.08479
		Median	1.95060
		Q25, Q75	1.59690, 2.52480
		Min, Max	0.6615, 5.0905

Note: THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: Cohorts 1 (subjects 1 - 64) and 2 (subjects 66 - 123) had the full filter analyzed, the remaining subjects had the mouthpiece analyzed.

Appendix 15.3.6.17

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_filter.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.62 Descriptive Statistics of Filter Analysis from the THS 2.2 Products - FAS**

Variable (units)	Timepoint	Statistic	THS 2.2 (N=80)
Analysis Full Filter Nicotine Amount per mL (mg/mL) (total over visit)	Day 2	n (THS 2.2 product smoked)	1384
		n (filters analysed)	1376
		n (subjects)	80
		Geometric Mean (CV%)	2.36301 (37.22%)
		95% CI	2.18099, 2.56022
		Median	2.38035
		Q25, Q75	1.97580, 2.94015
		Min, Max	0.9900, 5.7500
	Day 3	n (THS 2.2 product smoked)	1457
		n (filters analysed)	1453
		n (subjects)	80
		Geometric Mean (CV%)	2.57281 (37.45%)
		95% CI	2.37354, 2.78881
		Median	2.61010
		Q25, Q75	2.09724, 3.23180
		Min, Max	0.7098, 6.2003

Note: THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: Cohorts 1 (subjects 1 - 64) and 2 (subjects 66 - 123) had the full filter analyzed, the remaining subjects had the mouthpiece analyzed.

Appendix 15.3.6.17

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_filter.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.62 Descriptive Statistics of Filter Analysis from the THS 2.2 Products - FAS**

Variable (units)	Timepoint	Statistic	THS 2.2 (N=80)
Analysis Full Filter Nicotine Amount per mL (mg/mL) (total over visit)	Day 4	n (THS 2.2 product smoked)	1459
		n (filters analysed)	1452
		n (subjects)	79
		Geometric Mean (CV%)	2.39243 (38.13%)
		95% CI	2.20294, 2.59821
		Median	2.35300
		Q25, Q75	1.84200, 3.05230
		Min, Max	0.7821, 6.1942
	Day 5	n (THS 2.2 product smoked)	1639
		n (filters analysed)	1638
		n (subjects)	79
		Geometric Mean (CV%)	3.00859 (36.44%)
		95% CI	2.77983, 3.25618
		Median	3.05630
		Q25, Q75	2.43870, 3.70950
		Min, Max	1.1480, 7.1721

Note: THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: Cohorts 1 (subjects 1 - 64) and 2 (subjects 66 - 123) had the full filter analyzed, the remaining subjects had the mouthpiece analyzed.

Appendix 15.3.6.17

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_filter.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.62 Descriptive Statistics of Filter Analysis from the THS 2.2 Products - FAS**

Variable (units)	Timepoint	Statistic	THS 2.2 (N=80)
Analysis Full Filter Absolute UV TAR Absorbance (total over visit)	Day 1	n (THS 2.2 product smoked)	1191
		n (filters analysed)	1180
		n (subjects)	80
		Geometric Mean (CV%)	5.67327 (37.66%)
		95% CI	5.23167, 6.15215
		Median	5.95475
		Q25, Q75	4.50135, 7.49230
		Min, Max	1.6560, 14.2999
	Day 2	n (THS 2.2 product smoked)	1384
		n (filters analysed)	1376
		n (subjects)	80
		Geometric Mean (CV%)	7.01623 (38.26%)
		95% CI	6.46233, 7.61761
		Median	7.34015
		Q25, Q75	5.85070, 9.03495
		Min, Max	2.4930, 15.1430

Note: THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: Cohorts 1 (subjects 1 - 64) and 2 (subjects 66 - 123) had the full filter analyzed, the remaining subjects had the mouthpiece analyzed.

Appendix 15.3.6.17

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_filter.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.62 Descriptive Statistics of Filter Analysis from the THS 2.2 Products - FAS**

Variable (units)	Timepoint	Statistic	THS 2.2 (N=80)
Analysis Full Filter Absolute UV TAR Absorbance (total over visit)	Day 3	n (THS 2.2 product smoked)	1457
		n (filters analysed)	1453
		n (subjects)	80
		Geometric Mean (CV%)	7.37720 (37.26%)
		95% CI	6.80843, 7.99349
		Median	8.09905
		Q25, Q75	6.23044, 9.05601
		Min, Max	1.7697, 14.7490
	Day 4	n (THS 2.2 product smoked)	1459
		n (filters analysed)	1452
		n (subjects)	79
		Geometric Mean (CV%)	6.95239 (39.54%)
		95% CI	6.38360, 7.57187
		Median	6.89780
		Q25, Q75	5.64680, 9.12960
		Min, Max	1.4922, 19.2492

Note: THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: Cohorts 1 (subjects 1 - 64) and 2 (subjects 66 - 123) had the full filter analyzed, the remaining subjects had the mouthpiece analyzed.

Appendix 15.3.6.17

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_filter.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.62 Descriptive Statistics of Filter Analysis from the THS 2.2 Products - FAS**

Variable (units)	Timepoint	Statistic	THS 2.2 (N=80)
Analysis Full Filter Absolute UV TAR Absorbance (total over visit)	Day 5	n (THS 2.2 product smoked)	1639
		n (filters analysed)	1638
		n (subjects)	79
		Geometric Mean (CV%)	8.36063 (38.34%)
		95% CI	7.69510, 9.08371
		Median	8.62540
		Q25, Q75	6.77580, 10.70340
		Min, Max	3.2740, 19.6837
Analysis Full Filter Normalized UV Absorbance (per filter) (total over visit)	Day 1	n (THS 2.2 product smoked)	1191
		n (filters analysed)	1180
		n (subjects)	80
		Geometric Mean (CV%)	34.00694 (37.72%)
		95% CI	31.35587, 36.88216
		Median	35.72850
		Q25, Q75	26.88630, 44.95380
		Min, Max	9.9360, 85.7994

Note: THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: Cohorts 1 (subjects 1 - 64) and 2 (subjects 66 - 123) had the full filter analyzed, the remaining subjects had the mouthpiece analyzed.

Appendix 15.3.6.17

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_filter.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.62 Descriptive Statistics of Filter Analysis from the THS 2.2 Products - FAS**

Variable (units)	Timepoint	Statistic	THS 2.2 (N=80)
Analysis Full Filter Normalized UV Absorbance (per filter) (total over visit)	Day 2	n (THS 2.2 product smoked)	1384
		n (filters analysed)	1376
		n (subjects)	80
		Geometric Mean (CV%)	41.94787 (39.22%)
		95% CI	38.56199, 45.63104
		Median	44.04090
		Q25, Q75	35.10420, 54.20970
		Min, Max	14.8572, 90.8580
	Day 3	n (THS 2.2 product smoked)	1457
		n (filters analysed)	1453
		n (subjects)	80
		Geometric Mean (CV%)	44.03079 (38.97%)
		95% CI	40.49673, 47.87326
		Median	48.50510
		Q25, Q75	37.38270, 54.33600
		Min, Max	8.2674, 88.4940

Note: THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: Cohorts 1 (subjects 1 - 64) and 2 (subjects 66 - 123) had the full filter analyzed, the remaining subjects had the mouthpiece analyzed.

Appendix 15.3.6.17

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_filter.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.62 Descriptive Statistics of Filter Analysis from the THS 2.2 Products - FAS**

Variable (units)	Timepoint	Statistic	THS 2.2 (N=80)
Analysis Full Filter Normalized UV Absorbance (per filter) (total over visit)	Day 4	n (THS 2.2 product smoked)	1459
		n (filters analysed)	1452
		n (subjects)	79
		Geometric Mean (CV%)	41.50950 (39.97%)
		95% CI	38.08035, 45.24745
		Median	41.38680
		Q25, Q75	33.88080, 54.77760
		Min, Max	8.9532, 115.4952
	Day 5	n (THS 2.2 product smoked)	1639
		n (filters analysed)	1638
		n (subjects)	79
		Geometric Mean (CV%)	49.96771 (38.52%)
		95% CI	45.97388, 54.30850
		Median	51.75240
		Q25, Q75	39.09179, 64.22040
		Min, Max	19.6440, 117.8826

Note: THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: Cohorts 1 (subjects 1 - 64) and 2 (subjects 66 - 123) had the full filter analyzed, the remaining subjects had the mouthpiece analyzed.

Appendix 15.3.6.17

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_filter.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.62 Descriptive Statistics of Filter Analysis from the THS 2.2 Products - FAS**

Variable (units)	Timepoint	Statistic	THS 2.2 (N=80)
Analysis Mouthpiece Nicotine Amount per filter (mg/filter) (total over visit)	Day 1	n (THS 2.2 product smoked)	1191
		n (filters analysed)	731
		n (subjects)	49
		Geometric Mean (CV%)	3.76305 (46.42%)
		95% CI	3.31470, 4.27204
		Median	3.92930
		Q25, Q75	3.09900, 4.83370
		Min, Max	0.7345, 13.2495
	Day 2	n (THS 2.2 product smoked)	1384
		n (filters analysed)	861
		n (subjects)	49
		Geometric Mean (CV%)	4.66075 (43.07%)
		95% CI	4.14007, 5.24692
		Median	5.09230
		Q25, Q75	3.89800, 5.95360
		Min, Max	1.1219, 8.4520

Note: THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: Cohorts 1 (subjects 1 - 64) and 2 (subjects 66 - 123) had the full filter analyzed, the remaining subjects had the mouthpiece analyzed.

Appendix 15.3.6.17

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_filter.sas

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**Table 15.2.4.62 Descriptive Statistics of Filter Analysis from the THS 2.2 Products - FAS**

Variable (units)	Timepoint	Statistic	THS 2.2 (N=80)
Analysis Mouthpiece Nicotine Amount per filter (mg/filter) (total over visit)	Day 3	n (THS 2.2 product smoked)	1457
		n (filters analysed)	873
		n (subjects)	49
		Geometric Mean (CV%)	4.75788 (46.67%)
		95% CI	4.18832, 5.40488
		Median	5.19290
		Q25, Q75	3.96820, 6.04360
		Min, Max	0.6459, 9.0110
	Day 4	n (THS 2.2 product smoked)	1459
		n (filters analysed)	860
		n (subjects)	49
		Geometric Mean (CV%)	4.64127 (46.15%)
		95% CI	4.09103, 5.26551
		Median	4.82470
		Q25, Q75	3.87539, 6.34330
		Min, Max	0.7416, 10.1742

Note: THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: Cohorts 1 (subjects 1 - 64) and 2 (subjects 66 - 123) had the full filter analyzed, the remaining subjects had the mouthpiece analyzed.

Appendix 15.3.6.17

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_filter.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.62 Descriptive Statistics of Filter Analysis from the THS 2.2 Products - FAS**

Variable (units)	Timepoint	Statistic	THS 2.2 (N=80)
Analysis Mouthpiece Nicotine Amount per filter (mg/filter) (total over visit)	Day 5	n (THS 2.2 product smoked)	1639
		n (filters analysed)	1026
		n (subjects)	49
		Geometric Mean (CV%)	5.32515 (49.55%)
		95% CI	4.65464, 6.09226
		Median	5.62240
		Q25, Q75	4.23880, 7.24960
		Min, Max	1.3204, 11.7739
Analysis Mouthpiece Nicotine Amount per mL (mg/mL) Day 1 (total over visit)		n (THS 2.2 product smoked)	1191
		n (filters analysed)	731
		n (subjects)	49
		Geometric Mean (CV%)	0.37628 (46.41%)
		95% CI	0.33145, 0.42718
		Median	0.39290
		Q25, Q75	0.30990, 0.48320
		Min, Max	0.0735, 1.3246

Note: THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: Cohorts 1 (subjects 1 - 64) and 2 (subjects 66 - 123) had the full filter analyzed, the remaining subjects had the mouthpiece analyzed.

Appendix 15.3.6.17

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_filter.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.62 Descriptive Statistics of Filter Analysis from the THS 2.2 Products - FAS**

Variable (units)	Timepoint	Statistic	THS 2.2 (N=80)
Analysis Mouthpiece Nicotine Amount per mL (mg/mL) Day 2 (total over visit)		n (THS 2.2 product smoked)	1384
		n (filters analysed)	861
		n (subjects)	49
		Geometric Mean (CV%)	0.46602 (43.06%)
		95% CI	0.41395, 0.52463
		Median	0.50880
		Q25, Q75	0.38930, 0.59480
		Min, Max	0.1122, 0.8451
	Day 3	n (THS 2.2 product smoked)	1457
		n (filters analysed)	873
		n (subjects)	49
		Geometric Mean (CV%)	0.47582 (46.65%)
		95% CI	0.41888, 0.54051
		Median	0.51900
		Q25, Q75	0.39670, 0.60420
		Min, Max	0.0648, 0.9006

Note: THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: Cohorts 1 (subjects 1 - 64) and 2 (subjects 66 - 123) had the full filter analyzed, the remaining subjects had the mouthpiece analyzed.

Appendix 15.3.6.17

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_filter.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.62 Descriptive Statistics of Filter Analysis from the THS 2.2 Products - FAS**

Variable (units)	Timepoint	Statistic	THS 2.2 (N=80)
Analysis Mouthpiece Nicotine Amount per mL (mg/mL) Day 4 (total over visit)		n (THS 2.2 product smoked)	1459
		n (filters analysed)	860
		n (subjects)	49
		Geometric Mean (CV%)	0.46414 (46.16%)
		95% CI	0.40910, 0.52659
		Median	0.48250
		Q25, Q75	0.38770, 0.63430
		Min, Max	0.0741, 1.0174
	Day 5	n (THS 2.2 product smoked)	1639
		n (filters analysed)	1026
		n (subjects)	49
		Geometric Mean (CV%)	0.53254 (49.54%)
		95% CI	0.46549, 0.60925
		Median	0.56220
		Q25, Q75	0.42380, 0.72520
		Min, Max	0.1320, 1.1772

Note: THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: Cohorts 1 (subjects 1 - 64) and 2 (subjects 66 - 123) had the full filter analyzed, the remaining subjects had the mouthpiece analyzed.

Appendix 15.3.6.17

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_filter.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.62 Descriptive Statistics of Filter Analysis from the THS 2.2 Products - FAS**

Variable (units)	Timepoint	Statistic	THS 2.2 (N=80)
Analysis Mouthpiece Absolute UV TAR Absorbance (total over visit)	Day 1	n (THS 2.2 product smoked)	1191
		n (filters analysed)	731
		n (subjects)	49
		Geometric Mean (CV%)	1.41723 (47.88%)
		95% CI	1.24385, 1.61476
		Median	1.44430
		Q25, Q75	1.18400, 1.87190
		Min, Max	0.3480, 4.2836
	Day 2	n (THS 2.2 product smoked)	1384
		n (filters analysed)	861
		n (subjects)	49
		Geometric Mean (CV%)	1.78348 (38.74%)
		95% CI	1.60186, 1.98570
		Median	1.94370
		Q25, Q75	1.43580, 2.34400
		Min, Max	0.4211, 3.3012

Note: THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: Cohorts 1 (subjects 1 - 64) and 2 (subjects 66 - 123) had the full filter analyzed, the remaining subjects had the mouthpiece analyzed.

Appendix 15.3.6.17

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_filter.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.62 Descriptive Statistics of Filter Analysis from the THS 2.2 Products - FAS**

Variable (units)	Timepoint	Statistic	THS 2.2 (N=80)
Analysis Mouthpiece Absolute UV TAR Absorbance (total over visit)	Day 3	n (THS 2.2 product smoked)	1457
		n (filters analysed)	873
		n (subjects)	49
		Geometric Mean (CV%)	1.75426 (35.43%)
		95% CI	1.58930, 1.93634
		Median	1.86000
		Q25, Q75	1.40990, 2.28090
		Min, Max	0.5877, 3.1993
	Day 4	n (THS 2.2 product smoked)	1459
		n (filters analysed)	860
		n (subjects)	49
		Geometric Mean (CV%)	1.67115 (47.68%)
		95% CI	1.46744, 1.90315
		Median	1.64810
		Q25, Q75	1.35430, 2.23120
		Min, Max	0.2292, 3.5030

Note: THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: Cohorts 1 (subjects 1 - 64) and 2 (subjects 66 - 123) had the full filter analyzed, the remaining subjects had the mouthpiece analyzed.

Appendix 15.3.6.17

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_filter.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.62 Descriptive Statistics of Filter Analysis from the THS 2.2 Products - FAS**

Variable (units)	Timepoint	Statistic	THS 2.2 (N=80)
Analysis Mouthpiece Absolute UV TAR Absorbance (total over visit)	Day 5	n (THS 2.2 product smoked)	1639
		n (filters analysed)	1026
		n (subjects)	49
		Geometric Mean (CV%)	1.88061 (53.09%)
		95% CI	1.62983, 2.16998
		Median	2.16380
		Q25, Q75	1.47840, 2.64730
		Min, Max	0.4207, 4.6923
Analysis Mouthpiece Normalized UV Absorbance (per filter) (total over visit)	Day 1	n (THS 2.2 product smoked)	1191
		n (filters analysed)	731
		n (subjects)	49
		Geometric Mean (CV%)	8.43864 (48.71%)
		95% CI	7.39134, 9.63432
		Median	8.66580
		Q25, Q75	7.10400, 11.23140
		Min, Max	2.0880, 25.7016

Note: THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: Cohorts 1 (subjects 1 - 64) and 2 (subjects 66 - 123) had the full filter analyzed, the remaining subjects had the mouthpiece analyzed.

Appendix 15.3.6.17

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_filter.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.62 Descriptive Statistics of Filter Analysis from the THS 2.2 Products - FAS**

Variable (units)	Timepoint	Statistic	THS 2.2 (N=80)
Analysis Mouthpiece Normalized UV Absorbance (per filter) (total over visit)	Day 2	n (THS 2.2 product smoked)	1384
		n (filters analysed)	861
		n (subjects)	49
		Geometric Mean (CV%)	10.46366 (44.64%)
		95% CI	9.25783, 11.82654
		Median	11.66220
		Q25, Q75	8.61480, 14.06400
		Min, Max	2.4462, 19.8072
	Day 3	n (THS 2.2 product smoked)	1457
		n (filters analysed)	873
		n (subjects)	49
		Geometric Mean (CV%)	10.10987 (49.10%)
		95% CI	8.84669, 11.55342
		Median	11.16000
		Q25, Q75	8.45940, 13.46700
		Min, Max	1.1754, 19.1958

Note: THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: Cohorts 1 (subjects 1 - 64) and 2 (subjects 66 - 123) had the full filter analyzed, the remaining subjects had the mouthpiece analyzed.

Appendix 15.3.6.17

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_filter.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.62 Descriptive Statistics of Filter Analysis from the THS 2.2 Products - FAS**

Variable (units)	Timepoint	Statistic	THS 2.2 (N=80)
Analysis Mouthpiece Normalized UV Absorbance (per filter) (total over visit)	Day 4	n (THS 2.2 product smoked)	1459
		n (filters analysed)	860
		n (subjects)	49
		Geometric Mean (CV%)	9.66207 (51.59%)
		95% CI	8.40373, 11.10884
		Median	9.88860
		Q25, Q75	7.78680, 13.38720
		Min, Max	1.3752, 21.0180
	Day 5	n (THS 2.2 product smoked)	1639
		n (filters analysed)	1026
		n (subjects)	49
		Geometric Mean (CV%)	11.00479 (53.38%)
		95% CI	9.53078, 12.70677
		Median	12.48360
		Q25, Q75	8.53200, 15.57420
		Min, Max	2.5242, 26.0466

Note: THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: Cohorts 1 (subjects 1 - 64) and 2 (subjects 66 - 123) had the full filter analyzed, the remaining subjects had the mouthpiece analyzed.

Appendix 15.3.6.17

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_filter.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.62 Descriptive Statistics of Filter Analysis from the THS 2.2 Products - FAS**

Variable (units)	Timepoint	Statistic	THS 2.2 (N=80)
Analysis PLA + Hat Nicotine Amount per filter (mg/filter) (total over visit)	Day 1	n (THS 2.2 product smoked)	1191
		n (filters analysed)	731
		n (subjects)	49
		Geometric Mean (CV%)	15.70244 (32.03%)
		95% CI	14.35447, 17.17700
		Median	15.67950
		Q25, Q75	12.78740, 19.93800
		Min, Max	6.4395, 37.6594
	Day 2	n (THS 2.2 product smoked)	1384
		n (filters analysed)	861
		n (subjects)	49
		Geometric Mean (CV%)	19.53099 (32.86%)
		95% CI	17.81505, 21.41221
		Median	19.08640
		Q25, Q75	15.70980, 24.14260
		Min, Max	9.1427, 40.4537

Note: THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: Cohorts 1 (subjects 1 - 64) and 2 (subjects 66 - 123) had the full filter analyzed, the remaining subjects had the mouthpiece analyzed.

Appendix 15.3.6.17

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_filter.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.62 Descriptive Statistics of Filter Analysis from the THS 2.2 Products - FAS**

Variable (units)	Timepoint	Statistic	THS 2.2 (N=80)
Analysis PLA + Hat Nicotine Amount per filter (mg/filter) (total over visit)	Day 3	n (THS 2.2 product smoked)	1457
		n (filters analysed)	873
		n (subjects)	49
		Geometric Mean (CV%)	21.17921 (32.28%)
		95% CI	19.34806, 23.18367
		Median	20.72850
		Q25, Q75	18.53820, 26.67240
		Min, Max	6.4497, 44.4685
	Day 4	n (THS 2.2 product smoked)	1459
		n (filters analysed)	860
		n (subjects)	49
		Geometric Mean (CV%)	19.20392 (34.73%)
		95% CI	17.43014, 21.15821
		Median	19.23490
		Q25, Q75	16.00460, 23.67790
		Min, Max	7.0788, 40.3703

Note: THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: Cohorts 1 (subjects 1 - 64) and 2 (subjects 66 - 123) had the full filter analyzed, the remaining subjects had the mouthpiece analyzed.

Appendix 15.3.6.17

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_filter.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.62 Descriptive Statistics of Filter Analysis from the THS 2.2 Products - FAS**

Variable (units)	Timepoint	Statistic	THS 2.2 (N=80)
Analysis PLA + Hat Nicotine Amount per filter (mg/filter) (total over visit)	Day 5	n (THS 2.2 product smoked)	1639
		n (filters analysed)	1026
		n (subjects)	49
		Geometric Mean (CV%)	26.13131 (36.96%)
		95% CI	23.57897, 28.95992
		Median	26.24570
		Q25, Q75	20.04880, 33.24861
		Min, Max	11.4755, 59.9497
Analysis PLA + Hat Nicotine Amount per mL (mg/mL) (total over visit)	Day 1	n (THS 2.2 product smoked)	1191
		n (filters analysed)	731
		n (subjects)	49
		Geometric Mean (CV%)	1.57026 (32.03%)
		95% CI	1.43544, 1.71773
		Median	1.56770
		Q25, Q75	1.27880, 1.99400
		Min, Max	0.6440, 3.7659

Note: THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: Cohorts 1 (subjects 1 - 64) and 2 (subjects 66 - 123) had the full filter analyzed, the remaining subjects had the mouthpiece analyzed.

Appendix 15.3.6.17

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_filter.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.62 Descriptive Statistics of Filter Analysis from the THS 2.2 Products - FAS**

Variable (units)	Timepoint	Statistic	THS 2.2 (N=80)
Analysis PLA + Hat Nicotine Amount per mL (mg/mL) (total over visit)	Day 2	n (THS 2.2 product smoked)	1384
		n (filters analysed)	861
		n (subjects)	49
		Geometric Mean (CV%)	1.95312 (32.86%)
		95% CI	1.78153, 2.14124
		Median	1.90880
		Q25, Q75	1.57080, 2.41450
		Min, Max	0.9144, 4.0453
	Day 3	n (THS 2.2 product smoked)	1457
		n (filters analysed)	873
		n (subjects)	49
		Geometric Mean (CV%)	2.11796 (32.28%)
		95% CI	1.93483, 2.31842
		Median	2.07310
		Q25, Q75	1.85360, 2.66760
		Min, Max	0.6450, 4.4467

Note: THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: Cohorts 1 (subjects 1 - 64) and 2 (subjects 66 - 123) had the full filter analyzed, the remaining subjects had the mouthpiece analyzed.

Appendix 15.3.6.17

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_filter.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.62 Descriptive Statistics of Filter Analysis from the THS 2.2 Products - FAS**

Variable (units)	Timepoint	Statistic	THS 2.2 (N=80)
Analysis PLA + Hat Nicotine Amount per mL (mg/mL) (total over visit)	Day 4	n (THS 2.2 product smoked)	1459
		n (filters analysed)	860
		n (subjects)	49
		Geometric Mean (CV%)	1.92039 (34.73%)
		95% CI	1.74301, 2.11582
		Median	1.92360
		Q25, Q75	1.60040, 2.36760
		Min, Max	0.7080, 4.0366
	Day 5	n (THS 2.2 product smoked)	1639
		n (filters analysed)	1026
		n (subjects)	49
		Geometric Mean (CV%)	2.61316 (36.96%)
		95% CI	2.35792, 2.89602
		Median	2.62450
		Q25, Q75	2.00490, 3.32480
		Min, Max	1.1475, 5.9949

Note: THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: Cohorts 1 (subjects 1 - 64) and 2 (subjects 66 - 123) had the full filter analyzed, the remaining subjects had the mouthpiece analyzed.

Appendix 15.3.6.17

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_filter.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.62 Descriptive Statistics of Filter Analysis from the THS 2.2 Products - FAS**

Variable (units)	Timepoint	Statistic	THS 2.2 (N=80)
Analysis PLA + Hat Absolute UV TAR Absorbance (total over visit)	Day 1	n (THS 2.2 product smoked)	1191
		n (filters analysed)	731
		n (subjects)	49
		Geometric Mean (CV%)	4.36105 (35.02%)
		95% CI	3.95517, 4.80858
		Median	4.41010
		Q25, Q75	3.63440, 5.45940
		Min, Max	1.3080, 10.0163
	Day 2	n (THS 2.2 product smoked)	1384
		n (filters analysed)	861
		n (subjects)	49
		Geometric Mean (CV%)	5.61103 (32.83%)
		95% CI	5.11841, 6.15105
		Median	5.53110
		Q25, Q75	5.05830, 6.67910
		Min, Max	2.0685, 9.5928

Note: THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: Cohorts 1 (subjects 1 - 64) and 2 (subjects 66 - 123) had the full filter analyzed, the remaining subjects had the mouthpiece analyzed.

Appendix 15.3.6.17

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_filter.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.62 Descriptive Statistics of Filter Analysis from the THS 2.2 Products - FAS**

Variable (units)	Timepoint	Statistic	THS 2.2 (N=80)
Analysis PLA + Hat Absolute UV TAR Absorbance (total over visit)	Day 3	n (THS 2.2 product smoked)	1457
		n (filters analysed)	873
		n (subjects)	49
		Geometric Mean (CV%)	5.68688 (34.73%)
		95% CI	5.16161, 6.26560
		Median	5.94080
		Q25, Q75	4.93969, 7.02610
		Min, Max	1.1820, 11.5323
	Day 4	n (THS 2.2 product smoked)	1459
		n (filters analysed)	860
		n (subjects)	49
		Geometric Mean (CV%)	5.22084 (37.68%)
		95% CI	4.70208, 5.79683
		Median	5.24590
		Q25, Q75	4.39490, 6.47690
		Min, Max	1.2630, 9.9903

Note: THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: Cohorts 1 (subjects 1 - 64) and 2 (subjects 66 - 123) had the full filter analyzed, the remaining subjects had the mouthpiece analyzed.

Appendix 15.3.6.17

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_filter.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.62 Descriptive Statistics of Filter Analysis from the THS 2.2 Products - FAS**

Variable (units)	Timepoint	Statistic	THS 2.2 (N=80)
Analysis PLA + Hat Absolute UV TAR Absorbance (total over visit)	Day 5	n (THS 2.2 product smoked)	1639
		n (filters analysed)	1026
		n (subjects)	49
		Geometric Mean (CV%)	6.74474 (39.93%)
		95% CI	6.03936, 7.53251
		Median	6.95170
		Q25, Q75	5.30940, 8.79420
		Min, Max	2.7153, 14.9914
Analysis PLA + Hat Normalized UV Absorbance (per filter) (total over visit)	Day 1	n (THS 2.2 product smoked)	1191
		n (filters analysed)	731
		n (subjects)	49
		Geometric Mean (CV%)	26.16630 (35.02%)
		95% CI	23.73106, 28.85143
		Median	26.46060
		Q25, Q75	21.80640, 32.75640
		Min, Max	7.8480, 60.0978

Note: THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: Cohorts 1 (subjects 1 - 64) and 2 (subjects 66 - 123) had the full filter analyzed, the remaining subjects had the mouthpiece analyzed.

Appendix 15.3.6.17

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_filter.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.62 Descriptive Statistics of Filter Analysis from the THS 2.2 Products - FAS**

Variable (units)	Timepoint	Statistic	THS 2.2 (N=80)
Analysis PLA + Hat Normalized UV Absorbance (per filter) (total over visit)	Day 2	n (THS 2.2 product smoked)	1384
		n (filters analysed)	861
		n (subjects)	49
		Geometric Mean (CV%)	33.66616 (32.83%)
		95% CI	30.71049, 36.90630
		Median	33.18660
		Q25, Q75	30.34980, 40.07460
		Min, Max	12.4110, 57.5568
	Day 3	n (THS 2.2 product smoked)	1457
		n (filters analysed)	873
		n (subjects)	49
		Geometric Mean (CV%)	34.12125 (34.73%)
		95% CI	30.96967, 37.59355
		Median	35.64480
		Q25, Q75	29.63820, 42.15660
		Min, Max	7.0920, 69.1938

Note: THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: Cohorts 1 (subjects 1 - 64) and 2 (subjects 66 - 123) had the full filter analyzed, the remaining subjects had the mouthpiece analyzed.

Appendix 15.3.6.17

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_filter.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.4.62 Descriptive Statistics of Filter Analysis from the THS 2.2 Products - FAS**

Variable (units)	Timepoint	Statistic	THS 2.2 (N=80)
Analysis PLA + Hat Normalized UV Absorbance (per filter) (total over visit)	Day 4	n (THS 2.2 product smoked)	1459
		n (filters analysed)	860
		n (subjects)	49
		Geometric Mean (CV%)	31.32505 (37.68%)
		95% CI	28.21251, 34.78098
		Median	31.47540
		Q25, Q75	26.36940, 38.86141
		Min, Max	7.5780, 59.9418
	Day 5	n (THS 2.2 product smoked)	1639
		n (filters analysed)	1026
		n (subjects)	49
		Geometric Mean (CV%)	40.46845 (39.93%)
		95% CI	36.23620, 45.19501
		Median	41.71020
		Q25, Q75	31.85640, 52.76520
		Min, Max	16.2918, 89.9484

Note: THS = Tobacco Heating System.

Note: Geometric: mean, CV% and 95% confidence interval (CI) are reported.

Note: Cohorts 1 (subjects 1 - 64) and 2 (subjects 66 - 123) had the full filter analyzed, the remaining subjects had the mouthpiece analyzed.

Appendix 15.3.6.17

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_filter.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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15.2.5 Compliance

**Table 15.2.5.1 Summary of Compliance as Measured by Exhaled CO (ppm) in the SA Arm- Safety Population**

Timepoint	Statistic	SA (N=39)
Day 1 08:00-10:00 AM	n	39
	n (%) <=10ppm	31 (79.5%)
	n (%) >10ppm	8 (20.5%)
Day 1 12:00-02:00 PM	n	39
	n (%) <=10ppm	36 (92.3%)
	n (%) >10ppm	3 (7.7%)
Day 1 04:00-06:00 PM	n	39
	n (%) <=10ppm	39 (100 %)
	n (%) >10ppm	0
Day 1 08:00-10:00 PM	n	39
	n (%) <=10ppm	39 (100 %)
	n (%) >10ppm	0

Note: SA = Smoking abstinence.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_comp.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.5.1 Summary of Compliance as Measured by Exhaled CO (ppm) in the SA Arm- Safety Population**

Timepoint	Statistic	SA (N=39)
Day 2 08:00-10:00 AM	n	39
	n (%) ≤10ppm	39 (100 %)
	n (%) >10ppm	0
Day 2 12:00-02:00 PM	n	39
	n (%) ≤10ppm	39 (100 %)
	n (%) >10ppm	0
Day 2 04:00-06:00 PM	n	39
	n (%) ≤10ppm	39 (100 %)
	n (%) >10ppm	0
Day 2 08:00-10:00 PM	n	39
	n (%) ≤10ppm	39 (100 %)
	n (%) >10ppm	0

Note: SA = Smoking abstinence.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_comp.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.5.1 Summary of Compliance as Measured by Exhaled CO (ppm) in the SA Arm- Safety Population**

Timepoint	Statistic	SA (N=39)
Day 3 08:00-10:00 AM	n	39
	n (%) ≤10ppm	39 (100 %)
	n (%) >10ppm	0
Day 3 12:00-02:00 PM	n	39
	n (%) ≤10ppm	39 (100 %)
	n (%) >10ppm	0
Day 3 04:00-06:00 PM	n	39
	n (%) ≤10ppm	39 (100 %)
	n (%) >10ppm	0
Day 3 08:00-10:00 PM	n	39
	n (%) ≤10ppm	39 (100 %)
	n (%) >10ppm	0

Note: SA = Smoking abstinence.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_comp.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.5.1 Summary of Compliance as Measured by Exhaled CO (ppm) in the SA Arm- Safety Population**

Timepoint	Statistic	SA (N=39)
Day 4 08:00-10:00 AM	n	39
	n (%) ≤10ppm	39 (100 %)
	n (%) >10ppm	0
Day 4 12:00-02:00 PM	n	39
	n (%) ≤10ppm	39 (100 %)
	n (%) >10ppm	0
Day 4 04:00-06:00 PM	n	39
	n (%) ≤10ppm	39 (100 %)
	n (%) >10ppm	0
Day 4 08:00-10:00 PM	n	39
	n (%) ≤10ppm	39 (100 %)
	n (%) >10ppm	0

Note: SA = Smoking abstinence.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_comp.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.5.1 Summary of Compliance as Measured by Exhaled CO (ppm) in the SA Arm- Safety Population**

Timepoint	Statistic	SA (N=39)
Day 5 08:00-10:00 AM	n	39
	n (%) ≤10ppm	39 (100 %)
	n (%) >10ppm	0
Day 5 12:00-02:00 PM	n	39
	n (%) ≤10ppm	39 (100 %)
	n (%) >10ppm	0
Day 5 04:00-06:00 PM	n	39
	n (%) ≤10ppm	39 (100 %)
	n (%) >10ppm	0
Day 5 08:00-10:00 PM	n	39
	n (%) ≤10ppm	39 (100 %)
	n (%) >10ppm	0

Note: SA = Smoking abstinence.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_comp.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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15.2.6 Safety Summary Tables

**Table 15.2.6.1 Summary of Adverse Events - Safety Population**

	THS 2.2 (N=80)			CC (N=41)			SA (N=39)			Exposed Not Randomized (N=9)			Overall Safety (N=169)		
	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events
Adverse events (AE)	50	(62.5%)	98	29	(70.7%)	63	24	(61.5%)	49	9	(100 %)	17	112	(66.3%)	227
AE serious															
Yes	0			0			0			0			0		
No	50	(62.5%)	98	29	(70.7%)	63	24	(61.5%)	49	9	(100 %)	17	112	(66.3%)	227
AE related to THS 2.2/CC															
Not related	36	(45.0%)	83	22	(53.7%)	54	24	(61.5%)	49	7	(77.8%)	15	89	(52.7%)	201
Related	14	(17.5%)	15	7	(17.1%)	9	0			2	(22.2%)	2	23	(13.6%)	26
Expected	3	(3.8%)	3	7	(17.1%)	9				2	(22.2%)	2	12	(7.1%)	14
Not expected	11	(13.8%)	12	0						0			11	(6.5%)	12

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Exposed Not Randomized refers to all subjects exposed to THS 2.2 but not randomized. Overall Safety refers to all subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.6.1.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_adv1.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.1 Summary of Adverse Events - Safety Population**

	THS 2.2 (N=80)			CC (N=41)			SA (N=39)			Exposed Not Randomized (N=9)			Overall Safety (N=169)		
	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events
AE related to study procedure															
Not related	39	(48.8%)	87	23	(56.1%)	55	23	(59.0%)	48	7	(77.8%)	15	92	(54.4%)	205
Related	11	(13.8%)	11	6	(14.6%)	8	1	(2.6%)	1	2	(22.2%)	2	20	(11.8%)	22
AE severity															
Mild	28	(35.0%)	71	13	(31.7%)	42	11	(28.2%)	29	3	(33.3%)	10	55	(32.5%)	152
Moderate	22	(27.5%)	27	15	(36.6%)	20	13	(33.3%)	20	6	(66.7%)	7	56	(33.1%)	74
Severe	0			1	(2.4%)	1	0			0			1	(0.6%)	1

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Exposed Not Randomized refers to all subjects exposed to THS 2.2 but not randomized. Overall Safety refers to all subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.6.1.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_adv1.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.1 Summary of Adverse Events - Safety Population**

	THS 2.2 (N=80)			CC (N=41)			SA (N=39)			Exposed Not Randomized (N=9)			Overall Safety (N=169)		
	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events
Action taken due to AE															
Product use interrupted	0			0			0			0			0		
Product use stopped	0			0			0			0			0		
Product use reduced	0			1 (2.4%)		1	0			0			1 (0.6%)		1
Not applicable	0			0			22 (56.4%)		37	2 (22.2%)		5	24 (14.2%)		42
None	50 (62.5%)		98	29 (70.7%)		62	8 (20.5%)		12	7 (77.8%)		12	94 (55.6%)		184
Treatment given															
Yes	10 (12.5%)		12	9 (22.0%)		10	8 (20.5%)		11	3 (33.3%)		5	30 (17.8%)		38
No	40 (50.0%)		86	20 (48.8%)		53	16 (41.0%)		38	6 (66.7%)		12	82 (48.5%)		189
Other action taken	0			2 (4.9%)		2	1 (2.6%)		1	9 (100 %)		9	12 (7.1%)		12

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Exposed Not Randomized refers to all subjects exposed to THS 2.2 but not randomized. Overall Safety refers to all subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.6.1.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_adv1.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.2.1 Summary of Adverse Events by System Organ Class and Preferred Term - Safety Population**

System Organ Class Preferred Term	THS 2.2 (N=80)			CC (N=41)			SA (N=39)			Exposed Not Randomized (N=9)			Overall Safety (N=169)		
	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events
Any adverse events	50	(62.5%)	98	29	(70.7%)	63	24	(61.5%)	49	9	(100 %)	17	112	(66.3%)	227
Nervous system disorders	27	(33.8%)	36	19	(46.3%)	23	13	(33.3%)	15	3	(33.3%)	3	62	(36.7%)	77
Headache	24	(30.0%)	30	16	(39.0%)	19	13	(33.3%)	15	3	(33.3%)	3	56	(33.1%)	67
Syncope	6	(7.5%)	6	4	(9.8%)	4	0			0			10	(5.9%)	10
Investigations	7	(8.8%)	7	6	(14.6%)	8	3	(7.7%)	3	3	(33.3%)	4	19	(11.2%)	22
Spirometry abnormal	4	(5.0%)	4	3	(7.3%)	3	2	(5.1%)	2	3	(33.3%)	3	12	(7.1%)	12
Lymphocyte count increased	1	(1.3%)	1	0			1	(2.6%)	1	1	(11.1%)	1	3	(1.8%)	3
Carboxyhaemoglobin increased	0			2	(4.9%)	4	0			0			2	(1.2%)	4
Blood bilirubin increased	1	(1.3%)	1	0			0			0			1	(0.6%)	1
Eosinophil count increased	1	(1.3%)	1	0			0			0			1	(0.6%)	1
Neutrophil count decreased	0			1	(2.4%)	1	0			0			1	(0.6%)	1

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Exposed Not Randomized refers to all subjects exposed to THS 2.2 but not randomized. Overall Safety refers to all subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.6.1.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_adv2.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.2.1 Summary of Adverse Events by System Organ Class and Preferred Term - Safety Population**

System Organ Class Preferred Term	THS 2.2 (N=80)			CC (N=41)			SA (N=39)			Exposed Not Randomized (N=9)			Overall Safety (N=169)		
	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events
Gastrointestinal disorders	7	(8.8%)	7	5	(12.2%)	6	3	(7.7%)	5	2	(22.2%)	2	17	(10.1%)	20
Constipation	3	(3.8%)	3	2	(4.9%)	2	0			1	(11.1%)	1	6	(3.6%)	6
Toothache	1	(1.3%)	1	1	(2.4%)	1	0			1	(11.1%)	1	3	(1.8%)	3
Gastrooesophageal reflux disease	0			2	(4.9%)	3	0			0			2	(1.2%)	3
Abdominal distension	0			0			2	(5.1%)	2	0			2	(1.2%)	2
Mouth ulceration	2	(2.5%)	2	0			0			0			2	(1.2%)	2
Abdominal pain upper	0			0			1	(2.6%)	1	0			1	(0.6%)	1
Nausea	0			0			1	(2.6%)	1	0			1	(0.6%)	1
Oral pain	1	(1.3%)	1	0			0			0			1	(0.6%)	1
Vomiting	0			0			1	(2.6%)	1	0			1	(0.6%)	1

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Exposed Not Randomized refers to all subjects exposed to THS 2.2 but not randomized. Overall Safety refers to all subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.6.1.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_adv2.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.2.1 Summary of Adverse Events by System Organ Class and Preferred Term - Safety Population**

System Organ Class Preferred Term	THS 2.2 (N=80)			CC (N=41)			SA (N=39)			Exposed Not Randomized (N=9)			Overall Safety (N=169)		
	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events
Respiratory, thoracic and mediastinal disorders	10	(12.5%)	10	4	(9.8%)	4	0			1	(11.1%)	1	15	(8.9%)	15
Oropharyngeal pain	7	(8.8%)	7	3	(7.3%)	3				0			10	(5.9%)	10
Cough	3	(3.8%)	3	0						1	(11.1%)	1	4	(2.4%)	4
Epistaxis	0			1	(2.4%)	1				0			1	(0.6%)	1
Renal and urinary disorders	7	(8.8%)	9	4	(9.8%)	6	2	(5.1%)	2	0			13	(7.7%)	17
Polyuria	6	(7.5%)	8	4	(9.8%)	6	2	(5.1%)	2				12	(7.1%)	16
Leukocyturia	1	(1.3%)	1	0			0						1	(0.6%)	1
Metabolism and nutrition disorders	4	(5.0%)	4	3	(7.3%)	3	4	(10.3%)	4	2	(22.2%)	2	13	(7.7%)	13
Hypertriglyceridaemia	3	(3.8%)	3	2	(4.9%)	2	2	(5.1%)	2	0			7	(4.1%)	7
Hypercholesterolaemia	0			1	(2.4%)	1	1	(2.6%)	1	0			2	(1.2%)	2
Hyperglycaemia	0			0			1	(2.6%)	1	1	(11.1%)	1	2	(1.2%)	2

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Exposed Not Randomized refers to all subjects exposed to THS 2.2 but not randomized. Overall Safety refers to all subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.6.1.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_adv2.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.2.1 Summary of Adverse Events by System Organ Class and Preferred Term - Safety Population**

System Organ Class Preferred Term	THS 2.2 (N=80)			CC (N=41)			SA (N=39)			Exposed Not Randomized (N=9)			Overall Safety (N=169)		
	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events
Hyperkalaemia	1	(1.3%)	1	0			0			1	(11.1%)	1	2	(1.2%)	2
Musculoskeletal and connective tissue disorders	6	(7.5%)	6	2	(4.9%)	3	4	(10.3%)	4	0			12	(7.1%)	13
Back pain	3	(3.8%)	3	2	(4.9%)	3	3	(7.7%)	3				8	(4.7%)	9
Arthralgia	1	(1.3%)	1	0			0						1	(0.6%)	1
Musculoskeletal chest pain	1	(1.3%)	1	0			0						1	(0.6%)	1
Myalgia	0			0			1	(2.6%)	1				1	(0.6%)	1
Pain in extremity	1	(1.3%)	1	0			0						1	(0.6%)	1
Vascular disorders	3	(3.8%)	4	2	(4.9%)	2	3	(7.7%)	7	2	(22.2%)	2	10	(5.9%)	15
Hypertension	1	(1.3%)	2	1	(2.4%)	1	2	(5.1%)	5	2	(22.2%)	2	6	(3.6%)	10
Hypotension	2	(2.5%)	2	1	(2.4%)	1	1	(2.6%)	2	0			4	(2.4%)	5

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Exposed Not Randomized refers to all subjects exposed to THS 2.2 but not randomized. Overall Safety refers to all subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.6.1.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_adv2.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.2.1 Summary of Adverse Events by System Organ Class and Preferred Term - Safety Population**

System Organ Class Preferred Term	THS 2.2 (N=80)			CC (N=41)			SA (N=39)			Exposed Not Randomized (N=9)			Overall Safety (N=169)		
	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events
Infections and infestations	4	(5.0%)	4	2	(4.9%)	3	1	(2.6%)	1	0			7	(4.1%)	8
Nasopharyngitis	3	(3.8%)	3	1	(2.4%)	2	1	(2.6%)	1				5	(3.0%)	6
Herpes virus infection	0			1	(2.4%)	1	0						1	(0.6%)	1
Urinary tract infection	1	(1.3%)	1	0			0						1	(0.6%)	1
General disorders and administration site conditions	2	(2.5%)	2	1	(2.4%)	1	3	(7.7%)	3	0			6	(3.6%)	6
Influenza like illness	0			1	(2.4%)	1	3	(7.7%)	3				4	(2.4%)	4
Vessel puncture site reaction	2	(2.5%)	2	0			0						2	(1.2%)	2
Ear and labyrinth disorders	3	(3.8%)	3	0			2	(5.1%)	2	0			5	(3.0%)	5
Vertigo	3	(3.8%)	3				2	(5.1%)	2				5	(3.0%)	5

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Exposed Not Randomized refers to all subjects exposed to THS 2.2 but not randomized. Overall Safety refers to all subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.6.1.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_adv2.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.2.1 Summary of Adverse Events by System Organ Class and Preferred Term - Safety Population**

System Organ Class Preferred Term	THS 2.2 (N=80)			CC (N=41)			SA (N=39)			Exposed Not Randomized (N=9)			Overall Safety (N=169)		
	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events
Reproductive system and breast disorders	3	(3.8%)	3	1	(2.4%)	1	1	(2.6%)	1	0			5	(3.0%)	5
Dysmenorrhoea	3	(3.8%)	3	1	(2.4%)	1	1	(2.6%)	1				5	(3.0%)	5
Cardiac disorders	1	(1.3%)	1	2	(4.9%)	3	0			1	(11.1%)	1	4	(2.4%)	5
Tachycardia	0			2	(4.9%)	3				0			2	(1.2%)	3
Myocardial ischaemia	1	(1.3%)	1	0						0			1	(0.6%)	1
Ventricular extrasystoles	0			0						1	(11.1%)	1	1	(0.6%)	1
Blood and lymphatic system disorders	0			0			0			2	(22.2%)	2	2	(1.2%)	2
Leukocytosis										2	(22.2%)	2	2	(1.2%)	2
Eye disorders	0			0			1	(2.6%)	1	0			1	(0.6%)	1
Conjunctivitis							1	(2.6%)	1				1	(0.6%)	1

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Exposed Not Randomized refers to all subjects exposed to THS 2.2 but not randomized. Overall Safety refers to all subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.6.1.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_adv2.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.2.1 Summary of Adverse Events by System Organ Class and Preferred Term - Safety Population**

System Organ Class Preferred Term	THS 2.2 (N=80)			CC (N=41)			SA (N=39)			Exposed Not Randomized (N=9)			Overall Safety (N=169)		
	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events
Hepatobiliary disorders	1	(1.3%)	1	0			0			0			1	(0.6%)	1
Hyperbilirubinaemia	1	(1.3%)	1										1	(0.6%)	1
Psychiatric disorders	0			0			1	(2.6%)	1	0			1	(0.6%)	1
Nicotine dependence							1	(2.6%)	1				1	(0.6%)	1
Skin and subcutaneous tissue disorders	1	(1.3%)	1	0			0			0			1	(0.6%)	1
Hyperhidrosis	1	(1.3%)	1										1	(0.6%)	1

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Exposed Not Randomized refers to all subjects exposed to THS 2.2 but not randomized. Overall Safety refers to all subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.6.1.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_adv2.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.2.2 Summary of Adverse Events by System Organ Class - Safety Population**

System Organ Class	THS 2.2 (N=80)			CC (N=41)			SA (N=39)			Exposed Not Randomized (N=9)			Overall Safety (N=169)		
	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events
Any adverse events	50	(62.5%)	98	29	(70.7%)	63	24	(61.5%)	49	9	(100 %)	17	112	(66.3%)	227
Nervous system disorders	27	(33.8%)	36	19	(46.3%)	23	13	(33.3%)	15	3	(33.3%)	3	62	(36.7%)	77
Investigations	7	(8.8%)	7	6	(14.6%)	8	3	(7.7%)	3	3	(33.3%)	4	19	(11.2%)	22
Gastrointestinal disorders	7	(8.8%)	7	5	(12.2%)	6	3	(7.7%)	5	2	(22.2%)	2	17	(10.1%)	20
Respiratory, thoracic and mediastinal disorders	10	(12.5%)	10	4	(9.8%)	4	0			1	(11.1%)	1	15	(8.9%)	15
Renal and urinary disorders	7	(8.8%)	9	4	(9.8%)	6	2	(5.1%)	2	0			13	(7.7%)	17
Metabolism and nutrition disorders	4	(5.0%)	4	3	(7.3%)	3	4	(10.3%)	4	2	(22.2%)	2	13	(7.7%)	13
Musculoskeletal and connective tissue disorders	6	(7.5%)	6	2	(4.9%)	3	4	(10.3%)	4	0			12	(7.1%)	13
Vascular disorders	3	(3.8%)	4	2	(4.9%)	2	3	(7.7%)	7	2	(22.2%)	2	10	(5.9%)	15
Infections and infestations	4	(5.0%)	4	2	(4.9%)	3	1	(2.6%)	1	0			7	(4.1%)	8

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Exposed Not Randomized refers to all subjects exposed to THS 2.2 but not randomized. Overall Safety refers to all subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.6.1.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_adv3.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.2.2 Summary of Adverse Events by System Organ Class - Safety Population**

System Organ Class	THS 2.2 (N=80)			CC (N=41)			SA (N=39)			Exposed Not Randomized (N=9)			Overall Safety (N=169)		
	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events
General disorders and administration site conditions	2	(2.5%)	2	1	(2.4%)	1	3	(7.7%)	3	0			6	(3.6%)	6
Ear and labyrinth disorders	3	(3.8%)	3	0			2	(5.1%)	2	0			5	(3.0%)	5
Reproductive system and breast disorders	3	(3.8%)	3	1	(2.4%)	1	1	(2.6%)	1	0			5	(3.0%)	5
Cardiac disorders	1	(1.3%)	1	2	(4.9%)	3	0			1	(11.1%)	1	4	(2.4%)	5
Blood and lymphatic system disorders	0			0			0			2	(22.2%)	2	2	(1.2%)	2
Eye disorders	0			0			1	(2.6%)	1	0			1	(0.6%)	1
Hepatobiliary disorders	1	(1.3%)	1	0			0			0			1	(0.6%)	1
Psychiatric disorders	0			0			1	(2.6%)	1	0			1	(0.6%)	1
Skin and subcutaneous tissue disorders	1	(1.3%)	1	0			0			0			1	(0.6%)	1

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Exposed Not Randomized refers to all subjects exposed to THS 2.2 but not randomized. Overall Safety refers to all subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.6.1.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_adv3.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.2.3 Summary of Adverse Events by Preferred Term - Safety Population**

Preferred Term	THS 2.2 (N=80)			CC (N=41)			SA (N=39)			Exposed Not Randomized (N=9)			Overall Safety (N=169)		
	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events
Any adverse events	50	(62.5%)	98	29	(70.7%)	63	24	(61.5%)	49	9	(100 %)	17	112	(66.3%)	227
Headache	24	(30.0%)	30	16	(39.0%)	19	13	(33.3%)	15	3	(33.3%)	3	56	(33.1%)	67
Polyuria	6	(7.5%)	8	4	(9.8%)	6	2	(5.1%)	2	0			12	(7.1%)	16
Spirometry abnormal	4	(5.0%)	4	3	(7.3%)	3	2	(5.1%)	2	3	(33.3%)	3	12	(7.1%)	12
Oropharyngeal pain	7	(8.8%)	7	3	(7.3%)	3	0			0			10	(5.9%)	10
Syncope	6	(7.5%)	6	4	(9.8%)	4	0			0			10	(5.9%)	10
Back pain	3	(3.8%)	3	2	(4.9%)	3	3	(7.7%)	3	0			8	(4.7%)	9
Hypertriglyceridaemia	3	(3.8%)	3	2	(4.9%)	2	2	(5.1%)	2	0			7	(4.1%)	7
Hypertension	1	(1.3%)	2	1	(2.4%)	1	2	(5.1%)	5	2	(22.2%)	2	6	(3.6%)	10
Constipation	3	(3.8%)	3	2	(4.9%)	2	0			1	(11.1%)	1	6	(3.6%)	6
Nasopharyngitis	3	(3.8%)	3	1	(2.4%)	2	1	(2.6%)	1	0			5	(3.0%)	6
Dysmenorrhoea	3	(3.8%)	3	1	(2.4%)	1	1	(2.6%)	1	0			5	(3.0%)	5
Vertigo	3	(3.8%)	3	0			2	(5.1%)	2	0			5	(3.0%)	5
Hypotension	2	(2.5%)	2	1	(2.4%)	1	1	(2.6%)	2	0			4	(2.4%)	5

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Exposed Not Randomized refers to all subjects exposed to THS 2.2 but not randomized. Overall Safety refers to all subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.6.1.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_adv4.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.2.3 Summary of Adverse Events by Preferred Term - Safety Population**

Preferred Term	THS 2.2 (N=80)			CC (N=41)			SA (N=39)			Exposed Not Randomized (N=9)			Overall Safety (N=169)		
	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events
Cough	3	(3.8%)	3	0			0			1	(11.1%)	1	4	(2.4%)	4
Influenza like illness	0			1	(2.4%)	1	3	(7.7%)	3	0			4	(2.4%)	4
Lymphocyte count increased	1	(1.3%)	1	0			1	(2.6%)	1	1	(11.1%)	1	3	(1.8%)	3
Toothache	1	(1.3%)	1	1	(2.4%)	1	0			1	(11.1%)	1	3	(1.8%)	3
Carboxyhaemoglobin increased	0			2	(4.9%)	4	0			0			2	(1.2%)	4
Gastroesophageal reflux disease	0			2	(4.9%)	3	0			0			2	(1.2%)	3
Tachycardia	0			2	(4.9%)	3	0			0			2	(1.2%)	3
Abdominal distension	0			0			2	(5.1%)	2	0			2	(1.2%)	2
Hypercholesterolaemia	0			1	(2.4%)	1	1	(2.6%)	1	0			2	(1.2%)	2
Hyperglycaemia	0			0			1	(2.6%)	1	1	(11.1%)	1	2	(1.2%)	2
Hyperkalaemia	1	(1.3%)	1	0			0			1	(11.1%)	1	2	(1.2%)	2
Leukocytosis	0			0			0			2	(22.2%)	2	2	(1.2%)	2
Mouth ulceration	2	(2.5%)	2	0			0			0			2	(1.2%)	2
Vessel puncture site reaction	2	(2.5%)	2	0			0			0			2	(1.2%)	2

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Exposed Not Randomized refers to all subjects exposed to THS 2.2 but not randomized. Overall Safety refers to all subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.6.1.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_adv4.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.2.3 Summary of Adverse Events by Preferred Term - Safety Population**

Preferred Term	THS 2.2 (N=80)			CC (N=41)			SA (N=39)			Exposed Not Randomized (N=9)			Overall Safety (N=169)		
	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events
Abdominal pain upper	0			0			1 (2.6%)	1		0			1 (0.6%)	1	
Arthralgia	1 (1.3%)		1	0			0			0			1 (0.6%)	1	
Blood bilirubin increased	1 (1.3%)		1	0			0			0			1 (0.6%)	1	
Conjunctivitis	0			0			1 (2.6%)	1		0			1 (0.6%)	1	
Eosinophil count increased	1 (1.3%)		1	0			0			0			1 (0.6%)	1	
Epistaxis	0			1 (2.4%)		1	0			0			1 (0.6%)	1	
Herpes virus infection	0			1 (2.4%)		1	0			0			1 (0.6%)	1	
Hyperbilirubinaemia	1 (1.3%)		1	0			0			0			1 (0.6%)	1	
Hyperhidrosis	1 (1.3%)		1	0			0			0			1 (0.6%)	1	
Leukocyturia	1 (1.3%)		1	0			0			0			1 (0.6%)	1	
Musculoskeletal chest pain	1 (1.3%)		1	0			0			0			1 (0.6%)	1	
Myalgia	0			0			1 (2.6%)	1		0			1 (0.6%)	1	
Myocardial ischaemia	1 (1.3%)		1	0			0			0			1 (0.6%)	1	
Nausea	0			0			1 (2.6%)	1		0			1 (0.6%)	1	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Exposed Not Randomized refers to all subjects exposed to THS 2.2 but not randomized. Overall Safety refers to all subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.6.1.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_adv4.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.2.3 Summary of Adverse Events by Preferred Term - Safety Population**

Preferred Term	THS 2.2 (N=80)			CC (N=41)			SA (N=39)			Exposed Not Randomized (N=9)			Overall Safety (N=169)		
	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events
Neutrophil count decreased	0			1 (2.4%)	1		0			0			1 (0.6%)	1	
Nicotine dependence	0			0			1 (2.6%)	1		0			1 (0.6%)	1	
Oral pain	1 (1.3%)	1		0			0			0			1 (0.6%)	1	
Pain in extremity	1 (1.3%)	1		0			0			0			1 (0.6%)	1	
Urinary tract infection	1 (1.3%)	1		0			0			0			1 (0.6%)	1	
Ventricular extrasystoles	0			0			0			1 (11.1%)	1		1 (0.6%)	1	
Vomiting	0			0			1 (2.6%)	1		0			1 (0.6%)	1	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Exposed Not Randomized refers to all subjects exposed to THS 2.2 but not randomized. Overall Safety refers to all subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.6.1.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_adv4.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.3 Summary of Adverse Events by System Organ Class, Preferred Term and Relationship to Study Product Exposure and Expectedness - Safety Population**

	THS 2.2 (N=80)			CC (N=41)			SA (N=39)			Exposed Not Randomized (N=9)			Overall Safety (N=169)		
	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events
Adverse events (AE)	50	(62.5%)	98	29	(70.7%)	63	24	(61.5%)	49	9	(100 %)	17	112	(66.3%)	227
Related to THS 2.2/CC	14	(17.5%)	15	7	(17.1%)	9	0			2	(22.2%)	2	23	(13.6%)	26
Expected	3	(3.8%)	3	7	(17.1%)	9				2	(22.2%)	2	12	(7.1%)	14
Nervous system disorders	27	(33.8%)	36	19	(46.3%)	23	13	(33.3%)	15	3	(33.3%)	3	62	(36.7%)	77
(Headache)	24	(30.0%)	30	16	(39.0%)	19	13	(33.3%)	15	3	(33.3%)	3	56	(33.1%)	67
Related to THS 2.2/CC	0			0			0			0			0		
(Syncope)	6	(7.5%)	6	4	(9.8%)	4							10	(5.9%)	10
Related to THS 2.2/CC	1	(1.3%)	1	2	(4.9%)	2							3	(1.8%)	3
Expected	0			2	(4.9%)	2							2	(1.2%)	2

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Exposed Not Randomized refers to all subjects exposed to THS 2.2 but not randomized. Overall Safety refers to all subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.6.1.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_adv10.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.3 Summary of Adverse Events by System Organ Class, Preferred Term and Relationship to Study Product Exposure and Expectedness - Safety Population**

	THS 2.2 (N=80)			CC (N=41)			SA (N=39)			Exposed Not Randomized (N=9)			Overall Safety (N=169)		
	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events
Investigations	7	(8.8%)	7	6	(14.6%)	8	3	(7.7%)	3	3	(33.3%)	4	19	(11.2%)	22
(Spirometry abnormal)	4	(5.0%)	4	3	(7.3%)	3	2	(5.1%)	2	3	(33.3%)	3	12	(7.1%)	12
Related to THS 2.2/CC	4	(5.0%)	4	3	(7.3%)	3	0			2	(22.2%)	2	9	(5.3%)	9
Expected	3	(3.8%)	3	3	(7.3%)	3				2	(22.2%)	2	8	(4.7%)	8
(Lymphocyte count increased)	1	(1.3%)	1				1	(2.6%)	1	1	(11.1%)	1	3	(1.8%)	3
Related to THS 2.2/CC	0						0			0			0		
(Carboxyhaemoglobin increased)				2	(4.9%)	4							2	(1.2%)	4
Related to THS 2.2/CC				2	(4.9%)	4							2	(1.2%)	4
Expected				2	(4.9%)	4							2	(1.2%)	4

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Exposed Not Randomized refers to all subjects exposed to THS 2.2 but not randomized. Overall Safety refers to all subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.6.1.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_adv10.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.3 Summary of Adverse Events by System Organ Class, Preferred Term and Relationship to Study Product Exposure and Expectedness - Safety Population**

	THS 2.2 (N=80)			CC (N=41)			SA (N=39)			Exposed Not Randomized (N=9)			Overall Safety (N=169)		
	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events
(Blood bilirubin increased)	1	(1.3%)	1										1	(0.6%)	1
Related to THS 2.2/CC	0												0		
(Eosinophil count increased)	1	(1.3%)	1										1	(0.6%)	1
Related to THS 2.2/CC	0												0		
(Neutrophil count decreased)				1	(2.4%)	1							1	(0.6%)	1
Related to THS 2.2/CC				0									0		
Gastrointestinal disorders	7	(8.8%)	7	5	(12.2%)	6	3	(7.7%)	5	2	(22.2%)	2	17	(10.1%)	20
(Constipation)	3	(3.8%)	3	2	(4.9%)	2				1	(11.1%)	1	6	(3.6%)	6
Related to THS 2.2/CC	0			0						0			0		

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Exposed Not Randomized refers to all subjects exposed to THS 2.2 but not randomized. Overall Safety refers to all subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.6.1.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_adv10.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.3 Summary of Adverse Events by System Organ Class, Preferred Term and Relationship to Study Product Exposure and Expectedness - Safety Population**

	THS 2.2 (N=80)			CC (N=41)			SA (N=39)			Exposed Not Randomized (N=9)			Overall Safety (N=169)		
	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events
(Toothache)	1	(1.3%)	1	1	(2.4%)	1				1	(11.1%)	1	3	(1.8%)	3
Related to THS 2.2/CC	0			0						0			0		
(Gastrooesophageal reflux disease)				2	(4.9%)	3							2	(1.2%)	3
Related to THS 2.2/CC				0									0		
(Abdominal distension)							2	(5.1%)	2				2	(1.2%)	2
Related to THS 2.2/CC							0						0		
(Mouth ulceration)	2	(2.5%)	2										2	(1.2%)	2
Related to THS 2.2/CC	0												0		

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Exposed Not Randomized refers to all subjects exposed to THS 2.2 but not randomized. Overall Safety refers to all subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.6.1.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_adv10.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.3 Summary of Adverse Events by System Organ Class, Preferred Term and Relationship to Study Product Exposure and Expectedness - Safety Population**

	THS 2.2 (N=80)			CC (N=41)			SA (N=39)			Exposed Not Randomized (N=9)			Overall Safety (N=169)		
	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events
(Abdominal pain upper)							1	(2.6%)	1				1	(0.6%)	1
Related to THS 2.2/CC							0						0		
(Nausea)							1	(2.6%)	1				1	(0.6%)	1
Related to THS 2.2/CC							0						0		
(Oral pain)	1	(1.3%)	1										1	(0.6%)	1
Related to THS 2.2/CC	1	(1.3%)	1										1	(0.6%)	1
Expected	0												0		
(Vomiting)							1	(2.6%)	1				1	(0.6%)	1
Related to THS 2.2/CC							0						0		

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Exposed Not Randomized refers to all subjects exposed to THS 2.2 but not randomized. Overall Safety refers to all subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.6.1.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_adv10.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.3 Summary of Adverse Events by System Organ Class, Preferred Term and Relationship to Study Product Exposure and Expectedness - Safety Population**

	THS 2.2 (N=80)			CC (N=41)			SA (N=39)			Exposed Not Randomized (N=9)			Overall Safety (N=169)		
	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events
Respiratory, thoracic and mediastinal disorders	10	(12.5%)	10	4	(9.8%)	4	0			1	(11.1%)	1	15	(8.9%)	15
(Oropharyngeal pain)	7	(8.8%)	7	3	(7.3%)	3							10	(5.9%)	10
Related to THS 2.2/CC	1	(1.3%)	1	0									1	(0.6%)	1
Expected	0												0		
(Cough)	3	(3.8%)	3							1	(11.1%)	1	4	(2.4%)	4
Related to THS 2.2/CC	2	(2.5%)	2							0			2	(1.2%)	2
Expected	0												0		
(Epistaxis)				1	(2.4%)	1							1	(0.6%)	1
Related to THS 2.2/CC				0									0		

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Exposed Not Randomized refers to all subjects exposed to THS 2.2 but not randomized. Overall Safety refers to all subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.6.1.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_adv10.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.3 Summary of Adverse Events by System Organ Class, Preferred Term and Relationship to Study Product Exposure and Expectedness - Safety Population**

	THS 2.2 (N=80)			CC (N=41)			SA (N=39)			Exposed Not Randomized (N=9)			Overall Safety (N=169)		
	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events
Renal and urinary disorders	7	(8.8%)	9	4	(9.8%)	6	2	(5.1%)	2	0			13	(7.7%)	17
(Polyuria)	6	(7.5%)	8	4	(9.8%)	6	2	(5.1%)	2				12	(7.1%)	16
Related to THS 2.2/CC	0			0			0						0		
(Leukocyturia)	1	(1.3%)	1										1	(0.6%)	1
Related to THS 2.2/CC	0												0		
Metabolism and nutrition disorders	4	(5.0%)	4	3	(7.3%)	3	4	(10.3%)	4	2	(22.2%)	2	13	(7.7%)	13
(Hypertriglyceridaemia)	3	(3.8%)	3	2	(4.9%)	2	2	(5.1%)	2				7	(4.1%)	7
Related to THS 2.2/CC	1	(1.3%)	1	0			0						1	(0.6%)	1
Expected	0												0		

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Exposed Not Randomized refers to all subjects exposed to THS 2.2 but not randomized. Overall Safety refers to all subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.6.1.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_adv10.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.3 Summary of Adverse Events by System Organ Class, Preferred Term and Relationship to Study Product Exposure and Expectedness - Safety Population**

	THS 2.2 (N=80)			CC (N=41)			SA (N=39)			Exposed Not Randomized (N=9)			Overall Safety (N=169)		
	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events
(Hypercholesterolaemia)				1 (2.4%)	1		1 (2.6%)	1					2 (1.2%)	2	
Related to THS 2.2/CC				0			0						0		
(Hyperglycaemia)							1 (2.6%)	1		1 (11.1%)	1		2 (1.2%)	2	
Related to THS 2.2/CC							0			0			0		
(Hyperkalaemia)	1 (1.3%)		1							1 (11.1%)	1		2 (1.2%)	2	
Related to THS 2.2/CC	0									0			0		
Musculoskeletal and connective tissue disorders	6 (7.5%)		6	2 (4.9%)	3		4 (10.3%)	4		0			12 (7.1%)	13	
(Back pain)	3 (3.8%)		3	2 (4.9%)	3		3 (7.7%)	3					8 (4.7%)	9	
Related to THS 2.2/CC	0			0			0						0		

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Exposed Not Randomized refers to all subjects exposed to THS 2.2 but not randomized. Overall Safety refers to all subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.6.1.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_adv10.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.3 Summary of Adverse Events by System Organ Class, Preferred Term and Relationship to Study Product Exposure and Expectedness - Safety Population**

	THS 2.2 (N=80)			CC (N=41)			SA (N=39)			Exposed Not Randomized (N=9)			Overall Safety (N=169)		
	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events
(Arthralgia)	1	(1.3%)	1										1	(0.6%)	1
Related to THS 2.2/CC	0												0		
(Musculoskeletal chest pain)	1	(1.3%)	1										1	(0.6%)	1
Related to THS 2.2/CC	1	(1.3%)	1										1	(0.6%)	1
Expected	0												0		
(Myalgia)							1	(2.6%)	1				1	(0.6%)	1
Related to THS 2.2/CC							0						0		
(Pain in extremity)	1	(1.3%)	1										1	(0.6%)	1
Related to THS 2.2/CC	0												0		

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Exposed Not Randomized refers to all subjects exposed to THS 2.2 but not randomized. Overall Safety refers to all subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.6.1.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_adv10.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.3 Summary of Adverse Events by System Organ Class, Preferred Term and Relationship to Study Product Exposure and Expectedness - Safety Population**

	THS 2.2 (N=80)			CC (N=41)			SA (N=39)			Exposed Not Randomized (N=9)			Overall Safety (N=169)		
	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events
Vascular disorders	3	(3.8%)	4	2	(4.9%)	2	3	(7.7%)	7	2	(22.2%)	2	10	(5.9%)	15
(Hypertension)	1	(1.3%)	2	1	(2.4%)	1	2	(5.1%)	5	2	(22.2%)	2	6	(3.6%)	10
Related to THS 2.2/CC	0			0			0			0			0		
(Hypotension)	2	(2.5%)	2	1	(2.4%)	1	1	(2.6%)	2				4	(2.4%)	5
Related to THS 2.2/CC	0			0			0						0		
Infections and infestations	4	(5.0%)	4	2	(4.9%)	3	1	(2.6%)	1	0			7	(4.1%)	8
(Nasopharyngitis)	3	(3.8%)	3	1	(2.4%)	2	1	(2.6%)	1				5	(3.0%)	6
Related to THS 2.2/CC	0			0			0						0		

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Exposed Not Randomized refers to all subjects exposed to THS 2.2 but not randomized. Overall Safety refers to all subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.6.1.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_adv10.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.3 Summary of Adverse Events by System Organ Class, Preferred Term and Relationship to Study Product Exposure and Expectedness - Safety Population**

	THS 2.2 (N=80)			CC (N=41)			SA (N=39)			Exposed Not Randomized (N=9)			Overall Safety (N=169)		
	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events
(Herpes virus infection)				1	(2.4%)	1							1	(0.6%)	1
Related to THS 2.2/CC				0									0		
(Urinary tract infection)	1	(1.3%)	1										1	(0.6%)	1
Related to THS 2.2/CC	0												0		
General disorders and administration site conditions	2	(2.5%)	2	1	(2.4%)	1	3	(7.7%)	3	0			6	(3.6%)	6
(Influenza like illness)				1	(2.4%)	1	3	(7.7%)	3				4	(2.4%)	4
Related to THS 2.2/CC				0			0						0		
(Vessel puncture site reaction)	2	(2.5%)	2										2	(1.2%)	2
Related to THS 2.2/CC	0												0		

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Exposed Not Randomized refers to all subjects exposed to THS 2.2 but not randomized. Overall Safety refers to all subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.6.1.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_adv10.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.3 Summary of Adverse Events by System Organ Class, Preferred Term and Relationship to Study Product Exposure and Expectedness - Safety Population**

	THS 2.2 (N=80)			CC (N=41)			SA (N=39)			Exposed Not Randomized (N=9)			Overall Safety (N=169)		
	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events
Ear and labyrinth disorders	3	(3.8%)	3	0			2	(5.1%)	2	0			5	(3.0%)	5
(Vertigo)	3	(3.8%)	3				2	(5.1%)	2				5	(3.0%)	5
Related to THS 2.2/CC	2	(2.5%)	2				0						2	(1.2%)	2
Expected	0												0		
Reproductive system and breast disorders	3	(3.8%)	3	1	(2.4%)	1	1	(2.6%)	1	0			5	(3.0%)	5
(Dysmenorrhoea)	3	(3.8%)	3	1	(2.4%)	1	1	(2.6%)	1				5	(3.0%)	5
Related to THS 2.2/CC	0			0			0						0		
Cardiac disorders	1	(1.3%)	1	2	(4.9%)	3	0			1	(11.1%)	1	4	(2.4%)	5

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Exposed Not Randomized refers to all subjects exposed to THS 2.2 but not randomized. Overall Safety refers to all subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.6.1.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_adv10.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.3 Summary of Adverse Events by System Organ Class, Preferred Term and Relationship to Study Product Exposure and Expectedness - Safety Population**

	THS 2.2 (N=80)			CC (N=41)			SA (N=39)			Exposed Not Randomized (N=9)			Overall Safety (N=169)		
	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events
(Tachycardia)				2	(4.9%)	3							2	(1.2%)	3
Related to THS 2.2/CC				0									0		
(Myocardial ischaemia)	1	(1.3%)	1										1	(0.6%)	1
Related to THS 2.2/CC	1	(1.3%)	1										1	(0.6%)	1
Expected	0												0		
(Ventricular extrasystoles)										1	(11.1%)	1	1	(0.6%)	1
Related to THS 2.2/CC										0			0		
Blood and lymphatic system disorders	0			0			0			2	(22.2%)	2	2	(1.2%)	2

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Exposed Not Randomized refers to all subjects exposed to THS 2.2 but not randomized. Overall Safety refers to all subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.6.1.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_adv10.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.3 Summary of Adverse Events by System Organ Class, Preferred Term and Relationship to Study Product Exposure and Expectedness - Safety Population**

	THS 2.2 (N=80)			CC (N=41)			SA (N=39)			Exposed Not Randomized (N=9)			Overall Safety (N=169)		
	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events
(Leukocytosis) Related to THS 2.2/CC										2 (22.2%)	2		2 (1.2%)	2	
										0			0		
Eye disorders	0			0			1 (2.6%)	1		0			1 (0.6%)	1	
(Conjunctivitis)							1 (2.6%)	1					1 (0.6%)	1	
Related to THS 2.2/CC							0						0		
Hepatobiliary disorders	1 (1.3%)		1	0			0			0			1 (0.6%)	1	
(Hyperbilirubinaemia)	1 (1.3%)		1										1 (0.6%)	1	
Related to THS 2.2/CC	0												0		

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Exposed Not Randomized refers to all subjects exposed to THS 2.2 but not randomized. Overall Safety refers to all subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.6.1.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_adv10.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.3 Summary of Adverse Events by System Organ Class, Preferred Term and Relationship to Study Product Exposure and Expectedness - Safety Population**

	THS 2.2 (N=80)			CC (N=41)			SA (N=39)			Exposed Not Randomized (N=9)			Overall Safety (N=169)		
	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events
Psychiatric disorders	0			0			1 (2.6%)	1		0			1 (0.6%)	1	
(Nicotine dependence)							1 (2.6%)	1					1 (0.6%)	1	
Related to THS 2.2/CC							0						0		
Skin and subcutaneous tissue disorders	1 (1.3%)	1		0			0			0			1 (0.6%)	1	
(Hyperhidrosis)	1 (1.3%)	1											1 (0.6%)	1	
Related to THS 2.2/CC	1 (1.3%)	1											1 (0.6%)	1	
Expected	0												0		

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Exposed Not Randomized refers to all subjects exposed to THS 2.2 but not randomized. Overall Safety refers to all subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.6.1.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_adv10.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.4 Summary of Adverse Events Leading to Product Discontinuation or Reduction by System Organ Class and Preferred Term - Safety Population**

System Organ Class Preferred Term	THS 2.2 (N=80)			CC (N=41)			SA (N=39)			Exposed Not Randomized (N=9)			Overall Safety (N=169)		
	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events
Any adverse events leading to product discontinuation or reduction	0			1 (2.4%)	1		0			0			1 (0.6%)	1	
Investigations	0			1 (2.4%)	1		0						1 (0.6%)	1	
Carboxyhaemoglobin increased				1 (2.4%)	1								1 (0.6%)	1	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Exposed Not Randomized refers to all subjects exposed to THS 2.2 but not randomized. Overall Safety refers to all subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.6.1.3

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_adv11.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.5 Summary of Adverse Events Leading to Study Discontinuation by System Organ Class and Preferred Term - Safety Population**

System Organ Class Preferred Term	THS 2.2 (N=80)			CC (N=41)			SA (N=39)			Exposed Not Randomized (N=9)			Overall Safety (N=169)		
	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events
Any adverse events leading to study discontinuation	0			0			0			8 (88.9%)	8		8 (4.7%)	8	
Investigations										3 (33.3%)	3		3 (1.8%)	3	
Spirometry abnormal										3 (33.3%)	3		3 (1.8%)	3	
Blood and lymphatic system disorders										2 (22.2%)	2		2 (1.2%)	2	
Leukocytosis										2 (22.2%)	2		2 (1.2%)	2	
Vascular disorders										2 (22.2%)	2		2 (1.2%)	2	
Hypertension										2 (22.2%)	2		2 (1.2%)	2	
Metabolism and nutrition disorders										1 (11.1%)	1		1 (0.6%)	1	
Hyperglycaemia										1 (11.1%)	1		1 (0.6%)	1	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Exposed Not Randomized refers to all subjects exposed to THS 2.2 but not randomized. Overall Safety refers to all subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.6.1.3

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_adv5.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.6 Summary of Adverse Events Related to Study Procedure by System Organ Class and Preferred Term - Safety Population**

System Organ Class Preferred Term	THS 2.2 (N=80)			CC (N=41)			SA (N=39)			Exposed Not Randomized (N=9)			Overall Safety (N=169)		
	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events
Any adverse events related to study procedure	11	(13.8%)	11	6	(14.6%)	8	1	(2.6%)	1	2	(22.2%)	2	20	(11.8%)	22
Investigations	2	(2.5%)	2	3	(7.3%)	5	0			2	(22.2%)	2	7	(4.1%)	9
Spirometry abnormal	2	(2.5%)	2	1	(2.4%)	1				2	(22.2%)	2	5	(3.0%)	5
Carboxyhaemoglobin increased	0			2	(4.9%)	4				0			2	(1.2%)	4
Nervous system disorders	4	(5.0%)	4	3	(7.3%)	3	0			0			7	(4.1%)	7
Syncope	4	(5.0%)	4	3	(7.3%)	3							7	(4.1%)	7
General disorders and administration site conditions	2	(2.5%)	2	0			0			0			2	(1.2%)	2
Vessel puncture site reaction	2	(2.5%)	2										2	(1.2%)	2

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Exposed Not Randomized refers to all subjects exposed to THS 2.2 but not randomized. Overall Safety refers to all subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.6.1.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_adv6.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.6 Summary of Adverse Events Related to Study Procedure by System Organ Class and Preferred Term - Safety Population**

System Organ Class Preferred Term	THS 2.2 (N=80)			CC (N=41)			SA (N=39)			Exposed Not Randomized (N=9)			Overall Safety (N=169)		
	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events
Ear and labyrinth disorders	1	(1.3%)	1	0			0			0			1	(0.6%)	1
Vertigo	1	(1.3%)	1										1	(0.6%)	1
Musculoskeletal and connective tissue disorders	1	(1.3%)	1	0			0			0			1	(0.6%)	1
Musculoskeletal chest pain	1	(1.3%)	1										1	(0.6%)	1
Psychiatric disorders	0			0			1	(2.6%)	1	0			1	(0.6%)	1
Nicotine dependence							1	(2.6%)	1				1	(0.6%)	1
Respiratory, thoracic and mediastinal disorders	1	(1.3%)	1	0			0			0			1	(0.6%)	1
Oropharyngeal pain	1	(1.3%)	1										1	(0.6%)	1

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Exposed Not Randomized refers to all subjects exposed to THS 2.2 but not randomized. Overall Safety refers to all subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.6.1.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_adv6.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.7 Summary of Adverse Events by System Organ Class, Preferred Term and Severity - Safety Population**

System Organ Class Preferred Term	THS 2.2 (N=80)			CC (N=41)			SA (N=39)			Exposed Not Randomized (N=9)			Overall Safety (N=169)		
	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events
Adverse events	50	(62.5%)	98	29	(70.7%)	63	24	(61.5%)	49	9	(100 %)	17	112	(66.3%)	227
Mild	28	(35.0%)	71	13	(31.7%)	42	11	(28.2%)	29	3	(33.3%)	10	55	(32.5%)	152
Moderate	22	(27.5%)	27	15	(36.6%)	20	13	(33.3%)	20	6	(66.7%)	7	56	(33.1%)	74
Severe	0			1	(2.4%)	1	0			0			1	(0.6%)	1
Nervous system disorders	27	(33.8%)	36	19	(46.3%)	23	13	(33.3%)	15	3	(33.3%)	3	62	(36.7%)	77
Headache	24	(30.0%)	30	16	(39.0%)	19	13	(33.3%)	15	3	(33.3%)	3	56	(33.1%)	67
Mild	16	(20.0%)	21	10	(24.4%)	12	7	(17.9%)	9	0			33	(19.5%)	42
Moderate	8	(10.0%)	9	6	(14.6%)	7	6	(15.4%)	6	3	(33.3%)	3	23	(13.6%)	25
Severe	0			0			0			0			0		

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Exposed Not Randomized refers to all subjects exposed to THS 2.2 but not randomized. Overall Safety refers to all subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.6.1.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_adv7.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.7 Summary of Adverse Events by System Organ Class, Preferred Term and Severity - Safety Population**

System Organ Class Preferred Term	THS 2.2 (N=80)			CC (N=41)			SA (N=39)			Exposed Not Randomized (N=9)			Overall Safety (N=169)		
	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events
Syncope	6	(7.5%)	6	4	(9.8%)	4	0			0			10	(5.9%)	10
Mild	5	(6.3%)	5	2	(4.9%)	2							7	(4.1%)	7
Moderate	1	(1.3%)	1	2	(4.9%)	2							3	(1.8%)	3
Severe	0			0									0		
Investigations	7	(8.8%)	7	6	(14.6%)	8	3	(7.7%)	3	3	(33.3%)	4	19	(11.2%)	22
Spirometry abnormal	4	(5.0%)	4	3	(7.3%)	3	2	(5.1%)	2	3	(33.3%)	3	12	(7.1%)	12
Mild	4	(5.0%)	4	3	(7.3%)	3	2	(5.1%)	2	3	(33.3%)	3	12	(7.1%)	12
Moderate	0			0			0			0			0		
Severe	0			0			0			0			0		

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Exposed Not Randomized refers to all subjects exposed to THS 2.2 but not randomized. Overall Safety refers to all subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.6.1.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_adv7.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.7 Summary of Adverse Events by System Organ Class, Preferred Term and Severity - Safety Population**

System Organ Class Preferred Term	THS 2.2 (N=80)			CC (N=41)			SA (N=39)			Exposed Not Randomized (N=9)			Overall Safety (N=169)		
	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events
Lymphocyte count increased	1	(1.3%)	1	0			1	(2.6%)	1	1	(11.1%)	1	3	(1.8%)	3
Mild	0						0			0			0		
Moderate	1	(1.3%)	1				1	(2.6%)	1	1	(11.1%)	1	3	(1.8%)	3
Severe	0						0			0			0		
Carboxyhaemoglobin increased	0			2	(4.9%)	4	0			0			2	(1.2%)	4
Mild				2	(4.9%)	4							2	(1.2%)	4
Moderate				0									0		
Severe				0									0		
Blood bilirubin increased	1	(1.3%)	1	0			0			0			1	(0.6%)	1
Mild	0												0		
Moderate	1	(1.3%)	1										1	(0.6%)	1
Severe	0												0		

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Exposed Not Randomized refers to all subjects exposed to THS 2.2 but not randomized. Overall Safety refers to all subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.6.1.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_adv7.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.7 Summary of Adverse Events by System Organ Class, Preferred Term and Severity - Safety Population**

System Organ Class Preferred Term	THS 2.2 (N=80)			CC (N=41)			SA (N=39)			Exposed Not Randomized (N=9)			Overall Safety (N=169)		
	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events
Eosinophil count increased	1	(1.3%)	1	0			0			0			1	(0.6%)	1
Mild	1	(1.3%)	1										1	(0.6%)	1
Moderate	0												0		
Severe	0												0		
Neutrophil count decreased	0			1	(2.4%)	1	0			0			1	(0.6%)	1
Mild				0									0		
Moderate				1	(2.4%)	1							1	(0.6%)	1
Severe				0									0		
Gastrointestinal disorders	7	(8.8%)	7	5	(12.2%)	6	3	(7.7%)	5	2	(22.2%)	2	17	(10.1%)	20

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Exposed Not Randomized refers to all subjects exposed to THS 2.2 but not randomized. Overall Safety refers to all subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.6.1.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_adv7.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.7 Summary of Adverse Events by System Organ Class, Preferred Term and Severity - Safety Population**

System Organ Class Preferred Term	THS 2.2 (N=80)			CC (N=41)			SA (N=39)			Exposed Not Randomized (N=9)			Overall Safety (N=169)		
	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events
Constipation	3	(3.8%)	3	2	(4.9%)	2	0			1	(11.1%)	1	6	(3.6%)	6
Mild	3	(3.8%)	3	1	(2.4%)	1				0			4	(2.4%)	4
Moderate	0			1	(2.4%)	1				1	(11.1%)	1	2	(1.2%)	2
Severe	0			0						0			0		
Toothache	1	(1.3%)	1	1	(2.4%)	1	0			1	(11.1%)	1	3	(1.8%)	3
Mild	0			0						0			0		
Moderate	1	(1.3%)	1	1	(2.4%)	1				1	(11.1%)	1	3	(1.8%)	3
Severe	0			0						0			0		
Gastrooesophageal reflux disease	0			2	(4.9%)	3	0			0			2	(1.2%)	3
Mild				2	(4.9%)	3							2	(1.2%)	3
Moderate				0									0		
Severe				0									0		

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Exposed Not Randomized refers to all subjects exposed to THS 2.2 but not randomized. Overall Safety refers to all subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.6.1.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_adv7.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.7 Summary of Adverse Events by System Organ Class, Preferred Term and Severity - Safety Population**

System Organ Class Preferred Term	THS 2.2 (N=80)			CC (N=41)			SA (N=39)			Exposed Not Randomized (N=9)			Overall Safety (N=169)		
	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events
Abdominal distension	0			0			2 (5.1%)	2		0			2 (1.2%)	2	
Mild							1 (2.6%)	1					1 (0.6%)	1	
Moderate							1 (2.6%)	1					1 (0.6%)	1	
Severe							0						0		
Mouth ulceration	2 (2.5%)		2	0			0			0			2 (1.2%)	2	
Mild	1 (1.3%)		1										1 (0.6%)	1	
Moderate	1 (1.3%)		1										1 (0.6%)	1	
Severe	0												0		
Abdominal pain upper	0			0			1 (2.6%)	1		0			1 (0.6%)	1	
Mild							0						0		
Moderate							1 (2.6%)	1					1 (0.6%)	1	
Severe							0						0		

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Exposed Not Randomized refers to all subjects exposed to THS 2.2 but not randomized. Overall Safety refers to all subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.6.1.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_adv7.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.7 Summary of Adverse Events by System Organ Class, Preferred Term and Severity - Safety Population**

System Organ Class Preferred Term	THS 2.2 (N=80)			CC (N=41)			SA (N=39)			Exposed Not Randomized (N=9)			Overall Safety (N=169)		
	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events
Nausea	0			0			1 (2.6%)	1		0			1 (0.6%)	1	
Mild							0						0		
Moderate							1 (2.6%)	1					1 (0.6%)	1	
Severe							0						0		
Oral pain	1 (1.3%)	1		0			0			0			1 (0.6%)	1	
Mild	1 (1.3%)	1											1 (0.6%)	1	
Moderate	0												0		
Severe	0												0		
Vomiting	0			0			1 (2.6%)	1		0			1 (0.6%)	1	
Mild							0						0		
Moderate							1 (2.6%)	1					1 (0.6%)	1	
Severe							0						0		

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Exposed Not Randomized refers to all subjects exposed to THS 2.2 but not randomized. Overall Safety refers to all subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.6.1.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_adv7.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.7 Summary of Adverse Events by System Organ Class, Preferred Term and Severity - Safety Population**

System Organ Class Preferred Term	THS 2.2 (N=80)			CC (N=41)			SA (N=39)			Exposed Not Randomized (N=9)			Overall Safety (N=169)		
	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events
Respiratory, thoracic and mediastinal disorders	10	(12.5%)	10	4	(9.8%)	4	0			1	(11.1%)	1	15	(8.9%)	15
Oropharyngeal pain	7	(8.8%)	7	3	(7.3%)	3				0			10	(5.9%)	10
Mild	5	(6.3%)	5	2	(4.9%)	2							7	(4.1%)	7
Moderate	2	(2.5%)	2	1	(2.4%)	1							3	(1.8%)	3
Severe	0			0									0		
Cough	3	(3.8%)	3	0						1	(11.1%)	1	4	(2.4%)	4
Mild	2	(2.5%)	2							1	(11.1%)	1	3	(1.8%)	3
Moderate	1	(1.3%)	1							0			1	(0.6%)	1
Severe	0									0			0		

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Exposed Not Randomized refers to all subjects exposed to THS 2.2 but not randomized. Overall Safety refers to all subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.6.1.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_adv7.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.7 Summary of Adverse Events by System Organ Class, Preferred Term and Severity - Safety Population**

System Organ Class Preferred Term	THS 2.2 (N=80)			CC (N=41)			SA (N=39)			Exposed Not Randomized (N=9)			Overall Safety (N=169)		
	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events
Epistaxis	0			1 (2.4%)		1				0			1 (0.6%)		1
Mild				1 (2.4%)		1							1 (0.6%)		1
Moderate				0									0		
Severe				0									0		
Renal and urinary disorders	7 (8.8%)		9	4 (9.8%)		6	2 (5.1%)		2	0			13 (7.7%)		17
Polyuria	6 (7.5%)		8	4 (9.8%)		6	2 (5.1%)		2				12 (7.1%)		16
Mild	6 (7.5%)		8	4 (9.8%)		6	2 (5.1%)		2				12 (7.1%)		16
Moderate	0			0			0						0		
Severe	0			0			0						0		

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Exposed Not Randomized refers to all subjects exposed to THS 2.2 but not randomized. Overall Safety refers to all subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.6.1.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_adv7.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.7 Summary of Adverse Events by System Organ Class, Preferred Term and Severity - Safety Population**

System Organ Class Preferred Term	THS 2.2 (N=80)			CC (N=41)			SA (N=39)			Exposed Not Randomized (N=9)			Overall Safety (N=169)		
	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events
Leukocyturia	1	(1.3%)	1	0			0						1	(0.6%)	1
Mild	1	(1.3%)	1										1	(0.6%)	1
Moderate	0												0		
Severe	0												0		
Metabolism and nutrition disorders	4	(5.0%)	4	3	(7.3%)	3	4	(10.3%)	4	2	(22.2%)	2	13	(7.7%)	13
Hypertriglyceridaemia	3	(3.8%)	3	2	(4.9%)	2	2	(5.1%)	2	0			7	(4.1%)	7
Mild	0			0			0						0		
Moderate	3	(3.8%)	3	1	(2.4%)	1	2	(5.1%)	2				6	(3.6%)	6
Severe	0			1	(2.4%)	1	0						1	(0.6%)	1

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Exposed Not Randomized refers to all subjects exposed to THS 2.2 but not randomized. Overall Safety refers to all subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.6.1.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_adv7.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.7 Summary of Adverse Events by System Organ Class, Preferred Term and Severity - Safety Population**

System Organ Class Preferred Term	THS 2.2 (N=80)			CC (N=41)			SA (N=39)			Exposed Not Randomized (N=9)			Overall Safety (N=169)		
	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events
Hypercholesterolaemia	0			1 (2.4%)	1		1 (2.6%)	1		0			2 (1.2%)	2	
Mild				0			0						0		
Moderate				1 (2.4%)	1		1 (2.6%)	1					2 (1.2%)	2	
Severe				0			0						0		
Hyperglycaemia	0			0			1 (2.6%)	1		1 (11.1%)	1		2 (1.2%)	2	
Mild							1 (2.6%)	1		1 (11.1%)	1		2 (1.2%)	2	
Moderate							0			0			0		
Severe							0			0			0		
Hyperkalaemia	1 (1.3%)	1		0			0			1 (11.1%)	1		2 (1.2%)	2	
Mild	0									0			0		
Moderate	1 (1.3%)	1								1 (11.1%)	1		2 (1.2%)	2	
Severe	0									0			0		

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Exposed Not Randomized refers to all subjects exposed to THS 2.2 but not randomized. Overall Safety refers to all subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.6.1.1

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.7 Summary of Adverse Events by System Organ Class, Preferred Term and Severity - Safety Population**

System Organ Class Preferred Term	THS 2.2 (N=80)			CC (N=41)			SA (N=39)			Exposed Not Randomized (N=9)			Overall Safety (N=169)		
	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events
Musculoskeletal and connective tissue disorders	6	(7.5%)	6	2	(4.9%)	3	4	(10.3%)	4	0			12	(7.1%)	13
Back pain	3	(3.8%)	3	2	(4.9%)	3	3	(7.7%)	3				8	(4.7%)	9
Mild	1	(1.3%)	1	1	(2.4%)	2	3	(7.7%)	3				5	(3.0%)	6
Moderate	2	(2.5%)	2	1	(2.4%)	1	0						3	(1.8%)	3
Severe	0			0			0						0		
Arthralgia	1	(1.3%)	1	0			0						1	(0.6%)	1
Mild	1	(1.3%)	1										1	(0.6%)	1
Moderate	0												0		
Severe	0												0		

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Exposed Not Randomized refers to all subjects exposed to THS 2.2 but not randomized. Overall Safety refers to all subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.6.1.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_adv7.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.7 Summary of Adverse Events by System Organ Class, Preferred Term and Severity - Safety Population**

System Organ Class Preferred Term	THS 2.2 (N=80)			CC (N=41)			SA (N=39)			Exposed Not Randomized (N=9)			Overall Safety (N=169)		
	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events
Musculoskeletal chest pain	1	(1.3%)	1	0			0						1	(0.6%)	1
Mild	1	(1.3%)	1										1	(0.6%)	1
Moderate	0												0		
Severe	0												0		
Myalgia	0			0			1	(2.6%)	1				1	(0.6%)	1
Mild							1	(2.6%)	1				1	(0.6%)	1
Moderate							0						0		
Severe							0						0		
Pain in extremity	1	(1.3%)	1	0			0						1	(0.6%)	1
Mild	0												0		
Moderate	1	(1.3%)	1										1	(0.6%)	1
Severe	0												0		

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Exposed Not Randomized refers to all subjects exposed to THS 2.2 but not randomized. Overall Safety refers to all subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.6.1.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_adv7.sas

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**Table 15.2.6.7 Summary of Adverse Events by System Organ Class, Preferred Term and Severity - Safety Population**

System Organ Class Preferred Term	THS 2.2 (N=80)			CC (N=41)			SA (N=39)			Exposed Not Randomized (N=9)			Overall Safety (N=169)		
	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events
Vascular disorders	3	(3.8%)	4	2	(4.9%)	2	3	(7.7%)	7	2	(22.2%)	2	10	(5.9%)	15
Hypertension	1	(1.3%)	2	1	(2.4%)	1	2	(5.1%)	5	2	(22.2%)	2	6	(3.6%)	10
Mild	1	(1.3%)	2	1	(2.4%)	1	2	(5.1%)	5	2	(22.2%)	2	6	(3.6%)	10
Moderate	0			0			0			0			0		
Severe	0			0			0			0			0		
Hypotension	2	(2.5%)	2	1	(2.4%)	1	1	(2.6%)	2	0			4	(2.4%)	5
Mild	2	(2.5%)	2	1	(2.4%)	1	1	(2.6%)	2				4	(2.4%)	5
Moderate	0			0			0						0		
Severe	0			0			0						0		

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Exposed Not Randomized refers to all subjects exposed to THS 2.2 but not randomized. Overall Safety refers to all subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.6.1.1

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.7 Summary of Adverse Events by System Organ Class, Preferred Term and Severity - Safety Population**

System Organ Class Preferred Term	THS 2.2 (N=80)			CC (N=41)			SA (N=39)			Exposed Not Randomized (N=9)			Overall Safety (N=169)		
	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events
Infections and infestations	4	(5.0%)	4	2	(4.9%)	3	1	(2.6%)	1	0			7	(4.1%)	8
Nasopharyngitis	3	(3.8%)	3	1	(2.4%)	2	1	(2.6%)	1				5	(3.0%)	6
Mild	2	(2.5%)	2	0		1	1	(2.6%)	1				3	(1.8%)	4
Moderate	1	(1.3%)	1	1	(2.4%)	1	0						2	(1.2%)	2
Severe	0			0			0						0		
Herpes virus infection	0			1	(2.4%)	1	0						1	(0.6%)	1
Mild				0									0		
Moderate				1	(2.4%)	1							1	(0.6%)	1
Severe				0									0		

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Exposed Not Randomized refers to all subjects exposed to THS 2.2 but not randomized. Overall Safety refers to all subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.6.1.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_adv7.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.7 Summary of Adverse Events by System Organ Class, Preferred Term and Severity - Safety Population**

System Organ Class Preferred Term	THS 2.2 (N=80)			CC (N=41)			SA (N=39)			Exposed Not Randomized (N=9)			Overall Safety (N=169)		
	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events
Urinary tract infection	1	(1.3%)	1	0			0						1	(0.6%)	1
Mild	1	(1.3%)	1										1	(0.6%)	1
Moderate	0												0		
Severe	0												0		
General disorders and administration site conditions	2	(2.5%)	2	1	(2.4%)	1	3	(7.7%)	3	0			6	(3.6%)	6
Influenza like illness	0			1	(2.4%)	1	3	(7.7%)	3				4	(2.4%)	4
Mild				0			0						0		
Moderate				1	(2.4%)	1	3	(7.7%)	3				4	(2.4%)	4
Severe				0			0						0		

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Exposed Not Randomized refers to all subjects exposed to THS 2.2 but not randomized. Overall Safety refers to all subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.6.1.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_adv7.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.7 Summary of Adverse Events by System Organ Class, Preferred Term and Severity - Safety Population**

System Organ Class Preferred Term	THS 2.2 (N=80)			CC (N=41)			SA (N=39)			Exposed Not Randomized (N=9)			Overall Safety (N=169)		
	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events
Vessel puncture site reaction	2	(2.5%)	2	0			0						2	(1.2%)	2
Mild	2	(2.5%)	2										2	(1.2%)	2
Moderate	0												0		
Severe	0												0		
Ear and labyrinth disorders	3	(3.8%)	3	0			2	(5.1%)	2	0			5	(3.0%)	5
Vertigo	3	(3.8%)	3				2	(5.1%)	2				5	(3.0%)	5
Mild	3	(3.8%)	3				0						3	(1.8%)	3
Moderate	0						2	(5.1%)	2				2	(1.2%)	2
Severe	0						0						0		

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Exposed Not Randomized refers to all subjects exposed to THS 2.2 but not randomized. Overall Safety refers to all subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.6.1.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_adv7.sas

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**Table 15.2.6.7 Summary of Adverse Events by System Organ Class, Preferred Term and Severity - Safety Population**

System Organ Class Preferred Term	THS 2.2 (N=80)			CC (N=41)			SA (N=39)			Exposed Not Randomized (N=9)			Overall Safety (N=169)		
	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events
Reproductive system and breast disorders	3	(3.8%)	3	1	(2.4%)	1	1	(2.6%)	1	0			5	(3.0%)	5
Dysmenorrhoea	3	(3.8%)	3	1	(2.4%)	1	1	(2.6%)	1				5	(3.0%)	5
Mild	2	(2.5%)	2	0		0	0		0				2	(1.2%)	2
Moderate	1	(1.3%)	1	1	(2.4%)	1	1	(2.6%)	1				3	(1.8%)	3
Severe	0		0	0		0	0		0				0		0
Cardiac disorders	1	(1.3%)	1	2	(4.9%)	3	0		0	1	(11.1%)	1	4	(2.4%)	5
Tachycardia	0		0	2	(4.9%)	3				0			2	(1.2%)	3
Mild				2	(4.9%)	3							2	(1.2%)	3
Moderate				0		0							0		0
Severe				0		0							0		0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Exposed Not Randomized refers to all subjects exposed to THS 2.2 but not randomized. Overall Safety refers to all subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.6.1.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_adv7.sas

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**Table 15.2.6.7 Summary of Adverse Events by System Organ Class, Preferred Term and Severity - Safety Population**

System Organ Class Preferred Term	THS 2.2 (N=80)			CC (N=41)			SA (N=39)			Exposed Not Randomized (N=9)			Overall Safety (N=169)		
	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events
Myocardial ischaemia	1	(1.3%)	1	0						0			1	(0.6%)	1
Mild	1	(1.3%)	1										1	(0.6%)	1
Moderate	0												0		
Severe	0												0		
Ventricular extrasystoles	0			0						1	(11.1%)	1	1	(0.6%)	1
Mild										1	(11.1%)	1	1	(0.6%)	1
Moderate										0			0		
Severe										0			0		
Blood and lymphatic system disorders	0			0			0			2	(22.2%)	2	2	(1.2%)	2

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Exposed Not Randomized refers to all subjects exposed to THS 2.2 but not randomized. Overall Safety refers to all subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.6.1.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_adv7.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.7 Summary of Adverse Events by System Organ Class, Preferred Term and Severity - Safety Population**

System Organ Class Preferred Term	THS 2.2 (N=80)			CC (N=41)			SA (N=39)			Exposed Not Randomized (N=9)			Overall Safety (N=169)		
	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events
Leukocytosis										2 (22.2%)	2		2 (1.2%)	2	
Mild										2 (22.2%)	2		2 (1.2%)	2	
Moderate										0			0		
Severe										0			0		
Eye disorders	0			0			1 (2.6%)	1		0			1 (0.6%)	1	
Conjunctivitis							1 (2.6%)	1					1 (0.6%)	1	
Mild							1 (2.6%)	1					1 (0.6%)	1	
Moderate							0						0		
Severe							0						0		

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Exposed Not Randomized refers to all subjects exposed to THS 2.2 but not randomized. Overall Safety refers to all subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.6.1.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_adv7.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.7 Summary of Adverse Events by System Organ Class, Preferred Term and Severity - Safety Population**

System Organ Class Preferred Term	THS 2.2 (N=80)			CC (N=41)			SA (N=39)			Exposed Not Randomized (N=9)			Overall Safety (N=169)		
	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events
Hepatobiliary disorders	1	(1.3%)	1	0			0			0			1	(0.6%)	1
Hyperbilirubinaemia	1	(1.3%)	1										1	(0.6%)	1
Mild	0												0		
Moderate	1	(1.3%)	1										1	(0.6%)	1
Severe	0												0		
Psychiatric disorders	0			0			1	(2.6%)	1	0			1	(0.6%)	1
Nicotine dependence							1	(2.6%)	1				1	(0.6%)	1
Mild							1	(2.6%)	1				1	(0.6%)	1
Moderate							0						0		
Severe							0						0		

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Exposed Not Randomized refers to all subjects exposed to THS 2.2 but not randomized. Overall Safety refers to all subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.6.1.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_adv7.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.7 Summary of Adverse Events by System Organ Class, Preferred Term and Severity - Safety Population**

System Organ Class Preferred Term	THS 2.2 (N=80)			CC (N=41)			SA (N=39)			Exposed Not Randomized (N=9)			Overall Safety (N=169)		
	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events
Skin and subcutaneous tissue disorders	1	(1.3%)	1	0			0			0			1	(0.6%)	1
Hyperhidrosis	1	(1.3%)	1										1	(0.6%)	1
Mild	1	(1.3%)	1										1	(0.6%)	1
Moderate	0												0		
Severe	0												0		

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Exposed Not Randomized refers to all subjects exposed to THS 2.2 but not randomized. Overall Safety refers to all subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.6.1.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_adv7.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.8 Summary of Serious Adverse Events - Safety Population**

THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Exposed Not Randomized (N=9)	Overall Safety (N=169)
n (%) Events	n (%) Events	n (%) Events	n (%) Events	n (%) Events

No serious adverse events were reported

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Exposed Not Randomized refers to all subjects exposed to THS 2.2 but not randomized. Overall Safety refers to all subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.6.1.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_adv8.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.9 Summary of Serious Adverse Events by System Organ Class and Preferred Term - Safety Population**

System Organ Class Preferred Term	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Exposed Not Randomized (N=9)	Overall Safety (N=169)
	n (%) Events	n (%) Events	n (%) Events	n (%) Events	n (%) Events

No serious adverse events were reported

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Exposed Not Randomized refers to all subjects exposed to THS 2.2 but not randomized. Overall Safety refers to all subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.6.1.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_adv9.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.10 Summary of THS 2.2 Device Events and Malfunctions - Safety Population**

	THS 2.2 (N=80)			CC (N=41)			SA (N=39)			Enrolled Not Randomized (N=9)			Overall Safety (N=169)		
	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events
Device events	12	(15.0%)	19	0		0			0				12	(7.1%)	19
AE relationship															
Related	0												0		
Not related	12	(15.0%)	19										12	(7.1%)	19
Major	12	(15.0%)	19										12	(7.1%)	19
Charging issue	8	(10.0%)	9										8	(4.7%)	9
Device inoperable	3	(3.8%)	3										3	(1.8%)	3
Device stops intermittently	2	(2.5%)	3										2	(1.2%)	3
Device operates differently than expected	2	(2.5%)	2										2	(1.2%)	2
Device difficult to setup or prepare	1	(1.3%)	1										1	(0.6%)	1
Output issue	1	(1.3%)	1										1	(0.6%)	1
Minor	0												0		

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System

Note: Enrolled Not Randomized refers to all subjects who were enrolled but not randomized. Overall Safety refers to all subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.6.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_device.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.11.1 Summary of Prior Medication by Anatomical Therapeutic Classes (ATC) 1 and 2 - Safety Population**

ATC1 ATC2	THS 2.2 (N=80)			CC (N=41)			SA (N=39)			Exposed Not Randomized (N=9)			Overall Safety (N=169)		
	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events
Any medication	7	(8.8%)	7	1	(2.4%)	1	4	(10.3%)	4	1	(11.1%)	1	13	(7.7%)	13
Musculo-skeletal system	3	(3.8%)	3	1	(2.4%)	1	1	(2.6%)	1	0			5	(3.0%)	5
Antiinflammatory and antirheumatic products	3	(3.8%)	3	1	(2.4%)	1	1	(2.6%)	1				5	(3.0%)	5
Nervous system	1	(1.3%)	1	0			2	(5.1%)	2	0			3	(1.8%)	3
Analgesics	1	(1.3%)	1				0						1	(0.6%)	1
Anesthetics	0						1	(2.6%)	1				1	(0.6%)	1
Psychoanaleptics	0						1	(2.6%)	1				1	(0.6%)	1
Respiratory system	2	(2.5%)	2	0			0			0			2	(1.2%)	2
Antihistamines for systemic use	1	(1.3%)	1										1	(0.6%)	1
Cough and cold preparations	1	(1.3%)	1										1	(0.6%)	1

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Exposed Not Randomized refers to all subjects exposed to THS 2.2 but not randomized. Overall Safety refers to all subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.6.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_priatc.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.11.1 Summary of Prior Medication by Anatomical Therapeutic Classes (ATC) 1 and 2 - Safety Population**

ATC1 ATC2	THS 2.2 (N=80)			CC (N=41)			SA (N=39)			Exposed Not Randomized (N=9)			Overall Safety (N=169)		
	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events
Antiinfectives for systemic use	0			0			0			1 (11.1%)	1		1 (0.6%)	1	
Antibacterials for systemic use										1 (11.1%)	1		1 (0.6%)	1	
Genito urinary system and sex hormones	0			0			1 (2.6%)	1		0			1 (0.6%)	1	
Sex hormones and modulators of the genital system							1 (2.6%)	1					1 (0.6%)	1	
Various	1 (1.3%)		1	0			0			0			1 (0.6%)	1	
Unspecified herbal and traditional medicine	1 (1.3%)		1										1 (0.6%)	1	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Exposed Not Randomized refers to all subjects exposed to THS 2.2 but not randomized. Overall Safety refers to all subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.6.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_priatc.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.11.2 Summary of Prior Medication by Preferred Drug Name - Safety Population**

Preferred Term	THS 2.2 (N=80)			CC (N=41)			SA (N=39)			Exposed Not Randomized (N=9)			Overall Safety (N=169)		
	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events
Any medication	7	(8.8%)	7	1	(2.4%)	1	4	(10.3%)	4	1	(11.1%)	1	13	(7.7%)	13
Ibuprofen	3	(3.8%)	3	1	(2.4%)	1	1	(2.6%)	1	0			5	(3.0%)	5
Amoxicillin trihydrate	0			0			0			1	(11.1%)	1	1	(0.6%)	1
Cetirizine hydrochloride	1	(1.3%)	1	0			0			0			1	(0.6%)	1
Cimicifuga racemosa extract	1	(1.3%)	1	0			0			0			1	(0.6%)	1
Lafamme	0			0			1	(2.6%)	1	0			1	(0.6%)	1
Opipramol hydrochloride	0			0			1	(2.6%)	1	0			1	(0.6%)	1
Other local anesthetics	0			0			1	(2.6%)	1	0			1	(0.6%)	1
Paracetamol	1	(1.3%)	1	0			0			0			1	(0.6%)	1
Prolex dh /01539501/	1	(1.3%)	1	0			0			0			1	(0.6%)	1

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Exposed Not Randomized refers to all subjects exposed to THS 2.2 but not randomized. Overall Safety refers to all subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.6.3.1

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_primed.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.12.1 Summary of Concomitant Medication by Anatomical Therapeutic Classes (ATC) 1 and 2 - Safety Population**

ATC1 ATC2	THS 2.2 (N=80)			CC (N=41)			SA (N=39)			Exposed Not Randomized (N=9)			Overall Safety (N=169)		
	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events
Any medication	11	(13.8%)	16	9	(22.0%)	12	10	(25.6%)	17	3	(33.3%)	5	33	(19.5%)	50
Nervous system	7	(8.8%)	10	7	(17.1%)	8	8	(20.5%)	13	3	(33.3%)	3	25	(14.8%)	34
Analgesics	7	(8.8%)	10	7	(17.1%)	8	8	(20.5%)	13	3	(33.3%)	3	25	(14.8%)	34
Alimentary tract and metabolism	2	(2.5%)	2	1	(2.4%)	1	1	(2.6%)	1	1	(11.1%)	1	5	(3.0%)	5
Drugs for constipation	1	(1.3%)	1	1	(2.4%)	1	0			1	(11.1%)	1	3	(1.8%)	3
Drugs for functional gastrointestinal disorders	1	(1.3%)	1	0			1	(2.6%)	1	0			2	(1.2%)	2
Respiratory system	1	(1.3%)	1	1	(2.4%)	1	1	(2.6%)	2	0			3	(1.8%)	4
Cough and cold preparations	1	(1.3%)	1	1	(2.4%)	1	0						2	(1.2%)	2
Nasal preparations	0			0			1	(2.6%)	1				1	(0.6%)	1
Throat preparations	0			0			1	(2.6%)	1				1	(0.6%)	1

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Exposed Not Randomized refers to all subjects exposed to THS 2.2 but not randomized. Overall Safety refers to all subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.6.3.2

Path: /cvm/projects/prj/development/000000106324/dev/tables/t_conatc.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.12.1 Summary of Concomitant Medication by Anatomical Therapeutic Classes (ATC) 1 and 2 - Safety Population**

ATC1 ATC2	THS 2.2 (N=80)			CC (N=41)			SA (N=39)			Exposed Not Randomized (N=9)			Overall Safety (N=169)		
	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events
Cardiovascular system	0			1 (2.4%)	1		1 (2.6%)	1		1 (11.1%)	1		3 (1.8%)	3	
Vasoprotectives				1 (2.4%)	1		1 (2.6%)	1		0			2 (1.2%)	2	
Calcium channel blockers				0			0			1 (11.1%)	1		1 (0.6%)	1	
Musculo-skeletal system	1 (1.3%)	1		1 (2.4%)	1		0			0			2 (1.2%)	2	
Antiinflammatory and antirheumatic products	1 (1.3%)	1		1 (2.4%)	1								2 (1.2%)	2	
Dermatologicals	1 (1.3%)	1		0			0			0			1 (0.6%)	1	
Antiseptics and disinfectants	1 (1.3%)	1											1 (0.6%)	1	
Various	1 (1.3%)	1		0			0			0			1 (0.6%)	1	
Unspecified herbal and traditional medicine	1 (1.3%)	1											1 (0.6%)	1	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Exposed Not Randomized refers to all subjects exposed to THS 2.2 but not randomized. Overall Safety refers to all subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.6.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_conatc.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.12.2 Summary of Concomitant Medication by Preferred Drug Name - Safety Population**

Preferred Term	THS 2.2 (N=80)			CC (N=41)			SA (N=39)			Exposed Not Randomized (N=9)			Overall Safety (N=169)		
	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events
Any medication	11	(13.8%)	16	9	(22.0%)	12	10	(25.6%)	17	3	(33.3%)	5	33	(19.5%)	50
Paracetamol	7	(8.8%)	10	7	(17.1%)	8	8	(20.5%)	12	3	(33.3%)	3	25	(14.8%)	33
Glyceol /00744501/	1	(1.3%)	1	1	(2.4%)	1	0			1	(11.1%)	1	3	(1.8%)	3
Ascorutin	0			1	(2.4%)	1	1	(2.6%)	1	0			2	(1.2%)	2
Benadryl cold and flu	1	(1.3%)	1	1	(2.4%)	1	0			0			2	(1.2%)	2
Drotaverine hydrochloride	1	(1.3%)	1	0			1	(2.6%)	1	0			2	(1.2%)	2
Ibuprofen	1	(1.3%)	1	1	(2.4%)	1	0			0			2	(1.2%)	2
Acetylsalicylic acid	0			0			1	(2.6%)	1	0			1	(0.6%)	1
Aluminium acetotartrate	1	(1.3%)	1	0			0			0			1	(0.6%)	1
Amlodipine besilate	0			0			0			1	(11.1%)	1	1	(0.6%)	1
Arnica montana	1	(1.3%)	1	0			0			0			1	(0.6%)	1
Nasal preparations	0			0			1	(2.6%)	1	0			1	(0.6%)	1
Throat preparations	0			0			1	(2.6%)	1	0			1	(0.6%)	1

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Exposed Not Randomized refers to all subjects exposed to THS 2.2 but not randomized. Overall Safety refers to all subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Appendix 15.3.6.3.2

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_conmed.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.13 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Alanine aminotransferase (IU/L) 0-31	Baseline	n	41	20	19	4	84
		Mean (SD)	12.5 (4.38)	13.4 (5.59)	13.7 (6.30)	16.8 (8.81)	13.2 (5.36)
		Median	12.0	12.5	14.0	14.5	12.0
		Min, Max	6, 28	7, 28	5, 29	9, 29	5, 29
		Low value - n (%)	0	0	0	0	0
		Normal value - n (%)	41 (51.3%)	20 (48.8%)	19 (48.7%)	4 (44.4%)	84 (49.7%)
		High value - n (%)	0	0	0	0	0
		Abnormal clinically significant - n (%)	0	0	0	0	0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.4

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_labchem.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.13 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Alanine aminotransferase (IU/L) 0-31	Day 6/ Discharge	n	40	20	19	0	79
		Mean (SD)	15.5 (5.71)	18.3 (9.18)	21.1 (9.15)		17.5 (7.84)
		Median	14.0	15.0	20.0		15.0
		Min, Max	7, 29	10, 40	9, 37		7, 40
		Low value - n (%)	0	0	0		0
		Normal value - n (%)	40 (50.0%)	17 (41.5%)	16 (41.0%)		73 (43.2%)
		High value - n (%)	0	3 (7.3%)	3 (7.7%)		6 (3.6%)
		Abnormal clinically significant - n (%)	0	0	0		0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.4

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_labchem.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.13 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Alanine aminotransferase (IU/L) 0-31	Change from n baseline		40	20	19	0	79
		Mean (SD)	2.9 (4.94)	4.9 (7.12)	7.4 (5.53)		4.5 (5.91)
		Median	2.0	4.0	6.0		3.0
		Min, Max	-9, 15	-4, 26	1, 18		-9, 26

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.4

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_labchem.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.13 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Alanine aminotransferase (IU/L) 0-41	Baseline	n	39	21	20	5	85
		Mean (SD)	19.1 (7.53)	22.6 (9.64)	21.7 (11.10)	24.4 (11.20)	20.9 (9.20)
		Median	17.0	21.0	18.5	20.0	19.0
		Min, Max	10, 42	10, 49	8, 54	12, 39	8, 54
		Low value - n (%)	0	0	0	0	0
		Normal value - n (%)	38 (47.5%)	20 (48.8%)	19 (48.7%)	5 (55.6%)	82 (48.5%)
		High value - n (%)	1 (1.3%)	1 (2.4%)	1 (2.6%)	0	3 (1.8%)
		Abnormal clinically significant - n (%)	0	0	0	0	0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.4

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_labchem.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.13 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Alanine aminotransferase (IU/L) 0-41	Day 6/ Discharge	n	39	20	20	0	79
		Mean (SD)	28.1 (15.55)	33.8 (19.89)	40.2 (17.02)		32.6 (17.61)
		Median	22.0	25.0	39.0		26.0
		Min, Max	12, 73	14, 86	19, 77		12, 86
		Low value - n (%)	0	0	0		0
		Normal value - n (%)	33 (41.3%)	14 (34.1%)	10 (25.6%)		57 (33.7%)
		High value - n (%)	6 (7.5%)	6 (14.6%)	10 (25.6%)		22 (13.0%)
		Abnormal clinically significant - n (%)	0	0	0		0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.4

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_labchem.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.13 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Alanine aminotransferase (IU/L) 0-41	Change from baseline	n	39	20	20	0	79
		Mean (SD)	8.9 (10.83)	10.9 (14.73)	18.6 (12.40)		11.9 (12.78)
		Median	4.0	6.5	15.0		8.0
		Min, Max	-3, 45	-8, 54	1, 47		-8, 54

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.4

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_labchem.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.13 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Aspartate aminotransferase (IU/L) 0-32	Baseline	n	41	20	19	4	84
		Mean (SD)	14.7 (2.50)	15.9 (2.45)	15.9 (3.29)	18.5 (2.39)	15.5 (2.78)
		Median	15.0	16.5	15.0	19.5	15.0
		Min, Max	10, 22	12, 20	11, 23	15, 20	10, 23
		Low value - n (%)	0	0	0	0	0
		Normal value - n (%)	41 (51.3%)	20 (48.8%)	19 (48.7%)	4 (44.4%)	84 (49.7%)
		High value - n (%)	0	0	0	0	0
		Abnormal clinically significant - n (%)	0	0	0	0	0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.4

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_labchem.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.13 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Aspartate aminotransferase (IU/L) 0-32	Day 6/ Discharge	n	40	20	19	0	79
		Mean (SD)	15.7 (3.58)	17.8 (5.66)	19.8 (7.44)		17.2 (5.48)
		Median	15.0	16.5	19.0		16.0
		Min, Max	10, 27	14, 39	12, 44		10, 44
		Low value - n (%)	0	0	0		0
		Normal value - n (%)	40 (50.0%)	19 (46.3%)	18 (46.2%)		77 (45.6%)
		High value - n (%)	0	1 (2.4%)	1 (2.6%)		2 (1.2%)
		Abnormal clinically significant - n (%)	0	0	0		0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.4

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_labchem.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.13 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Aspartate aminotransferase (IU/L) 0-32	Change from baseline	n	40	20	19	0	79
		Mean (SD)	1.0 (3.09)	1.9 (4.78)	3.9 (6.30)		1.9 (4.58)
		Median	1.0	2.0	2.0		1.0
		Min, Max	-6, 11	-3, 19	-2, 27		-6, 27

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.4

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_labchem.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.13 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Aspartate aminotransferase (IU/L) 0-38	Baseline	n	39	21	20	5	85
		Mean (SD)	18.2 (3.91)	18.6 (3.73)	19.6 (5.39)	16.4 (4.28)	18.5 (4.27)
		Median	18.0	18.0	18.5	14.0	18.0
		Min, Max	12, 28	13, 26	13, 36	13, 22	12, 36
		Low value - n (%)	0	0	0	0	0
		Normal value - n (%)	39 (48.8%)	21 (51.2%)	20 (51.3%)	5 (55.6%)	85 (50.3%)
		High value - n (%)	0	0	0	0	0
		Abnormal clinically significant - n (%)	0	0	0	0	0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.13 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Aspartate aminotransferase (IU/L) 0-38	Day 6/ Discharge	n	39	20	20	0	79
		Mean (SD)	20.1 (5.76)	21.6 (7.37)	25.0 (6.66)		21.7 (6.65)
		Median	19.0	19.5	26.0		20.0
		Min, Max	12, 36	14, 39	16, 37		12, 39
		Low value - n (%)	0	0	0		0
		Normal value - n (%)	39 (48.8%)	18 (43.9%)	20 (51.3%)		77 (45.6%)
		High value - n (%)	0	2 (4.9%)	0		2 (1.2%)
		Abnormal clinically significant - n (%)	0	0	0		0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

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Path: /cvn/projects/prj/development/000000106324/dev/tables/t_labchem.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.13 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Aspartate aminotransferase (IU/L) 0-38	Change from baseline	n	39	20	20	0	79
		Mean (SD)	1.9 (4.21)	3.1 (6.33)	5.4 (5.66)		3.1 (5.32)
		Median	1.0	0.5	4.5		2.0
		Min, Max	-4, 14	-5, 20	-5, 17		-5, 20

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.13 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Bilirubin (mg/dL) 0-1	Baseline	n	80	41	39	9	169
		Mean (SD)	0.58 (0.231)	0.59 (0.246)	0.57 (0.246)	0.38 (0.084)	0.57 (0.236)
		Median	0.50	0.60	0.50	0.40	0.50
		Min, Max	0.2, 1.2	0.2, 1.3	0.3, 1.5	0.3, 0.5	0.2, 1.5
		Low value - n (%)	0	0	0	0	0
		Normal value - n (%)	77 (96.3%)	39 (95.1%)	38 (97.4%)	9 (100 %)	163 (96.4%)
		High value - n (%)	3 (3.8%)	2 (4.9%)	1 (2.6%)	0	6 (3.6%)
		Abnormal clinically significant - n (%)	0	0	0	0	0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.4

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_labchem.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.13 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Bilirubin (mg/dL) 0-1	Day 6/ Discharge	n	79	40	39	0	158
		Mean (SD)	0.63 (0.293)	0.56 (0.190)	0.49 (0.199)		0.58 (0.254)
		Median	0.60	0.60	0.50		0.50
		Min, Max	0.3, 1.7	0.3, 1.0	0.2, 1.1		0.2, 1.7
		Low value - n (%)	0	0	0		0
		Normal value - n (%)	72 (90.0%)	40 (97.6%)	38 (97.4%)		150 (88.8%)
		High value - n (%)	7 (8.8%)	0	1 (2.6%)		8 (4.7%)
		Abnormal clinically significant - n (%)	2 (2.5%)	0	0		2 (1.2%)

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.13 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Bilirubin (mg/dL) 0-1	Change from baseline	n	79	40	39	0	158
		Mean (SD)	0.05 (0.221)	-0.02 (0.185)	-0.07 (0.178)		0.00 (0.208)
		Median	0.00	0.00	0.00		0.00
		Min, Max	-0.5, 0.8	-0.6, 0.3	-0.5, 0.2		-0.6, 0.8

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.4

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.13 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Direct bilirubin (mg/dL) 0-0.3	Baseline	n	80	41	39	9	169
		Mean (SD)	0.21 (0.084)	0.21 (0.078)	0.21 (0.081)	0.13 (0.050)	0.21 (0.082)
		Median	0.20	0.20	0.20	0.10	0.20
		Min, Max	0.1, 0.4	0.0, 0.4	0.1, 0.5	0.1, 0.2	0.0, 0.5
		Low value - n (%)	0	0	0	0	0
		Normal value - n (%)	74 (92.5%)	40 (97.6%)	37 (94.9%)	9 (100 %)	160 (94.7%)
		High value - n (%)	6 (7.5%)	1 (2.4%)	2 (5.1%)	0	9 (5.3%)
		Abnormal clinically significant - n (%)	0	0	0	0	0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.13 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Direct bilirubin (mg/dL) 0-0.3	Day 6/ Discharge	n	79	40	39	0	158
		Mean (SD)	0.22 (0.089)	0.19 (0.071)	0.19 (0.074)		0.21 (0.083)
		Median	0.20	0.20	0.20		0.20
		Min, Max	0.1, 0.6	0.1, 0.3	0.1, 0.4		0.1, 0.6
		Low value - n (%)	0	0	0		0
		Normal value - n (%)	72 (90.0%)	40 (97.6%)	38 (97.4%)		150 (88.8%)
		High value - n (%)	7 (8.8%)	0	1 (2.6%)		8 (4.7%)
		Abnormal clinically significant - n (%)	0	0	0		0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

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Path: /cvn/projects/prj/development/000000106324/dev/tables/t_labchem.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.13 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Direct bilirubin (mg/dL) 0-0.3	Change from baseline	n	79	40	39	0	158
		Mean (SD)	0.01 (0.074)	-0.02 (0.068)	-0.03 (0.068)		-0.01 (0.073)
		Median	0.00	0.00	0.00		0.00
		Min, Max	-0.2, 0.2	-0.1, 0.2	-0.2, 0.1		-0.2, 0.2

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.13 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Alkaline phosphatase (IU/L) 35-104	Baseline	n	41	20	19	4	84
		Mean (SD)	58.6 (19.29)	60.0 (9.15)	59.1 (15.44)	76.5 (28.22)	59.9 (17.13)
		Median	56.0	57.5	58.0	76.5	57.0
		Min, Max	35, 118	45, 79	42, 98	43, 110	35, 118
		Low value - n (%)	0	0	0	0	0
		Normal value - n (%)	40 (50.0%)	20 (48.8%)	19 (48.7%)	3 (33.3%)	82 (48.5%)
		High value - n (%)	1 (1.3%)	0	0	1 (11.1%)	2 (1.2%)
		Abnormal clinically significant - n (%)	0	0	0	0	0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.13 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Alkaline phosphatase (IU/L) 35-104	Day 6/ Discharge	n	40	20	19	0	79
		Mean (SD)	60.6 (19.01)	63.2 (9.69)	61.9 (15.48)		61.6 (16.13)
		Median	56.0	61.5	61.0		59.0
		Min, Max	38, 121	51, 82	40, 97		38, 121
		Low value - n (%)	0	0	0		0
		Normal value - n (%)	39 (48.8%)	20 (48.8%)	19 (48.7%)		78 (46.2%)
		High value - n (%)	1 (1.3%)	0	0		1 (0.6%)
		Abnormal clinically significant - n (%)	0	0	0		0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.13 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Alkaline phosphatase (IU/L) 35-104	Change from baseline		40	20	19	0	79
		Mean (SD)	2.1 (5.34)	3.2 (4.14)	2.8 (6.51)		2.6 (5.33)
		Median	1.0	3.0	2.0		1.0
		Min, Max	-9, 16	-4, 11	-10, 17		-10, 17

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.13 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Alkaline phosphatase (IU/L) 40-129	Baseline	n	39	21	20	5	85
		Mean (SD)	64.6 (13.15)	71.0 (18.13)	67.1 (14.63)	80.6 (8.80)	67.7 (15.03)
		Median	64.0	75.0	69.0	82.0	68.0
		Min, Max	38, 89	41, 109	40, 99	71, 89	38, 109
		Low value - n (%)	2 (2.5%)	0	0	0	2 (1.2%)
		Normal value - n (%)	37 (46.3%)	21 (51.2%)	20 (51.3%)	5 (55.6%)	83 (49.1%)
		High value - n (%)	0	0	0	0	0
		Abnormal clinically significant - n (%)	0	0	0	0	0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.4

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_labchem.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.13 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Alkaline phosphatase (IU/L) 40-129	Day 6/ Discharge	n	39	20	20	0	79
		Mean (SD)	66.5 (12.67)	74.5 (18.42)	72.2 (15.92)		70.0 (15.33)
		Median	67.0	73.0	73.0		71.0
		Min, Max	39, 100	46, 112	46, 107		39, 112
		Low value - n (%)	1 (1.3%)	0	0		1 (0.6%)
		Normal value - n (%)	38 (47.5%)	20 (48.8%)	20 (51.3%)		78 (46.2%)
		High value - n (%)	0	0	0		0
		Abnormal clinically significant - n (%)	0	0	0		0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.4

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_labchem.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.13 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Alkaline phosphatase (IU/L) 40-129	Change from baseline		39	20	20	0	79
		Mean (SD)	1.9 (4.95)	3.1 (5.07)	5.1 (4.84)		3.0 (5.06)
		Median	3.0	3.0	4.0		3.0
		Min, Max	-7, 12	-6, 12	-2, 17		-7, 17

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.4

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.13 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Gamma glutamyl transferase (IU/L) 5-36	Baseline	n	41	20	19	4	84
		Mean (SD)	14.6 (7.16)	21.8 (17.05)	14.3 (6.74)	28.0 (32.30)	16.9 (12.44)
		Median	13.0	16.0	14.0	14.0	14.0
		Min, Max	6, 48	7, 72	5, 33	8, 76	5, 76
		Low value - n (%)	0	0	0	0	0
		Normal value - n (%)	40 (50.0%)	17 (41.5%)	19 (48.7%)	3 (33.3%)	79 (46.7%)
		High value - n (%)	1 (1.3%)	3 (7.3%)	0	1 (11.1%)	5 (3.0%)
		Abnormal clinically significant - n (%)	0	0	0	0	0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.4

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_labchem.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.13 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Gamma glutamyl transferase (IU/L) 5-36	Day 6/ Discharge	n	40	20	19	0	79
		Mean (SD)	14.8 (7.25)	22.5 (13.46)	15.3 (6.37)		16.8 (9.52)
		Median	13.0	16.0	14.0		15.0
		Min, Max	6, 45	8, 55	6, 28		6, 55
		Low value - n (%)	0	0	0		0
		Normal value - n (%)	39 (48.8%)	16 (39.0%)	19 (48.7%)		74 (43.8%)
		High value - n (%)	1 (1.3%)	4 (9.8%)	0		5 (3.0%)
		Abnormal clinically significant - n (%)	0	0	0		0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.4

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.13 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Gamma glutamyl transferase (IU/L) 5-36	Change from n baseline		40	20	19	0	79
		Mean (SD)	0.2 (1.65)	0.7 (7.44)	1.0 (2.54)		0.5 (4.06)
		Median	0.0	1.0	1.0		0.0
		Min, Max	-3, 6	-17, 13	-5, 5		-17, 13

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

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Path: /cvn/projects/prj/development/000000106324/dev/tables/t_labchem.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.13 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Gamma glutamyl transferase (IU/L) 8-61	Baseline	n	39	21	20	5	85
		Mean (SD)	28.0 (14.89)	23.1 (11.71)	23.8 (14.54)	31.6 (13.05)	26.0 (13.99)
		Median	22.0	23.0	19.5	37.0	22.0
		Min, Max	6, 62	8, 63	9, 70	12, 46	6, 70
		Low value - n (%)	1 (1.3%)	0	0	0	1 (0.6%)
		Normal value - n (%)	37 (46.3%)	20 (48.8%)	19 (48.7%)	5 (55.6%)	81 (47.9%)
		High value - n (%)	1 (1.3%)	1 (2.4%)	1 (2.6%)	0	3 (1.8%)
		Abnormal clinically significant - n (%)	0	0	0	0	0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

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**Table 15.2.6.13 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Gamma glutamyl transferase (IU/L) 8-61	Day 6/ Discharge	n	39	20	20	0	79
		Mean (SD)	29.0 (15.37)	26.6 (18.07)	29.5 (16.39)		28.5 (16.17)
		Median	25.0	23.5	25.5		24.0
		Min, Max	7, 62	9, 89	10, 79		7, 89
		Low value - n (%)	1 (1.3%)	0	0		1 (0.6%)
		Normal value - n (%)	37 (46.3%)	19 (46.3%)	19 (48.7%)		75 (44.4%)
		High value - n (%)	1 (1.3%)	1 (2.4%)	1 (2.6%)		3 (1.8%)
		Abnormal clinically significant - n (%)	0	0	0		0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.4

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.13 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Gamma glutamyl transferase (IU/L) 8-61	Change from n baseline		39	20	20	0	79
		Mean (SD)	1.0 (5.17)	3.1 (8.78)	5.7 (4.82)		2.7 (6.42)
		Median	0.0	0.5	4.0		1.0
		Min, Max	-9, 18	-5, 26	0, 15		-9, 26

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.13 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Lactate dehydrogenase (IU/L) 240-480	Baseline	n	80	41	39	9	169
		Mean (SD)	289.1 (35.88)	294.2 (49.50)	296.8 (37.86)	324.8 (61.39)	294.0 (41.91)
		Median	285.0	299.0	287.0	317.0	288.0
		Min, Max	200, 387	143, 427	222, 391	260, 445	143, 445
		Low value - n (%)	6 (7.5%)	4 (9.8%)	1 (2.6%)	0	11 (6.5%)
		Normal value - n (%)	74 (92.5%)	37 (90.2%)	38 (97.4%)	9 (100 %)	158 (93.5%)
		High value - n (%)	0	0	0	0	0
		Abnormal clinically significant - n (%)	0	0	0	0	0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

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**Table 15.2.6.13 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Lactate dehydrogenase (IU/L) 240-480	Day 6/ Discharge	n	79	40	39	0	158
		Mean (SD)	281.8 (35.97)	295.1 (58.87)	286.8 (37.62)		286.4 (43.32)
		Median	282.0	296.0	283.0		283.0
		Min, Max	189, 404	138, 445	203, 357		138, 445
		Low value - n (%)	6 (7.5%)	4 (9.8%)	3 (7.7%)		13 (7.7%)
		Normal value - n (%)	73 (91.3%)	36 (87.8%)	36 (92.3%)		145 (85.8%)
		High value - n (%)	0	0	0		0
		Abnormal clinically significant - n (%)	0	0	0		0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.13 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Lactate dehydrogenase (IU/L) 240-480	Change from baseline	n	79	40	39	0	158
		Mean (SD)	-7.6 (28.85)	1.9 (33.75)	-10.0 (28.36)		-5.8 (30.20)
		Median	-8.0	-7.0	-8.0		-8.0
		Min, Max	-87, 62	-59, 81	-65, 50		-87, 81

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.13 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Glucose (mg/dL) 70-105	Baseline	n	80	41	39	9	169
		Mean (SD)	87.6 (9.09)	86.2 (9.15)	86.7 (10.78)	94.4 (21.34)	87.4 (10.48)
		Median	86.0	85.0	86.0	88.0	86.0
		Min, Max	73, 114	59, 113	70, 112	77, 145	59, 145
		Low value - n (%)	0	1 (2.4%)	0	0	1 (0.6%)
		Normal value - n (%)	76 (95.0%)	39 (95.1%)	36 (92.3%)	7 (77.8%)	158 (93.5%)
		High value - n (%)	4 (5.0%)	1 (2.4%)	3 (7.7%)	2 (22.2%)	10 (5.9%)
		Abnormal clinically significant - n (%)	0	0	0	1 (11.1%)	1 (0.6%)

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.13 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Glucose (mg/dL) 70-105	Day 6/ Discharge	n	79	40	39	0	158
		Mean (SD)	83.3 (9.97)	87.2 (13.26)	87.6 (14.05)		85.3 (12.05)
		Median	82.0	84.0	84.0		83.0
		Min, Max	58, 113	65, 119	61, 130		58, 130
		Low value - n (%)	4 (5.0%)	1 (2.4%)	1 (2.6%)		6 (3.6%)
		Normal value - n (%)	73 (91.3%)	34 (82.9%)	33 (84.6%)		140 (82.8%)
		High value - n (%)	2 (2.5%)	5 (12.2%)	5 (12.8%)		12 (7.1%)
		Abnormal clinically significant - n (%)	0	0	1 (2.6%)		1 (0.6%)

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.13 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Glucose (mg/dL) 70-105	Change from baseline	n	79	40	39	0	158
		Mean (SD)	-4.4 (10.98)	1.0 (10.60)	0.9 (11.73)		-1.7 (11.33)
		Median	-4.0	-0.5	0.0		-2.5
		Min, Max	-29, 31	-17, 28	-18, 49		-29, 49

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.13 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Creatinine (mg/dL) 0.5-0.9	Baseline	n	41	20	19	4	84
		Mean (SD)	0.622 (0.0881)	0.595 (0.0760)	0.584 (0.0959)	0.550 (0.1291)	0.604 (0.0898)
		Median	0.600	0.600	0.600	0.550	0.600
		Min, Max	0.50, 0.80	0.50, 0.80	0.40, 0.80	0.40, 0.70	0.40, 0.80
		Low value - n (%)	0	0	1 (2.6%)	1 (11.1%)	2 (1.2%)
		Normal value - n (%)	41 (51.3%)	20 (48.8%)	18 (46.2%)	3 (33.3%)	82 (48.5%)
		High value - n (%)	0	0	0	0	0
		Abnormal clinically significant - n (%)	0	0	0	0	0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.13 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Creatinine (mg/dL) 0.5-0.9	Day 6/ Discharge	n	40	20	19	0	79
		Mean (SD)	0.620 (0.0758)	0.595 (0.0888)	0.563 (0.1066)		0.600 (0.0892)
		Median	0.600	0.600	0.600		0.600
		Min, Max	0.50, 0.80	0.40, 0.80	0.40, 0.70		0.40, 0.80
		Low value - n (%)	0	1 (2.4%)	3 (7.7%)		4 (2.4%)
		Normal value - n (%)	40 (50.0%)	19 (46.3%)	16 (41.0%)		75 (44.4%)
		High value - n (%)	0	0	0		0
		Abnormal clinically significant - n (%)	0	0	0		0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.4

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_labchem.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.13 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Creatinine (mg/dL) 0.5-0.9	Change from baseline	n	40	20	19	0	79
		Mean (SD)	0.000 (0.0717)	0.000 (0.0562)	-0.021 (0.0631)		-0.005 (0.0659)
		Median	0.000	0.000	0.000		0.000
		Min, Max	-0.10, 0.10	-0.10, 0.10	-0.10, 0.10		-0.10, 0.10

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.4

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_labchem.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.13 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Creatinine (mg/dL) 0.7-1.2	Baseline	n	39	21	20	5	85
		Mean (SD)	0.754 (0.1023)	0.795 (0.0865)	0.800 (0.1214)	0.740 (0.1141)	0.774 (0.1049)
		Median	0.700	0.800	0.800	0.700	0.800
		Min, Max	0.60, 1.00	0.60, 1.00	0.50, 1.00	0.60, 0.90	0.50, 1.00
		Low value - n (%)	6 (7.5%)	1 (2.4%)	2 (5.1%)	1 (11.1%)	10 (5.9%)
		Normal value - n (%)	33 (41.3%)	20 (48.8%)	18 (46.2%)	4 (44.4%)	75 (44.4%)
		High value - n (%)	0	0	0	0	0
		Abnormal clinically significant - n (%)	0	0	0	0	0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.4

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_labchem.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.13 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Creatinine (mg/dL) 0.7-1.2	Day 6/ Discharge	n	39	20	20	0	79
		Mean (SD)	0.785 (0.0905)	0.800 (0.1170)	0.820 (0.0952)		0.797 (0.0987)
		Median	0.800	0.800	0.800		0.800
		Min, Max	0.60, 1.00	0.60, 1.10	0.70, 1.10		0.60, 1.10
		Low value - n (%)	1 (1.3%)	1 (2.4%)	0		2 (1.2%)
		Normal value - n (%)	38 (47.5%)	19 (46.3%)	20 (51.3%)		77 (45.6%)
		High value - n (%)	0	0	0		0
		Abnormal clinically significant - n (%)	0	0	0		0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.4

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_labchem.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.13 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Creatinine (mg/dL) 0.7-1.2	Change from baseline	n	39	20	20	0	79
		Mean (SD)	0.031 (0.0656)	0.000 (0.0859)	0.020 (0.0895)		0.020 (0.0775)
		Median	0.000	0.000	0.000		0.000
		Min, Max	-0.10, 0.10	-0.10, 0.20	-0.10, 0.20		-0.10, 0.20

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.4

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_labchem.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.13 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Protein (g/dL) 64-83	Baseline	n	80	41	39	9	169
		Mean (SD)	69.74 (3.822)	69.85 (3.954)	69.56 (4.198)	70.22 (2.949)	69.75 (3.875)
		Median	69.00	69.00	69.00	70.00	69.00
		Min, Max	61.0, 79.0	62.0, 78.0	59.0, 77.0	65.0, 74.0	59.0, 79.0
		Low value - n (%)	4 (5.0%)	1 (2.4%)	2 (5.1%)	0	7 (4.1%)
		Normal value - n (%)	76 (95.0%)	40 (97.6%)	37 (94.9%)	9 (100 %)	162 (95.9%)
		High value - n (%)	0	0	0	0	0
		Abnormal clinically significant - n (%)	0	0	0	0	0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.4

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_labchem.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.13 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Protein (g/dL) 64-83	Day 6/ Discharge	n	79	40	39	0	158
		Mean (SD)	69.84 (3.376)	69.73 (3.336)	70.38 (4.089)		69.94 (3.541)
		Median	70.00	70.00	71.00		70.00
		Min, Max	59.0, 78.0	62.0, 76.0	62.0, 80.0		59.0, 80.0
		Low value - n (%)	2 (2.5%)	2 (4.9%)	2 (5.1%)		6 (3.6%)
		Normal value - n (%)	77 (96.3%)	38 (92.7%)	37 (94.9%)		152 (89.9%)
		High value - n (%)	0	0	0		0
		Abnormal clinically significant - n (%)	0	0	0		0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.4

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_labchem.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.13 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Protein (g/dL) 64-83	Change from baseline	n	79	40	39	0	158
		Mean (SD)	0.10 (3.533)	0.08 (4.048)	0.82 (3.663)		0.27 (3.690)
		Median	0.00	0.00	1.00		0.00
		Min, Max	-8.0, 7.0	-7.0, 8.0	-8.0, 8.0		-8.0, 8.0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.4

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.13 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Blood urea nitrogen (mg/dL) 5-23	Baseline	n	80	41	39	9	169
		Mean (SD)	12.3 (2.72)	12.1 (2.36)	12.4 (2.92)	12.1 (2.38)	12.3 (2.65)
		Median	12.3	12.2	11.9	12.2	12.2
		Min, Max	6, 20	6, 16	8, 20	8, 16	6, 20
		Low value - n (%)	0	0	0	0	0
		Normal value - n (%)	80 (100 %)	41 (100 %)	39 (100 %)	9 (100 %)	169 (100 %)
		High value - n (%)	0	0	0	0	0
		Abnormal clinically significant - n (%)	0	0	0	0	0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.4

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_labchem.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.13 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Blood urea nitrogen (mg/dL) 5-23	Day 6/ Discharge	n	79	40	39	0	158
		Mean (SD)	13.1 (2.77)	12.6 (2.86)	13.2 (2.90)		13.0 (2.82)
		Median	12.8	12.1	12.9		12.6
		Min, Max	5, 21	9, 21	8, 20		5, 21
		Low value - n (%)	0	0	0		0
		Normal value - n (%)	79 (98.8%)	40 (97.6%)	39 (100 %)		158 (93.5%)
		High value - n (%)	0	0	0		0
		Abnormal clinically significant - n (%)	0	0	0		0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.4

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_labchem.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.13 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Blood urea nitrogen (mg/dL) 5-23	Change from baseline	n	79	40	39	0	158
		Mean (SD)	0.8 (2.32)	0.4 (3.22)	0.8 (2.26)		0.7 (2.55)
		Median	0.8	0.1	1.0		0.5
		Min, Max	-5, 7	-7, 8	-3, 6		-7, 8

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.4

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_labchem.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.13 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Triglycerides (mg/dL) 50-200	Baseline	n	80	41	39	9	169
		Mean (SD)	109.3 (45.07)	108.7 (51.19)	110.2 (46.53)	157.2 (56.94)	111.9 (48.37)
		Median	97.5	92.0	98.0	165.0	99.0
		Min, Max	39, 308	44, 273	45, 254	94, 225	39, 308
		Low value - n (%)	1 (1.3%)	1 (2.4%)	1 (2.6%)	0	3 (1.8%)
		Normal value - n (%)	78 (97.5%)	38 (92.7%)	36 (92.3%)	6 (66.7%)	158 (93.5%)
		High value - n (%)	1 (1.3%)	2 (4.9%)	2 (5.1%)	3 (33.3%)	8 (4.7%)
		Abnormal clinically significant - n (%)	1 (1.3%)	0	0	0	1 (0.6%)

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.4

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_labchem.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.13 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Triglycerides (mg/dL) 50-200	Day 6/ Discharge	n	79	40	39	0	158
		Mean (SD)	98.9 (64.68)	104.5 (86.27)	128.5 (71.48)		107.6 (72.89)
		Median	80.0	85.5	106.0		87.0
		Min, Max	34, 358	43, 552	38, 345		34, 552
		Low value - n (%)	5 (6.3%)	4 (9.8%)	1 (2.6%)		10 (5.9%)
		Normal value - n (%)	68 (85.0%)	34 (82.9%)	32 (82.1%)		134 (79.3%)
		High value - n (%)	6 (7.5%)	2 (4.9%)	6 (15.4%)		14 (8.3%)
		Abnormal clinically significant - n (%)	3 (3.8%)	1 (2.4%)	2 (5.1%)		6 (3.6%)

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.4

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_labchem.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.13 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Triglycerides (mg/dL) 50-200	Change from baseline	n	79	40	39	0	158
		Mean (SD)	-10.8 (46.13)	-4.9 (56.46)	18.4 (50.87)		-2.1 (51.19)
		Median	-19.0	-13.0	7.0		-12.0
		Min, Max	-95, 204	-77, 279	-90, 158		-95, 279

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.4

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.13 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Cholesterol (mg/dL) 0-200	Baseline	n	80	41	39	9	169
		Mean (SD)	192.4 (33.51)	186.5 (30.12)	181.9 (41.83)	224.4 (50.43)	190.3 (36.68)
		Median	189.5	178.0	178.0	235.0	185.0
		Min, Max	120, 266	145, 278	120, 327	147, 295	120, 327
		Low value - n (%)	0	0	0	0	0
		Normal value - n (%)	45 (56.3%)	29 (70.7%)	30 (76.9%)	4 (44.4%)	108 (63.9%)
		High value - n (%)	35 (43.8%)	12 (29.3%)	9 (23.1%)	5 (55.6%)	61 (36.1%)
		Abnormal clinically significant - n (%)	0	0	1 (2.6%)	0	1 (0.6%)

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.4

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.13 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Cholesterol (mg/dL) 0-200	Day 6/ Discharge	n	79	40	39	0	158
		Mean (SD)	190.6 (30.44)	189.6 (37.52)	192.3 (39.60)		190.8 (34.50)
		Median	189.0	187.5	186.0		188.0
		Min, Max	127, 266	137, 331	122, 285		122, 331
		Low value - n (%)	0	0	0		0
		Normal value - n (%)	46 (57.5%)	25 (61.0%)	26 (66.7%)		97 (57.4%)
		High value - n (%)	33 (41.3%)	15 (36.6%)	13 (33.3%)		61 (36.1%)
		Abnormal clinically significant - n (%)	0	1 (2.4%)	0		1 (0.6%)

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.4

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_labchem.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.13 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Cholesterol (mg/dL) 0-200	Change from baseline	n	79	40	39	0	158
		Mean (SD)	-2.7 (18.16)	2.7 (22.33)	10.4 (15.83)		1.9 (19.42)
		Median	0.0	-2.5	14.0		2.0
		Min, Max	-49, 40	-44, 57	-42, 42		-49, 57

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.13 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Albumin (g/L) 35-52	Baseline	n	80	41	39	9	169
		Mean (SD)	47.7 (2.54)	47.4 (2.36)	47.7 (2.54)	47.4 (2.07)	47.6 (2.46)
		Median	48.0	48.0	48.0	47.0	48.0
		Min, Max	40, 53	42, 52	42, 52	45, 50	40, 53
		Low value - n (%)	0	0	0	0	0
		Normal value - n (%)	79 (98.8%)	41 (100 %)	39 (100 %)	9 (100 %)	168 (99.4%)
		High value - n (%)	1 (1.3%)	0	0	0	1 (0.6%)
		Abnormal clinically significant - n (%)	0	0	0	0	0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.4

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_labchem.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.13 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Albumin (g/L) 35-52	Day 6/ Discharge	n	79	40	39	0	158
		Mean (SD)	48.7 (2.63)	48.5 (2.25)	49.5 (2.51)		48.8 (2.52)
		Median	49.0	49.0	50.0		49.0
		Min, Max	43, 56	42, 52	42, 55		42, 56
		Low value - n (%)	0	0	0		0
		Normal value - n (%)	75 (93.8%)	40 (97.6%)	34 (87.2%)		149 (88.2%)
		High value - n (%)	4 (5.0%)	0	5 (12.8%)		9 (5.3%)
		Abnormal clinically significant - n (%)	0	0	0		0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.4

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_labchem.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.13 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Albumin (g/L) 35-52	Change from baseline	n	79	40	39	0	158
		Mean (SD)	1.0 (2.07)	1.2 (2.68)	1.8 (2.01)		1.2 (2.24)
		Median	1.0	2.0	2.0		1.0
		Min, Max	-5, 6	-5, 7	-2, 6		-5, 7

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.4

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_labchem.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.13 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Sodium (mmol/L) 136-146	Baseline	n	80	41	39	9	169
		Mean (SD)	141.3 (1.57)	141.0 (1.38)	141.4 (1.29)	142.1 (1.46)	141.3 (1.47)
		Median	141.0	141.0	141.0	142.0	141.0
		Min, Max	137, 145	138, 144	139, 144	140, 144	137, 145
		Low value - n (%)	0	0	0	0	0
		Normal value - n (%)	80 (100 %)	41 (100 %)	39 (100 %)	9 (100 %)	169 (100 %)
		High value - n (%)	0	0	0	0	0
		Abnormal clinically significant - n (%)	0	0	0	0	0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.4

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_labchem.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.13 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Sodium (mmol/L) 136-146	Day 6/ Discharge	n	79	40	39	0	158
		Mean (SD)	140.5 (1.78)	140.3 (1.71)	140.6 (1.62)		140.4 (1.72)
		Median	140.0	140.0	141.0		140.0
		Min, Max	138, 147	137, 144	137, 145		137, 147
		Low value - n (%)	0	0	0		0
		Normal value - n (%)	78 (97.5%)	40 (97.6%)	39 (100 %)		157 (92.9%)
		High value - n (%)	1 (1.3%)	0	0		1 (0.6%)
		Abnormal clinically significant - n (%)	0	0	0		0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.4

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_labchem.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.13 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Sodium (mmol/L) 136-146	Change from baseline	n	79	40	39	0	158
		Mean (SD)	-0.8 (1.68)	-0.7 (1.92)	-0.8 (1.44)		-0.8 (1.68)
		Median	-1.0	-1.0	-1.0		-1.0
		Min, Max	-4, 6	-5, 5	-3, 2		-5, 6

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.4

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_labchem.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.13 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Potassium (mmol/L) 3.5-5.1	Baseline	n	80	41	39	9	169
		Mean (SD)	4.39 (0.317)	4.42 (0.261)	4.34 (0.350)	4.63 (0.548)	4.40 (0.331)
		Median	4.30	4.40	4.30	4.50	4.40
		Min, Max	3.5, 5.4	3.9, 5.1	3.9, 5.4	4.0, 5.8	3.5, 5.8
		Low value - n (%)	0	0	0	0	0
		Normal value - n (%)	78 (97.5%)	41 (100 %)	38 (97.4%)	8 (88.9%)	165 (97.6%)
		High value - n (%)	2 (2.5%)	0	1 (2.6%)	1 (11.1%)	4 (2.4%)
		Abnormal clinically significant - n (%)	0	0	0	1 (11.1%)	1 (0.6%)

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.4

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_labchem.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.13 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Potassium (mmol/L) 3.5-5.1	Day 6/ Discharge	n	79	40	39	0	158
		Mean (SD)	4.55 (0.353)	4.52 (0.319)	4.35 (0.248)		4.49 (0.331)
		Median	4.60	4.50	4.40		4.50
		Min, Max	3.9, 5.8	4.1, 5.7	3.8, 4.9		3.8, 5.8
		Low value - n (%)	0	0	0		0
		Normal value - n (%)	76 (95.0%)	39 (95.1%)	39 (100 %)		154 (91.1%)
		High value - n (%)	3 (3.8%)	1 (2.4%)	0		4 (2.4%)
		Abnormal clinically significant - n (%)	1 (1.3%)	0	0		1 (0.6%)

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.4

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_labchem.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.13 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Potassium (mmol/L) 3.5-5.1	Change from baseline	n	79	40	39	0	158
		Mean (SD)	0.15 (0.427)	0.10 (0.425)	0.01 (0.347)		0.10 (0.410)
		Median	0.10	0.10	-0.10		0.10
		Min, Max	-0.8, 1.7	-0.8, 1.3	-1.0, 1.0		-1.0, 1.7

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.4

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_labchem.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.14 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Erythrocytes (T/L) 3.5-5 (Females)	Baseline	n	41	20	19	4	84
		Mean (SD)	4.386 (0.3020)	4.220 (0.2338)	4.351 (0.3086)	4.110 (0.3727)	4.325 (0.2984)
		Median	4.390	4.245	4.280	4.225	4.310
		Min, Max	3.77, 4.96	3.74, 4.54	3.85, 4.87	3.57, 4.42	3.57, 4.96
		Low value - n (%)	0	0	0	0	0
		Normal value - n (%)	41 (51.3%)	20 (48.8%)	19 (48.7%)	4 (44.4%)	84 (49.7%)
		High value - n (%)	0	0	0	0	0
		Abnormal clinically significant - n (%)	0	0	0	0	0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.5

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_labhem.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.14 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Erythrocytes (T/L) 3.5-5 (Females)	Day 6/ Discharge	n	40	20	19	0	79
		Mean (SD)	4.318 (0.2496)	4.277 (0.2356)	4.302 (0.2596)		4.304 (0.2460)
		Median	4.270	4.270	4.290		4.290
		Min, Max	3.85, 5.06	3.84, 4.73	3.75, 4.68		3.75, 5.06
		Low value - n (%)	0	0	0		0
		Normal value - n (%)	39 (48.8%)	20 (48.8%)	19 (48.7%)		78 (46.2%)
		High value - n (%)	1 (1.3%)	0	0		1 (0.6%)
		Abnormal clinically significant - n (%)	0	0	0		0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.5

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_labhem.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.14 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Erythrocytes (T/L) 3.5-5 (Females)	Change from baseline	n	40	20	19	0	79
		Mean (SD)	-0.073 (0.1747)	0.058 (0.2037)	-0.049 (0.1912)		-0.034 (0.1918)
		Median	-0.080	0.075	-0.090		-0.070
		Min, Max	-0.39, 0.45	-0.30, 0.32	-0.41, 0.31		-0.41, 0.45

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.5

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_labhem.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.14 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Erythrocytes (T/L) 4.5-5.7 (Males)	Baseline	n	39	21	20	5	85
		Mean (SD)	4.974 (0.3932)	4.973 (0.2475)	4.901 (0.3017)	4.758 (0.3506)	4.944 (0.3378)
		Median	5.030	4.950	4.960	4.660	4.970
		Min, Max	3.80, 5.94	4.47, 5.37	4.31, 5.37	4.38, 5.25	3.80, 5.94
		Low value - n (%)	5 (6.3%)	1 (2.4%)	3 (7.7%)	1 (11.1%)	10 (5.9%)
		Normal value - n (%)	33 (41.3%)	20 (48.8%)	17 (43.6%)	4 (44.4%)	74 (43.8%)
		High value - n (%)	1 (1.3%)	0	0	0	1 (0.6%)
		Abnormal clinically significant - n (%)	0	0	0	1 (11.1%)	1 (0.6%)

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.5

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_labhem.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.14 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Erythrocytes (T/L) 4.5-5.7 (Males)	Day 6/ Discharge	n	39	21	20	0	80
		Mean (SD)	4.918 (0.3361)	4.926 (0.2478)	4.938 (0.3296)		4.925 (0.3099)
		Median	4.920	4.870	4.945		4.925
		Min, Max	3.84, 5.78	4.50, 5.40	4.37, 5.62		3.84, 5.78
		Low value - n (%)	5 (6.3%)	0	2 (5.1%)		7 (4.1%)
		Normal value - n (%)	33 (41.3%)	21 (51.2%)	18 (46.2%)		72 (42.6%)
		High value - n (%)	1 (1.3%)	0	0		1 (0.6%)
		Abnormal clinically significant - n (%)	0	0	0		0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.5

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_labhem.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.14 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Erythrocytes (T/L) 4.5-5.7 (Males)	Change from baseline	n	39	21	20	0	80
		Mean (SD)	-0.056 (0.3923)	-0.048 (0.2192)	0.038 (0.1835)		-0.030 (0.3096)
		Median	-0.030	-0.080	-0.010		-0.025
		Min, Max	-1.79, 1.17	-0.41, 0.47	-0.20, 0.48		-1.79, 1.17

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.5

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.14 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Hemoglobin (g/dL) 12-16	Baseline	n	41	20	19	4	84
		Mean (SD)	13.75 (1.040)	13.34 (0.830)	13.49 (0.945)	13.95 (0.786)	13.60 (0.965)
		Median	13.70	13.25	13.20	14.15	13.50
		Min, Max	11.0, 15.7	11.9, 14.8	12.4, 15.8	12.9, 14.6	11.0, 15.8
		Low value - n (%)	1 (1.3%)	1 (2.4%)	0	0	2 (1.2%)
		Normal value - n (%)	40 (50.0%)	19 (46.3%)	19 (48.7%)	4 (44.4%)	82 (48.5%)
		High value - n (%)	0	0	0	0	0
		Abnormal clinically significant - n (%)	0	0	0	0	0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.5

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.14 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Hemoglobin (g/dL) 12-16	Day 6/ Discharge	n	40	20	19	0	79
		Mean (SD)	13.52 (0.842)	13.42 (0.834)	13.39 (0.710)		13.46 (0.802)
		Median	13.60	13.35	13.20		13.50
		Min, Max	11.2, 15.3	11.8, 15.6	12.4, 14.9		11.2, 15.6
		Low value - n (%)	3 (3.8%)	1 (2.4%)	0		4 (2.4%)
		Normal value - n (%)	37 (46.3%)	19 (46.3%)	19 (48.7%)		75 (44.4%)
		High value - n (%)	0	0	0		0
		Abnormal clinically significant - n (%)	0	0	0		0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.5

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_labhem.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.14 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Hemoglobin (g/dL) 12-16	Change from baseline	n	40	20	19	0	79
		Mean (SD)	-0.25 (0.558)	0.08 (0.653)	-0.10 (0.612)		-0.13 (0.603)
		Median	-0.30	0.20	-0.10		-0.20
		Min, Max	-1.3, 1.3	-1.1, 0.9	-1.2, 1.1		-1.3, 1.3

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.5

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_labhem.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.14 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Hemoglobin (g/dL) 12-16.8	Baseline	n	39	21	20	5	85
		Mean (SD)	15.71 (1.320)	15.62 (0.732)	15.51 (1.077)	15.30 (0.652)	15.62 (1.101)
		Median	15.50	15.60	15.60	15.00	15.60
		Min, Max	13.9, 21.8	14.3, 17.1	13.5, 18.0	14.7, 16.1	13.5, 21.8
		Low value - n (%)	0	0	0	0	0
		Normal value - n (%)	36 (45.0%)	19 (46.3%)	19 (48.7%)	5 (55.6%)	79 (46.7%)
		High value - n (%)	3 (3.8%)	2 (4.9%)	1 (2.6%)	0	6 (3.6%)
		Abnormal clinically significant - n (%)	0	0	0	0	0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.5

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_labhem.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.14 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Hemoglobin (g/dL) 12-16.8	Day 6/ Discharge	n	39	21	20	0	80
		Mean (SD)	15.39 (0.845)	15.48 (0.745)	15.60 (1.147)		15.47 (0.899)
		Median	15.30	15.50	15.70		15.45
		Min, Max	14.1, 17.5	14.1, 17.0	13.5, 18.2		13.5, 18.2
		Low value - n (%)	0	0	0		0
		Normal value - n (%)	37 (46.3%)	20 (48.8%)	18 (46.2%)		75 (44.4%)
		High value - n (%)	2 (2.5%)	1 (2.4%)	2 (5.1%)		5 (3.0%)
		Abnormal clinically significant - n (%)	0	0	0		0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.5

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_labhem.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.14 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Hemoglobin (g/dL) 12-16.8	Change from baseline	n	39	21	20	0	80
		Mean (SD)	-0.32 (1.332)	-0.14 (0.637)	0.09 (0.513)		-0.17 (1.024)
		Median	-0.20	-0.10	0.15		-0.10
		Min, Max	-7.5, 1.2	-1.3, 1.2	-0.8, 1.2		-7.5, 1.2

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.5

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.14 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Hematocrit (%) 37-47	Baseline	n	41	20	19	4	84
		Mean (SD)	39.76 (2.589)	38.66 (2.351)	39.16 (2.454)	40.25 (3.060)	39.38 (2.527)
		Median	40.20	38.55	38.80	40.85	39.20
		Min, Max	33.0, 45.4	34.0, 43.1	36.7, 45.7	36.3, 43.0	33.0, 45.7
		Low value - n (%)	6 (7.5%)	6 (14.6%)	4 (10.3%)	1 (11.1%)	17 (10.1%)
		Normal value - n (%)	35 (43.8%)	14 (34.1%)	15 (38.5%)	3 (33.3%)	67 (39.6%)
		High value - n (%)	0	0	0	0	0
		Abnormal clinically significant - n (%)	0	0	0	0	0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

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Path: /cvn/projects/prj/development/000000106324/dev/tables/t_labhem.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.14 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Hematocrit (%) 37-47	Day 6/ Discharge	n	40	20	19	0	79
		Mean (SD)	38.86 (2.062)	38.91 (2.379)	38.50 (1.887)		38.78 (2.086)
		Median	38.95	38.50	38.10		38.70
		Min, Max	33.3, 43.2	35.1, 45.4	35.4, 42.3		33.3, 45.4
		Low value - n (%)	6 (7.5%)	3 (7.3%)	5 (12.8%)		14 (8.3%)
		Normal value - n (%)	34 (42.5%)	17 (41.5%)	14 (35.9%)		65 (38.5%)
		High value - n (%)	0	0	0		0
		Abnormal clinically significant - n (%)	0	0	0		0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.5

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.14 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Hematocrit (%) 37-47	Change from baseline	n	40	20	19	0	79
		Mean (SD)	-0.95 (1.610)	0.25 (1.917)	-0.66 (1.676)		-0.58 (1.757)
		Median	-1.15	0.50	-0.70		-0.90
		Min, Max	-3.4, 4.3	-3.4, 2.7	-3.9, 2.1		-3.9, 4.3

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.5

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.14 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Hematocrit (%) 40-49.5	Baseline	n	39	21	20	5	85
		Mean (SD)	44.89 (3.215)	44.66 (1.996)	43.87 (2.467)	43.38 (1.762)	44.51 (2.720)
		Median	44.50	44.80	43.85	44.40	44.50
		Min, Max	40.5, 59.8	41.2, 47.9	39.7, 50.3	41.0, 44.9	39.7, 59.8
		Low value - n (%)	0	0	1 (2.6%)	0	1 (0.6%)
		Normal value - n (%)	38 (47.5%)	21 (51.2%)	18 (46.2%)	5 (55.6%)	82 (48.5%)
		High value - n (%)	1 (1.3%)	0	1 (2.6%)	0	2 (1.2%)
		Abnormal clinically significant - n (%)	0	0	0	0	0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.14 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Hematocrit (%) 40-49.5	Day 6/ Discharge	n	39	21	20	0	80
		Mean (SD)	43.89 (2.181)	43.90 (1.906)	43.84 (2.479)		43.88 (2.165)
		Median	43.40	44.50	44.20		43.80
		Min, Max	40.0, 47.9	41.2, 46.7	39.9, 49.8		39.9, 49.8
		Low value - n (%)	0	0	1 (2.6%)		1 (0.6%)
		Normal value - n (%)	39 (48.8%)	21 (51.2%)	18 (46.2%)		78 (46.2%)
		High value - n (%)	0	0	1 (2.6%)		1 (0.6%)
		Abnormal clinically significant - n (%)	0	0	0		0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.14 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Hematocrit (%) 40-49.5	Change from baseline	n	39	21	20	0	80
		Mean (SD)	-1.00 (3.491)	-0.75 (1.941)	-0.04 (1.655)		-0.69 (2.763)
		Median	-0.50	-0.80	-0.50		-0.50
		Min, Max	-18.7, 5.5	-4.0, 4.2	-2.6, 4.1		-18.7, 5.5

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.5

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.14 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Leukocytes (G/L) 3.5-10	Baseline	n	80	41	39	9	169
		Mean (SD)	7.04 (1.592)	6.86 (1.417)	7.23 (1.412)	8.92 (2.548)	7.14 (1.621)
		Median	6.86	6.86	6.96	9.05	6.95
		Min, Max	3.7, 10.8	4.2, 9.6	5.1, 11.0	5.5, 13.2	3.7, 13.2
		Low value - n (%)	0	0	0	0	0
		Normal value - n (%)	78 (97.5%)	41 (100 %)	37 (94.9%)	7 (77.8%)	163 (96.4%)
		High value - n (%)	2 (2.5%)	0	2 (5.1%)	2 (22.2%)	6 (3.6%)
		Abnormal clinically significant - n (%)	0	0	0	2 (22.2%)	2 (1.2%)

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.5

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.14 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Leukocytes (G/L) 3.5-10	Day 6/ Discharge	n	79	41	39	0	159
		Mean (SD)	6.99 (1.789)	7.45 (1.211)	6.84 (1.263)		7.07 (1.545)
		Median	6.73	7.36	6.68		6.98
		Min, Max	4.2, 13.7	5.0, 10.3	4.6, 9.5		4.2, 13.7
		Low value - n (%)	0	0	0		0
		Normal value - n (%)	75 (93.8%)	40 (97.6%)	39 (100 %)		154 (91.1%)
		High value - n (%)	4 (5.0%)	1 (2.4%)	0		5 (3.0%)
		Abnormal clinically significant - n (%)	0	0	0		0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.5

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.14 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Leukocytes (G/L) 3.5-10	Change from baseline	n	79	41	39	0	159
		Mean (SD)	-0.02 (1.397)	0.59 (0.894)	-0.40 (1.179)		0.05 (1.276)
		Median	-0.02	0.58	-0.35		0.03
		Min, Max	-4.4, 6.8	-2.1, 3.0	-4.3, 1.9		-4.4, 6.8

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.5

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.14 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Neutrophils (G/L) 1.8-7.7	Baseline	n	80	41	39	9	169
		Mean (SD)	3.77 (1.206)	3.66 (1.066)	3.94 (1.051)	5.09 (2.265)	3.85 (1.242)
		Median	3.60	3.57	3.77	4.81	3.65
		Min, Max	1.5, 7.6	1.2, 6.0	2.0, 7.0	2.7, 9.5	1.2, 9.5
		Low value - n (%)	1 (1.3%)	1 (2.4%)	0	0	2 (1.2%)
		Normal value - n (%)	79 (98.8%)	40 (97.6%)	39 (100 %)	7 (77.8%)	165 (97.6%)
		High value - n (%)	0	0	0	2 (22.2%)	2 (1.2%)
		Abnormal clinically significant - n (%)	0	1 (2.4%)	0	1 (11.1%)	2 (1.2%)

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.14 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Neutrophils (G/L) 1.8-7.7	Day 6/ Discharge	n	79	41	39	0	159
		Mean (SD)	3.73 (1.362)	4.05 (0.980)	4.05 (0.920)		3.89 (1.179)
		Median	3.38	3.78	3.89		3.69
		Min, Max	1.6, 10.3	2.2, 6.1	2.4, 6.4		1.6, 10.3
		Low value - n (%)	1 (1.3%)	0	0		1 (0.6%)
		Normal value - n (%)	77 (96.3%)	41 (100 %)	39 (100 %)		157 (92.9%)
		High value - n (%)	1 (1.3%)	0	0		1 (0.6%)
		Abnormal clinically significant - n (%)	0	0	0		0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.14 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Neutrophils (G/L) 1.8-7.7	Change from baseline	n	79	41	39	0	159
		Mean (SD)	0.00 (1.171)	0.39 (0.659)	0.12 (1.056)		0.13 (1.040)
		Median	0.08	0.31	0.13		0.13
		Min, Max	-3.3, 6.3	-1.1, 2.3	-3.2, 1.9		-3.3, 6.3

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.5

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.14 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Neutrophils/leukocytes (%) 45-70	Baseline	n	80	41	39	9	169
		Mean (SD)	53.0 (8.39)	52.8 (7.93)	53.9 (6.21)	55.3 (9.57)	53.3 (7.85)
		Median	51.4	52.5	53.1	54.5	52.5
		Min, Max	40, 77	29, 66	39, 66	43, 72	29, 77
		Low value - n (%)	17 (21.3%)	5 (12.2%)	3 (7.7%)	1 (11.1%)	26 (15.4%)
		Normal value - n (%)	60 (75.0%)	36 (87.8%)	36 (92.3%)	7 (77.8%)	139 (82.2%)
		High value - n (%)	3 (3.8%)	0	0	1 (11.1%)	4 (2.4%)
		Abnormal clinically significant - n (%)	0	0	0	1 (11.1%)	1 (0.6%)

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.5

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_labhem.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.14 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Neutrophils/leukocytes (%) 45-70	Day 6/ Discharge	n	79	41	39	0	159
		Mean (SD)	52.6 (8.03)	53.9 (6.95)	59.0 (5.79)		54.5 (7.69)
		Median	52.5	52.6	59.6		54.5
		Min, Max	35, 75	40, 70	45, 70		35, 75
		Low value - n (%)	13 (16.3%)	3 (7.3%)	1 (2.6%)		17 (10.1%)
		Normal value - n (%)	63 (78.8%)	38 (92.7%)	38 (97.4%)		139 (82.2%)
		High value - n (%)	3 (3.8%)	0	0		3 (1.8%)
		Abnormal clinically significant - n (%)	0	0	0		0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.5

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**Table 15.2.6.14 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Neutrophils/leukocytes (%) 45-70	Change from baseline	n	79	41	39	0	159
		Mean (SD)	-0.2 (6.58)	1.2 (4.68)	5.1 (6.98)		1.5 (6.58)
		Median	1.0	-0.4	4.7		1.7
		Min, Max	-17, 18	-7, 11	-10, 16		-17, 18

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

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**Table 15.2.6.14 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Lymphocytes (G/L) 1-5	Baseline	n	80	41	39	9	169
		Mean (SD)	2.39 (0.704)	2.31 (0.555)	2.37 (0.600)	2.78 (0.743)	2.39 (0.651)
		Median	2.31	2.31	2.32	2.63	2.32
		Min, Max	1.2, 4.4	0.9, 3.3	1.4, 4.5	1.7, 4.1	0.9, 4.5
		Low value - n (%)	0	1 (2.4%)	0	0	1 (0.6%)
		Normal value - n (%)	80 (100 %)	40 (97.6%)	39 (100 %)	9 (100 %)	168 (99.4%)
		High value - n (%)	0	0	0	0	0
		Abnormal clinically significant - n (%)	1 (1.3%)	0	1 (2.6%)	1 (11.1%)	3 (1.8%)

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.5

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_labhem.sas

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**Table 15.2.6.14 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Lymphocytes (G/L) 1-5	Day 6/ Discharge	n	79	41	39	0	159
		Mean (SD)	2.35 (0.644)	2.40 (0.512)	1.91 (0.475)		2.26 (0.603)
		Median	2.24	2.55	1.82		2.18
		Min, Max	1.1, 4.2	1.4, 3.2	1.1, 3.4		1.1, 4.2
		Low value - n (%)	0	0	0		0
		Normal value - n (%)	79 (98.8%)	41 (100 %)	39 (100 %)		159 (94.1%)
		High value - n (%)	0	0	0		0
		Abnormal clinically significant - n (%)	0	0	0		0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.5

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_labhem.sas

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**Table 15.2.6.14 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Lymphocytes (G/L) 1-5	Change from baseline	n	79	41	39	0	159
		Mean (SD)	-0.04 (0.431)	0.09 (0.386)	-0.45 (0.366)		-0.11 (0.451)
		Median	-0.01	0.06	-0.46		-0.10
		Min, Max	-1.1, 1.0	-0.8, 0.8	-1.2, 0.3		-1.2, 1.0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

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**Table 15.2.6.14 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Lymphocytes/leukocytes (%) 20-45	Baseline	n	80	41	39	9	169
		Mean (SD)	34.3 (7.46)	34.1 (7.22)	33.1 (6.87)	32.5 (8.34)	33.8 (7.27)
		Median	35.5	34.8	32.9	34.9	34.8
		Min, Max	14, 49	22, 56	20, 49	18, 42	14, 56
		Low value - n (%)	2 (2.5%)	0	0	1 (11.1%)	3 (1.8%)
		Normal value - n (%)	72 (90.0%)	39 (95.1%)	37 (94.9%)	8 (88.9%)	156 (92.3%)
		High value - n (%)	6 (7.5%)	2 (4.9%)	2 (5.1%)	0	10 (5.9%)
		Abnormal clinically significant - n (%)	0	0	1 (2.6%)	1 (11.1%)	2 (1.2%)

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

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**Table 15.2.6.14 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Lymphocytes/leukocytes (%) 20-45	Day 6/ Discharge	n	79	41	39	0	159
		Mean (SD)	34.2 (6.93)	32.4 (6.36)	28.1 (5.38)		32.3 (6.86)
		Median	34.1	32.1	27.3		32.1
		Min, Max	14, 51	19, 45	18, 39		14, 51
		Low value - n (%)	4 (5.0%)	1 (2.4%)	1 (2.6%)		6 (3.6%)
		Normal value - n (%)	71 (88.8%)	39 (95.1%)	38 (97.4%)		148 (87.6%)
		High value - n (%)	4 (5.0%)	1 (2.4%)	0		5 (3.0%)
		Abnormal clinically significant - n (%)	0	0	0		0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

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**Table 15.2.6.14 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Lymphocytes/leukocytes (%) 20-45	Change from baseline	n	79	41	39	0	159
		Mean (SD)	-0.2 (5.56)	-1.6 (4.66)	-4.9 (6.63)		-1.7 (5.92)
		Median	-0.4	-0.8	-5.2		-1.6
		Min, Max	-16, 17	-14, 6	-16, 12		-16, 17

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

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**Table 15.2.6.14 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Monocytes (G/L) 0-0.8	Baseline	n	80	41	39	9	169
		Mean (SD)	0.63 (0.168)	0.61 (0.130)	0.66 (0.169)	0.77 (0.225)	0.64 (0.166)
		Median	0.61	0.60	0.65	0.77	0.62
		Min, Max	0.3, 1.2	0.4, 0.9	0.4, 1.2	0.5, 1.1	0.3, 1.2
		Low value - n (%)	0	0	0	0	0
		Normal value - n (%)	69 (86.3%)	38 (92.7%)	34 (87.2%)	6 (66.7%)	147 (87.0%)
		High value - n (%)	11 (13.8%)	3 (7.3%)	5 (12.8%)	3 (33.3%)	22 (13.0%)
		Abnormal clinically significant - n (%)	0	0	0	0	0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

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**Table 15.2.6.14 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Monocytes (G/L) 0-0.8	Day 6/ Discharge	n	79	41	39	0	159
		Mean (SD)	0.68 (0.173)	0.73 (0.178)	0.67 (0.204)		0.69 (0.182)
		Median	0.67	0.71	0.65		0.67
		Min, Max	0.3, 1.2	0.4, 1.3	0.4, 1.3		0.3, 1.3
		Low value - n (%)	0	0	0		0
		Normal value - n (%)	63 (78.8%)	30 (73.2%)	32 (82.1%)		125 (74.0%)
		High value - n (%)	16 (20.0%)	11 (26.8%)	7 (17.9%)		34 (20.1%)
		Abnormal clinically significant - n (%)	0	0	0		0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

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**Table 15.2.6.14 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Monocytes (G/L) 0-0.8	Change from baseline	n	79	41	39	0	159
		Mean (SD)	0.05 (0.142)	0.12 (0.156)	0.01 (0.194)		0.06 (0.163)
		Median	0.05	0.08	0.00		0.05
		Min, Max	-0.3, 0.5	-0.1, 0.7	-0.6, 0.6		-0.6, 0.7

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

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**Table 15.2.6.14 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Monocytes/leukocytes (%) 4-12	Baseline	n	80	41	39	9	169
		Mean (SD)	9.1 (1.84)	9.0 (1.57)	9.2 (1.79)	8.8 (2.11)	9.1 (1.77)
		Median	8.9	8.9	9.0	8.3	8.9
		Min, Max	6, 14	6, 12	6, 13	7, 14	6, 14
		Low value - n (%)	0	0	0	0	0
		Normal value - n (%)	73 (91.3%)	40 (97.6%)	36 (92.3%)	8 (88.9%)	157 (92.9%)
		High value - n (%)	7 (8.8%)	1 (2.4%)	3 (7.7%)	1 (11.1%)	12 (7.1%)
		Abnormal clinically significant - n (%)	0	0	0	0	0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

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**Table 15.2.6.14 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Monocytes/leukocytes (%) 4-12	Day 6/ Discharge	n	79	41	39	0	159
		Mean (SD)	9.9 (1.87)	9.8 (2.26)	9.8 (2.22)		9.9 (2.05)
		Median	9.6	9.9	9.5		9.6
		Min, Max	6, 16	7, 19	6, 17		6, 19
		Low value - n (%)	0	0	0		0
		Normal value - n (%)	70 (87.5%)	37 (90.2%)	33 (84.6%)		140 (82.8%)
		High value - n (%)	9 (11.3%)	4 (9.8%)	6 (15.4%)		19 (11.2%)
		Abnormal clinically significant - n (%)	0	0	0		0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

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**Table 15.2.6.14 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Monocytes/leukocytes (%) 4-12	Change from n baseline		79	41	39	0	159
		Mean (SD)	0.8 (1.49)	0.8 (2.01)	0.6 (1.85)		0.8 (1.72)
		Median	0.7	0.7	0.4		0.6
		Min, Max	-3, 4	-2, 11	-4, 5		-4, 11

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

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**Table 15.2.6.14 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Eosinophils (G/L) 0-0.45	Baseline	n	80	41	39	9	169
		Mean (SD)	0.23 (0.121)	0.25 (0.109)	0.24 (0.128)	0.25 (0.082)	0.24 (0.117)
		Median	0.20	0.25	0.21	0.26	0.22
		Min, Max	0.0, 0.9	0.1, 0.6	0.1, 0.7	0.1, 0.4	0.0, 0.9
		Low value - n (%)	0	0	0	0	0
		Normal value - n (%)	78 (97.5%)	39 (95.1%)	36 (92.3%)	9 (100 %)	162 (95.9%)
		High value - n (%)	2 (2.5%)	2 (4.9%)	3 (7.7%)	0	7 (4.1%)
		Abnormal clinically significant - n (%)	1 (1.3%)	0	0	0	1 (0.6%)

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

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**Table 15.2.6.14 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Eosinophils (G/L) 0-0.45	Day 6/ Discharge	n	79	41	39	0	159
		Mean (SD)	0.20 (0.108)	0.24 (0.098)	0.17 (0.101)		0.20 (0.106)
		Median	0.16	0.23	0.15		0.18
		Min, Max	0.0, 0.5	0.1, 0.6	0.0, 0.6		0.0, 0.6
		Low value - n (%)	0	0	0		0
		Normal value - n (%)	77 (96.3%)	40 (97.6%)	38 (97.4%)		155 (91.7%)
		High value - n (%)	2 (2.5%)	1 (2.4%)	1 (2.6%)		4 (2.4%)
		Abnormal clinically significant - n (%)	0	0	0		0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

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**Table 15.2.6.14 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Eosinophils (G/L) 0-0.45	Change from baseline	n	79	41	39	0	159
		Mean (SD)	-0.03 (0.089)	0.00 (0.046)	-0.07 (0.070)		-0.03 (0.079)
		Median	-0.03	0.00	-0.08		-0.03
		Min, Max	-0.5, 0.3	-0.1, 0.1	-0.3, 0.1		-0.5, 0.3

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.5

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_labhem.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.14 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Eosinophils/leukocytes (%) 0-5	Baseline	n	80	41	39	9	169
		Mean (SD)	3.3 (1.84)	3.7 (1.69)	3.5 (1.99)	3.0 (1.18)	3.4 (1.81)
		Median	3.0	3.3	3.0	2.5	3.1
		Min, Max	1, 12	2, 11	1, 10	1, 5	1, 12
		Low value - n (%)	0	0	0	0	0
		Normal value - n (%)	70 (87.5%)	36 (87.8%)	34 (87.2%)	8 (88.9%)	148 (87.6%)
		High value - n (%)	10 (12.5%)	5 (12.2%)	5 (12.8%)	1 (11.1%)	21 (12.4%)
		Abnormal clinically significant - n (%)	1 (1.3%)	0	0	0	1 (0.6%)

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.5

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_labhem.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.14 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Eosinophils/leukocytes (%) 0-5	Day 6/ Discharge	n	79	41	39	0	159
		Mean (SD)	2.9 (1.57)	3.3 (1.46)	2.6 (1.58)		2.9 (1.56)
		Median	2.6	3.0	2.5		2.6
		Min, Max	1, 10	2, 9	1, 10		1, 10
		Low value - n (%)	0	0	0		0
		Normal value - n (%)	74 (92.5%)	39 (95.1%)	37 (94.9%)		150 (88.8%)
		High value - n (%)	5 (6.3%)	2 (4.9%)	2 (5.1%)		9 (5.3%)
		Abnormal clinically significant - n (%)	0	0	0		0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.5

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_labhem.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.14 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Eosinophils/leukocytes (%) 0-5	Change from baseline	n	79	41	39	0	159
		Mean (SD)	-0.5 (1.11)	-0.3 (0.68)	-0.9 (1.09)		-0.5 (1.03)
		Median	-0.4	-0.2	-0.8		-0.5
		Min, Max	-6, 3	-2, 1	-5, 1		-6, 3

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.5

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_labhem.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.14 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Basophils (G/L) 0-0.2	Baseline	n	80	41	39	9	169
		Mean (SD)	0.02 (0.014)	0.03 (0.014)	0.03 (0.013)	0.04 (0.020)	0.03 (0.015)
		Median	0.02	0.03	0.02	0.03	0.02
		Min, Max	0.0, 0.1	0.0, 0.1	0.0, 0.1	0.0, 0.1	0.0, 0.1
		Low value - n (%)	0	0	0	0	0
		Normal value - n (%)	80 (100 %)	41 (100 %)	39 (100 %)	9 (100 %)	169 (100 %)
		High value - n (%)	0	0	0	0	0
		Abnormal clinically significant - n (%)	0	0	0	0	0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.5

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_labhem.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.14 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Basophils (G/L) 0-0.2	Day 6/ Discharge	n	79	41	39	0	159
		Mean (SD)	0.03 (0.018)	0.03 (0.013)	0.03 (0.017)		0.03 (0.016)
		Median	0.02	0.03	0.03		0.03
		Min, Max	0.0, 0.1	0.0, 0.1	0.0, 0.1		0.0, 0.1
		Low value - n (%)	0	0	0		0
		Normal value - n (%)	79 (98.8%)	41 (100 %)	39 (100 %)		159 (94.1%)
		High value - n (%)	0	0	0		0
		Abnormal clinically significant - n (%)	0	0	0		0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.5

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_labhem.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.14 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Basophils (G/L) 0-0.2	Change from baseline	n	79	41	39	0	159
		Mean (SD)	0.00 (0.012)	0.00 (0.010)	0.01 (0.012)		0.00 (0.012)
		Median	0.00	0.00	0.01		0.00
		Min, Max	0.0, 0.0	0.0, 0.0	0.0, 0.0		0.0, 0.0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.5

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_labhem.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.14 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Basophils/leukocytes (%)	Baseline	n	80	41	39	9	169
0-1		Mean (SD)	0.4 (0.20)	0.5 (0.25)	0.4 (0.20)	0.5 (0.29)	0.4 (0.22)
		Median	0.3	0.4	0.3	0.4	0.3
		Min, Max	0, 1	0, 1	0, 1	0, 1	0, 1
		Low value - n (%)	0	0	0	0	0
		Normal value - n (%)	80 (100 %)	40 (97.6%)	39 (100 %)	8 (88.9%)	167 (98.8%)
		High value - n (%)	0	1 (2.4%)	0	1 (11.1%)	2 (1.2%)
		Abnormal clinically significant - n (%)	0	0	0	0	0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.5

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_labhem.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.14 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Basophils/leukocytes (%) 0-1	Day 6/ Discharge	n	79	41	39	0	159
		Mean (SD)	0.4 (0.25)	0.4 (0.21)	0.5 (0.23)		0.4 (0.24)
		Median	0.4	0.4	0.4		0.4
		Min, Max	0, 1	0, 1	0, 1		0, 1
		Low value - n (%)	0	0	0		0
		Normal value - n (%)	76 (95.0%)	41 (100 %)	38 (97.4%)		155 (91.7%)
		High value - n (%)	3 (3.8%)	0	1 (2.6%)		4 (2.4%)
		Abnormal clinically significant - n (%)	0	0	0		0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.5

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.14 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Basophils/leukocytes (%)	Change from baseline	n	79	41	39	0	159
0-1		Mean (SD)	0.1 (0.18)	0.0 (0.16)	0.1 (0.18)		0.0 (0.18)
		Median	0.1	0.0	0.1		0.0
		Min, Max	-1, 1	0, 0	0, 1		-1, 1

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.14 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Platelets (G/L) 125-400	Baseline	n	80	41	39	9	169
		Mean (SD)	223.3 (48.97)	206.9 (34.24)	207.8 (45.92)	262.2 (46.78)	217.8 (46.48)
		Median	223.0	204.0	198.0	272.0	215.0
		Min, Max	128, 333	139, 272	123, 333	164, 323	123, 333
		Low value - n (%)	0	0	1 (2.6%)	0	1 (0.6%)
		Normal value - n (%)	80 (100 %)	41 (100 %)	38 (97.4%)	9 (100 %)	168 (99.4%)
		High value - n (%)	0	0	0	0	0
		Abnormal clinically significant - n (%)	0	0	0	0	0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.5

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.14 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Platelets (G/L) 125-400	Day 6/ Discharge	n	79	41	39	0	159
		Mean (SD)	233.2 (47.49)	223.9 (39.97)	217.7 (47.51)		227.0 (45.87)
		Median	230.0	227.0	214.0		224.0
		Min, Max	140, 344	144, 316	129, 349		129, 349
		Low value - n (%)	0	0	0		0
		Normal value - n (%)	79 (98.8%)	41 (100 %)	39 (100 %)		159 (94.1%)
		High value - n (%)	0	0	0		0
		Abnormal clinically significant - n (%)	0	0	0		0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.14 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Platelets (G/L) 125-400	Change from baseline	n	79	41	39	0	159
		Mean (SD)	9.8 (23.07)	17.0 (23.57)	9.9 (19.03)		11.7 (22.36)
		Median	8.0	11.0	9.0		10.0
		Min, Max	-39, 96	-23, 70	-26, 53		-39, 96

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.14 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Ery. mean corpuscular hemoglobin (pg) 28-34	Baseline	n	80	41	39	9	169
		Mean (SD)	31.50 (1.707)	31.53 (1.152)	31.37 (1.619)	33.02 (1.519)	31.56 (1.586)
		Median	31.45	31.60	31.60	32.90	31.60
		Min, Max	27.6, 38.7	28.9, 35.3	28.4, 34.3	30.7, 36.1	27.6, 38.7
		Low value - n (%)	1 (1.3%)	0	0	0	1 (0.6%)
		Normal value - n (%)	76 (95.0%)	40 (97.6%)	37 (94.9%)	8 (88.9%)	161 (95.3%)
		High value - n (%)	3 (3.8%)	1 (2.4%)	2 (5.1%)	1 (11.1%)	7 (4.1%)
		Abnormal clinically significant - n (%)	0	0	0	0	0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.14 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Ery. mean corpuscular hemoglobin (pg) 28-34	Day 6/ Discharge	n	79	41	39	0	159
		Mean (SD)	31.34 (1.413)	31.41 (1.118)	31.41 (1.711)		31.38 (1.417)
		Median	31.30	31.50	31.50		31.40
		Min, Max	27.9, 37.2	28.8, 34.6	28.2, 34.4		27.9, 37.2
		Low value - n (%)	1 (1.3%)	0	0		1 (0.6%)
		Normal value - n (%)	77 (96.3%)	40 (97.6%)	37 (94.9%)		154 (91.1%)
		High value - n (%)	1 (1.3%)	1 (2.4%)	2 (5.1%)		4 (2.4%)
		Abnormal clinically significant - n (%)	0	0	0		0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.14 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Ery. mean corpuscular hemoglobin (pg) 28-34	Change from baseline	n	79	41	39	0	159
		Mean (SD)	-0.17 (0.841)	-0.12 (0.379)	0.04 (0.420)		-0.10 (0.660)
		Median	-0.10	-0.20	0.00		-0.10
		Min, Max	-6.5, 1.2	-0.8, 0.7	-0.8, 0.9		-6.5, 1.2

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.14 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Ery. mean corpuscular hgb concentration (g/dL) 31-35	Baseline	n	41	20	19	4	84
		Mean (SD)	34.56 (0.866)	34.51 (0.662)	34.46 (0.673)	34.68 (0.768)	34.53 (0.764)
		Median	34.60	34.45	34.50	34.75	34.50
		Min, Max	33.1, 36.8	33.1, 36.0	33.2, 35.7	33.7, 35.5	33.1, 36.8
		Low value - n (%)	0	0	0	0	0
		Normal value - n (%)	31 (38.8%)	18 (43.9%)	16 (41.0%)	3 (33.3%)	68 (40.2%)
		High value - n (%)	10 (12.5%)	2 (4.9%)	3 (7.7%)	1 (11.1%)	16 (9.5%)
		Abnormal clinically significant - n (%)	0	0	0	0	0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.14 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Ery. mean corpuscular hgb concentration (g/dL) 31-35	Day 6/ Discharge	n	40	20	19	0	79
		Mean (SD)	34.78 (0.780)	34.50 (0.882)	34.79 (0.744)		34.71 (0.799)
		Median	34.70	34.40	35.00		34.70
		Min, Max	33.3, 36.7	33.2, 36.8	33.6, 35.9		33.2, 36.8
		Low value - n (%)	0	0	0		0
		Normal value - n (%)	27 (33.8%)	16 (39.0%)	12 (30.8%)		55 (32.5%)
		High value - n (%)	13 (16.3%)	4 (9.8%)	7 (17.9%)		24 (14.2%)
		Abnormal clinically significant - n (%)	0	0	0		0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.5

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_labhem.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.14 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Ery. mean corpuscular hgb concentration (g/dL) 31-35	Change from baseline	n	40	20	19	0	79
		Mean (SD)	0.21 (0.502)	-0.02 (0.387)	0.33 (0.558)		0.18 (0.500)
		Median	0.10	0.00	0.30		0.10
		Min, Max	-0.7, 1.3	-0.8, 0.8	-0.9, 1.1		-0.9, 1.3

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.5

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_labhem.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.14 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Ery. mean corpuscular hgb concentration (g/dL) 32-36	Baseline	n	39	21	20	5	85
		Mean (SD)	34.96 (0.749)	35.00 (0.767)	35.34 (0.840)	35.30 (0.887)	35.08 (0.786)
		Median	34.80	35.00	35.55	35.70	35.10
		Min, Max	33.6, 36.5	33.3, 36.9	33.7, 36.9	33.8, 35.9	33.3, 36.9
		Low value - n (%)	0	0	0	0	0
		Normal value - n (%)	36 (45.0%)	20 (48.8%)	17 (43.6%)	5 (55.6%)	78 (46.2%)
		High value - n (%)	3 (3.8%)	1 (2.4%)	3 (7.7%)	0	7 (4.1%)
		Abnormal clinically significant - n (%)	0	0	0	0	0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.5

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_labhem.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.14 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Ery. mean corpuscular hgb concentration (g/dL) 32-36	Day 6/ Discharge	n	39	21	20	0	80
		Mean (SD)	35.06 (0.925)	35.26 (0.822)	35.56 (0.849)		35.24 (0.893)
		Median	35.20	35.40	35.75		35.20
		Min, Max	33.0, 37.2	33.4, 36.4	33.8, 36.7		33.0, 37.2
		Low value - n (%)	0	0	0		0
		Normal value - n (%)	34 (42.5%)	18 (43.9%)	13 (33.3%)		65 (38.5%)
		High value - n (%)	5 (6.3%)	3 (7.3%)	7 (17.9%)		15 (8.9%)
		Abnormal clinically significant - n (%)	0	0	0		0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.5

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_labhem.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.14 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Ery. mean corpuscular hgb concentration (g/dL) 32-36	Change from baseline	n	39	21	20	0	80
		Mean (SD)	0.09 (0.687)	0.26 (0.505)	0.22 (0.526)		0.17 (0.603)
		Median	0.20	0.20	0.30		0.20
		Min, Max	-1.7, 1.5	-0.9, 1.0	-0.7, 1.5		-1.7, 1.5

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.5

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_labhem.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.14 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Ery. mean corpuscular volume (fL) 83-103	Baseline	n	80	41	39	9	169
		Mean (SD)	90.6 (4.63)	90.8 (3.87)	89.9 (4.19)	94.4 (5.21)	90.7 (4.45)
		Median	90.2	90.8	89.4	95.3	90.5
		Min, Max	83, 110	84, 103	83, 101	86, 102	83, 110
		Low value - n (%)	3 (3.8%)	0	1 (2.6%)	0	4 (2.4%)
		Normal value - n (%)	75 (93.8%)	40 (97.6%)	38 (97.4%)	9 (100 %)	162 (95.9%)
		High value - n (%)	2 (2.5%)	1 (2.4%)	0	0	3 (1.8%)
		Abnormal clinically significant - n (%)	0	0	0	0	0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.5

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_labhem.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.14 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Ery. mean corpuscular volume (fL) 83-103	Day 6/ Discharge	n	79	41	39	0	159
		Mean (SD)	89.8 (4.26)	90.1 (3.77)	89.3 (4.18)		89.7 (4.10)
		Median	89.5	90.1	89.1		89.6
		Min, Max	82, 107	84, 102	81, 99		81, 107
		Low value - n (%)	3 (3.8%)	0	1 (2.6%)		4 (2.4%)
		Normal value - n (%)	75 (93.8%)	41 (100 %)	38 (97.4%)		154 (91.1%)
		High value - n (%)	1 (1.3%)	0	0		1 (0.6%)
		Abnormal clinically significant - n (%)	0	0	0		0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.5

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_labhem.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.14 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Ery. mean corpuscular volume (fL) 83-103	Change from n baseline		79	41	39	0	159
		Mean (SD)	-0.8 (1.79)	-0.7 (0.60)	-0.6 (0.89)		-0.7 (1.37)
		Median	-0.6	-0.7	-0.5		-0.6
		Min, Max	-15, 1	-2, 0	-3, 1		-15, 1

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.5

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_labhem.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.15 Summary of Urinalysis Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
pH 5-8	Baseline	n	80	41	39	9	169
		Mean (SD)	5.88 (0.475)	5.99 (0.531)	5.91 (0.525)	6.00 (0.500)	5.92 (0.500)
		Median	6.00	6.00	6.00	6.00	6.00
		Min, Max	5.0, 6.5	5.0, 7.0	5.0, 7.0	5.0, 7.0	5.0, 7.0
		Low value - n (%)	0	0	0	0	0
		Normal value - n (%)	80 (100 %)	41 (100 %)	39 (100 %)	9 (100 %)	169 (100 %)
		High value - n (%)	0	0	0	0	0
		Abnormal clinically significant - n (%)	0	0	0	0	0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.6

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_laburin.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.15 Summary of Urinalysis Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
pH 5-8	Day 6/ Discharge	n	79	41	39	0	159
		Mean (SD)	5.99 (0.631)	5.98 (0.671)	6.01 (0.568)		5.99 (0.623)
		Median	6.00	6.00	6.00		6.00
		Min, Max	5.0, 8.0	5.0, 7.0	5.0, 7.0		5.0, 8.0
		Low value - n (%)	0	0	0	0	0
		Normal value - n (%)	79 (98.8%)	41 (100 %)	39 (100 %)	0	159 (94.1%)
		High value - n (%)	0	0	0	0	0
		Abnormal clinically significant - n (%)	0	0	0	0	0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.6

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_laburin.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.15 Summary of Urinalysis Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
pH 5-8	Change from baseline	n	79	41	39	0	159
		Mean (SD)	0.10 (0.691)	-0.01 (0.712)	0.10 (0.609)		0.07 (0.675)
		Median	0.00	0.00	0.00		0.00
		Min, Max	-1.5, 1.5	-2.0, 1.5	-1.5, 1.0		-2.0, 1.5

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.6

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_laburin.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.15 Summary of Urinalysis Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Specific gravity (g/L) 1010-1030	Baseline	n	80	41	39	9	169
		Mean (SD)	1018.4 (3.96)	1017.7 (4.20)	1018.8 (4.52)	1018.3 (3.54)	1018.3 (4.12)
		Median	1020.0	1020.0	1020.0	1020.0	1020.0
		Min, Max	1010, 1025	1010, 1025	1010, 1030	1015, 1025	1010, 1030
		Low value - n (%)	0	0	0	0	0
		Normal value - n (%)	80 (100 %)	41 (100 %)	39 (100 %)	9 (100 %)	169 (100 %)
		High value - n (%)	0	0	0	0	0
		Abnormal clinically significant - n (%)	0	0	0	0	0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.6

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_laburin.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.15 Summary of Urinalysis Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Specific gravity (g/L) 1010-1030	Day 6/ Discharge	n	79	41	39	0	159
		Mean (SD)	1016.9 (4.69)	1015.6 (4.90)	1016.5 (5.15)		1016.5 (4.86)
		Median	1015.0	1015.0	1015.0		1015.0
		Min, Max	1005, 1025	1005, 1025	1000, 1025		1000, 1025
		Low value - n (%)	2 (2.5%)	3 (7.3%)	3 (7.7%)	0	8 (4.7%)
		Normal value - n (%)	77 (96.3%)	38 (92.7%)	36 (92.3%)	0	151 (89.3%)
		High value - n (%)	0	0	0	0	0
		Abnormal clinically significant - n (%)	0	0	0	0	0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.6

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.15 Summary of Urinalysis Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Specific gravity (g/L) 1010-1030	Change from baseline	n	79	41	39	0	159
		Mean (SD)	-1.5 (4.82)	-2.1 (6.02)	-2.3 (5.61)		-1.8 (5.32)
		Median	0.0	0.0	0.0		0.0
		Min, Max	-15, 10	-15, 10	-15, 10		-15, 10

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.15 Summary of Urinalysis Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Bilirubin (urine) (mg/dL) Negative	Baseline	Negative - n (%)	80 (100 %)	41 (100 %)	39 (100 %)	9 (100 %)	169 (100 %)
		Positive - n (%)	0	0	0	0	0
		Normal value - n (%)	80 (100 %)	41 (100 %)	39 (100 %)	9 (100 %)	169 (100 %)
		Abnormal value - n (%)	0	0	0	0	0
		Abnormal clinically significant - n (%)	0	0	0	0	0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.15 Summary of Urinalysis Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Bilirubin (urine) (mg/dL) Negative	Day 6/ Discharge	Negative - n (%)	79 (98.8%)	41 (100 %)	39 (100 %)	0	159 (94.1%)
		Positive - n (%)	0	0	0	0	0
		Normal value - n (%)	79 (98.8%)	41 (100 %)	39 (100 %)	0	159 (94.1%)
		Abnormal value - n (%)	0	0	0	0	0
		Abnormal clinically significant - n (%)	0	0	0	0	0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

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Path: /cvn/projects/prj/development/000000106324/dev/tables/t_laburin.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.15 Summary of Urinalysis Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Glucose (urine) (mg/dL) Negative	Baseline	Negative - n (%)	80 (100 %)	41 (100 %)	39 (100 %)	9 (100 %)	169 (100 %)
		Positive - n (%)	0	0	0	0	0
		Normal value - n (%)	80 (100 %)	41 (100 %)	39 (100 %)	9 (100 %)	169 (100 %)
		Abnormal value - n (%)	0	0	0	0	0
		Abnormal clinically significant - n (%)	0	0	0	0	0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.15 Summary of Urinalysis Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Glucose (urine) (mg/dL) Negative	Day 6/ Discharge	Negative - n (%)	79 (98.8%)	41 (100 %)	39 (100 %)	0	159 (94.1%)
		Positive - n (%)	0	0	0	0	0
		Normal value - n (%)	79 (98.8%)	41 (100 %)	39 (100 %)	0	159 (94.1%)
		Abnormal value - n (%)	0	0	0	0	0
		Abnormal clinically significant - n (%)	0	0	0	0	0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.15 Summary of Urinalysis Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Nitrite Negative	Baseline	Negative - n (%)	78 (97.5%)	41 (100 %)	39 (100 %)	9 (100 %)	167 (98.8%)
		Positive - n (%)	2 (2.5%)	0	0	0	2 (1.2%)
		Normal value - n (%)	78 (97.5%)	41 (100 %)	39 (100 %)	9 (100 %)	167 (98.8%)
		Abnormal value - n (%)	2 (2.5%)	0	0	0	2 (1.2%)
		Abnormal clinically significant - n (%)	0	0	0	0	0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.6

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_laburin.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.15 Summary of Urinalysis Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Nitrite Negative	Day 6/ Discharge	Negative - n (%)	78 (97.5%)	40 (97.6%)	39 (100 %)	0	157 (92.9%)
		Positive - n (%)	1 (1.3%)	1 (2.4%)	0	0	2 (1.2%)
		Normal value - n (%)	78 (97.5%)	40 (97.6%)	39 (100 %)	0	157 (92.9%)
		Abnormal value - n (%)	1 (1.3%)	1 (2.4%)	0	0	2 (1.2%)
		Abnormal clinically significant - n (%)	0	0	0	0	0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.6

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_laburin.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.15 Summary of Urinalysis Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Protein (urine) (mg/dL) Negative	Baseline	Negative - n (%)	80 (100 %)	41 (100 %)	39 (100 %)	9 (100 %)	169 (100 %)
		Positive - n (%)	0	0	0	0	0
		Normal value - n (%)	80 (100 %)	41 (100 %)	39 (100 %)	9 (100 %)	169 (100 %)
		Abnormal value - n (%)	0	0	0	0	0
		Abnormal clinically significant - n (%)	0	0	0	0	0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: If result is greater than 'Negative/Trace' then result is counted as 'Positive'.

Appendix 15.3.6.6

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_laburin.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.15 Summary of Urinalysis Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Protein (urine) (mg/dL) Negative	Day 6/ Discharge	n	2	1	0	0	3
		Mean (SD)	NC	NC			NC
		Median	NC	NC			NC
		Min, Max	NC	NC			NC
		Negative - n (%)	77 (96.3%)	40 (97.6%)	39 (100 %)	0	156 (92.3%)
		Positive - n (%)	2 (2.5%)	1 (2.4%)	0	0	3 (1.8%)
		Normal value - n (%)	77 (96.3%)	40 (97.6%)	39 (100 %)	0	156 (92.3%)
		Abnormal value - n (%)	2 (2.5%)	1 (2.4%)	0	0	3 (1.8%)
		Abnormal clinically significant - n (%)	0	0	0	0	0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: If result is greater than 'Negative/Trace' then result is counted as 'Positive'.

Note: NC = Not calculated.

Appendix 15.3.6.6

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_laburin.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.15 Summary of Urinalysis Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Protein (urine) (mg/dL) Negative	Change from baseline	n	0	0	0	0	0
		Mean (SD)	NC	NC			NC
		Median	NC	NC			NC
		Min, Max	NC	NC			NC

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: If result is greater than 'Negative/Trace' then result is counted as 'Positive'.

Note: NC = Not calculated.

Appendix 15.3.6.6

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_laburin.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.15 Summary of Urinalysis Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Occult blood (ERYT/uL) Negative	Baseline	n	6	1	2	1	10
		Mean (SD)	89.2 (92.76)	NC	NC	NC	88.5 (93.49)
		Median	50.0	NC	NC	NC	50.0
		Min, Max	10, 250	NC	NC	NC	10, 250
		Negative - n (%)	74 (92.5%)	40 (97.6%)	37 (94.9%)	8 (88.9%)	159 (94.1%)
		Positive - n (%)	6 (7.5%)	1 (2.4%)	2 (5.1%)	1 (11.1%)	10 (5.9%)
		Normal value - n (%)	74 (92.5%)	40 (97.6%)	37 (94.9%)	8 (88.9%)	159 (94.1%)
		Abnormal value - n (%)	6 (7.5%)	1 (2.4%)	2 (5.1%)	1 (11.1%)	10 (5.9%)
		Abnormal clinically significant - n (%)	0	0	0	0	0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: If result is greater than 'Negative/Trace' then result is counted as 'Positive'.

Note: NC = Not calculated.

Appendix 15.3.6.6

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_laburin.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.15 Summary of Urinalysis Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Occult blood (ERYT/uL) Negative	Day 6/ Discharge	n	8	2	2	0	12
		Mean (SD)	123.1 (113.83)	NC	NC		127.9 (113.57)
		Median	87.5	NC	NC		87.5
		Min, Max	10, 250	NC	NC		10, 250
		Negative - n (%)	71 (88.8%)	39 (95.1%)	37 (94.9%)	0	147 (87.0%)
		Positive - n (%)	8 (10.0%)	2 (4.9%)	2 (5.1%)	0	12 (7.1%)
		Normal value - n (%)	71 (88.8%)	39 (95.1%)	37 (94.9%)	0	147 (87.0%)
		Abnormal value - n (%)	8 (10.0%)	2 (4.9%)	2 (5.1%)	0	12 (7.1%)
		Abnormal clinically significant - n (%)	0	0	0	0	0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: If result is greater than 'Negative/Trace' then result is counted as 'Positive'.

Note: NC = Not calculated.

Appendix 15.3.6.6

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_laburin.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.15 Summary of Urinalysis Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Occult blood (ERYT/uL) Negative	Change from baseline	n	1	0	0	0	1
		Mean (SD)	NC	NC	NC		NC
		Median	NC	NC	NC		NC
		Min, Max	NC	NC	NC		NC

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: If result is greater than 'Negative/Trace' then result is counted as 'Positive'.

Note: NC = Not calculated.

Appendix 15.3.6.6

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_laburin.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.16 Summary of Supine Vital Signs - Safety Population**

Parameter (units)	Study Day	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)	
			Raw value	Change	Raw value	Change	Raw value	Change
Supine Systolic Blood Pressure (mmHg)		n	80		41		39	
		Mean (SD)	111.4 (13.03)		110.4 (13.71)		112.1 (12.86)	
		Median	111.0		111.0		114.0	
		Min, Max	74, 146		80, 141		91, 143	
	Day 1	n	80	80	41	41	39	39
		Mean (SD)	111.0 (12.48)	-0.5 (10.06)	110.3 (12.72)	-0.1 (9.65)	110.5 (11.25)	-1.6 (8.70)
		Median	110.0	-0.5	111.0	-2.0	111.0	-3.0
		Min, Max	90, 144	-22, 34	86, 134	-19, 28	88, 135	-17, 21
	Day 2	n	80	80	41	41	39	39
		Mean (SD)	110.5 (13.06)	-1.0 (9.21)	112.0 (12.08)	1.6 (6.93)	110.6 (11.50)	-1.4 (8.83)
		Median	110.0	-1.5	113.0	2.0	110.0	-1.0
		Min, Max	73, 151	-33, 21	89, 138	-11, 15	88, 146	-18, 14

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects who were enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Change is change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.7

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_vit.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.16 Summary of Supine Vital Signs - Safety Population**

Parameter (units)	Study Day	Statistic	Enrolled Not Randomized (N=9)		Overall Safety (N=169)	
			Raw value	Change	Raw value	Change
Supine Systolic Blood Pressure (mmHg)	Baseline	n	9		169	
		Mean (SD)	121.3 (15.96)		111.8 (13.40)	
		Median	118.0		112.0	
		Min, Max	101, 155		74, 155	
	Day 1	n	0	0	160	160
		Mean (SD)			110.7 (12.18)	-0.7 (9.60)
		Median			110.5	-2.0
		Min, Max			86, 144	-22, 34
	Day 2	n	0	0	160	160
		Mean (SD)			110.9 (12.39)	-0.4 (8.62)
		Median			110.5	-1.0
		Min, Max			73, 151	-33, 21

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects who were enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Change is change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.7

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_vit.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.16 Summary of Supine Vital Signs - Safety Population**

Parameter (units)	Study Day	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)	
			Raw value	Change	Raw value	Change	Raw value	Change
Supine Systolic Blood Pressure (mmHg)	Day 3	n	80	80	41	41	39	39
		Mean (SD)	112.7 (12.39)	1.3 (9.55)	111.3 (11.20)	1.0 (8.89)	111.9 (12.30)	-0.2 (8.21)
		Median	110.5	2.0	113.0	1.0	110.0	-1.0
		Min, Max	91, 170	-25, 24	90, 144	-18, 18	90, 142	-16, 16
	Day 4	n	79	79	41	41	39	39
		Mean (SD)	111.0 (11.73)	-0.6 (10.69)	110.3 (13.48)	-0.1 (10.75)	112.7 (13.88)	0.7 (9.15)
		Median	111.0	0.0	110.0	-1.0	113.0	1.0
		Min, Max	90, 150	-26, 37	84, 154	-17, 28	87, 151	-16, 25
	Day 5	n	79	79	41	41	39	39
		Mean (SD)	108.4 (12.16)	-3.2 (9.15)	109.5 (9.96)	-0.8 (8.02)	110.6 (12.56)	-1.5 (10.01)
		Median	108.0	-3.0	111.0	-1.0	111.0	-2.0
		Min, Max	82, 138	-27, 18	89, 132	-20, 18	85, 138	-20, 18

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects who were enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Change is change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.7

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_vit.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.16 Summary of Supine Vital Signs - Safety Population**

Parameter (units)	Study Day	Statistic	Enrolled Not Randomized (N=9)		Overall Safety (N=169)	
			Raw value	Change	Raw value	Change
Supine Systolic Blood Pressure (mmHg)	Day 3	n	0	0	160	160
		Mean (SD)			112.2 (12.01)	0.8 (9.04)
		Median			110.5	1.0
		Min, Max			90, 170	-25, 24
	Day 4	n	0	0	159	159
		Mean (SD)			111.2 (12.69)	-0.2 (10.30)
		Median			111.0	0.0
		Min, Max			84, 154	-26, 37
	Day 5	n	0	0	159	159
		Mean (SD)			109.2 (11.70)	-2.2 (9.10)
		Median			110.0	-2.0
		Min, Max			82, 138	-27, 18

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects who were enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Change is change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.7

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_vit.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.16 Summary of Supine Vital Signs - Safety Population**

Parameter (units)	Study Day	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)	
			Raw value	Change	Raw value	Change	Raw value	Change
Supine Systolic Blood Pressure (mmHg)	Day 6/Discharge	n	80	80	41	41	39	39
		Mean (SD)	107.2 (10.55)	-4.2 (9.29)	107.7 (10.96)	-2.7 (9.64)	111.4 (12.26)	-0.6 (9.24)
		Median	109.0	-4.0	106.0	0.0	111.0	-3.0
		Min, Max	90, 140	-30, 18	90, 135	-21, 16	90, 140	-20, 18
Supine Diastolic Blood Pressure (mmHg)	Baseline	n	80		41		39	
		Mean (SD)	69.4 (8.31)		67.6 (8.11)		71.3 (8.01)	
		Median	69.0		67.0		71.0	
		Min, Max	44, 90		51, 89		56, 90	
	Day 1	n	80	80	41	41	39	39
		Mean (SD)	70.0 (7.97)	0.6 (7.59)	68.7 (7.29)	1.1 (7.23)	67.9 (6.32)	-3.4 (6.75)
		Median	70.0	0.0	68.0	0.0	67.0	-3.0
		Min, Max	52, 89	-14, 21	53, 84	-17, 16	52, 82	-17, 10

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects who were enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Change is change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.7

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_vit.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.16 Summary of Supine Vital Signs - Safety Population**

Parameter (units)	Study Day	Statistic	Enrolled Not Randomized (N=9)		Overall Safety (N=169)	
			Raw value	Change	Raw value	Change
Supine Systolic Blood Pressure (mmHg)	Day 6/Discharge	n	2	2	162	162
		Mean (SD)	NC	NC	108.5 (11.21)	-2.8 (9.46)
		Median	NC	NC	108.0	-3.0
		Min, Max	NC	NC	90, 140	-30, 18
Supine Diastolic Blood Pressure (mmHg)	Baseline	n	9		169	
		Mean (SD)	79.8 (10.55)		70.0 (8.66)	
		Median	79.0		69.0	
		Min, Max	68, 101		44, 101	
	Day 1	n	0	0	160	160
		Mean (SD)			69.2 (7.44)	-0.3 (7.48)
		Median			68.0	-1.0
		Min, Max			52, 89	-17, 21

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects who were enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Change is change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: NC = Not calculated

Appendix 15.3.6.7

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_vit.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.16 Summary of Supine Vital Signs - Safety Population**

Parameter (units)	Study Day	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)	
			Raw value	Change	Raw value	Change	Raw value	Change
Supine Diastolic Blood Pressure (mmHg)	Day 2	n	80	80	41	41	39	39
		Mean (SD)	69.5 (7.83)	0.1 (7.39)	69.3 (7.92)	1.7 (9.16)	68.2 (6.68)	-3.2 (8.01)
		Median	69.0	0.0	69.0	2.0	68.0	-2.0
		Min, Max	48, 88	-16, 24	53, 87	-14, 23	52, 85	-18, 13
	Day 3	n	80	80	41	41	39	39
		Mean (SD)	69.7 (6.93)	0.3 (7.46)	68.3 (6.90)	0.7 (7.45)	68.4 (6.52)	-3.0 (8.29)
		Median	70.0	-1.0	68.0	1.0	67.0	-3.0
		Min, Max	55, 93	-17, 21	55, 85	-14, 16	49, 81	-18, 13
	Day 4	n	79	79	41	41	39	39
		Mean (SD)	70.1 (7.81)	0.6 (7.67)	67.5 (8.36)	-0.1 (8.84)	68.3 (7.18)	-3.0 (7.43)
		Median	69.0	-1.0	67.0	-2.0	68.0	-2.0
		Min, Max	56, 94	-14, 20	56, 98	-17, 20	54, 85	-17, 13

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects who were enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Change is change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.7

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_vit.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.16 Summary of Supine Vital Signs - Safety Population**

Parameter (units)	Study Day	Statistic	Enrolled Not Randomized (N=9)		Overall Safety (N=169)	
			Raw value	Change	Raw value	Change
Supine Diastolic Blood Pressure (mmHg)	Day 2	n	0	0	160	160
		Mean (SD)			69.1 (7.57)	-0.3 (8.16)
		Median			69.0	0.0
		Min, Max			48, 88	-18, 24
	Day 3	n	0	0	160	160
		Mean (SD)			69.0 (6.82)	-0.4 (7.76)
		Median			69.0	-1.0
		Min, Max			49, 93	-18, 21
	Day 4	n	0	0	159	159
		Mean (SD)			69.0 (7.84)	-0.5 (8.02)
		Median			68.0	-1.0
		Min, Max			54, 98	-17, 20

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects who were enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Change is change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.7

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_vit.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.16 Summary of Supine Vital Signs - Safety Population**

Parameter (units)	Study Day	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)	
			Raw value	Change	Raw value	Change	Raw value	Change
Supine Diastolic Blood Pressure (mmHg)	Day 5	n	79	79	41	41	39	39
		Mean (SD)	68.6 (6.47)	-0.9 (7.08)	69.1 (6.26)	1.5 (7.72)	68.8 (6.49)	-2.6 (8.21)
		Median	69.0	-2.0	68.0	2.0	68.0	-4.0
		Min, Max	54, 83	-16, 19	60, 82	-13, 19	59, 83	-20, 17
	Day 6/Discharge	n	80	80	41	41	39	39
		Mean (SD)	71.8 (7.02)	2.3 (7.50)	70.7 (7.54)	3.1 (8.55)	71.4 (7.19)	0.1 (8.36)
		Median	72.0	3.0	70.0	6.0	70.0	-1.0
		Min, Max	55, 88	-16, 21	56, 90	-17, 20	54, 87	-19, 16
	Baseline	n	80		41		39	
		Mean (SD)	74.4 (10.15)		75.0 (9.31)		75.4 (9.47)	
		Median	74.5		76.0		74.0	
		Min, Max	50, 102		57, 99		59, 91	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects who were enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Change is change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.7

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_vit.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.16 Summary of Supine Vital Signs - Safety Population**

Parameter (units)	Study Day	Statistic	Enrolled Not Randomized (N=9)		Overall Safety (N=169)	
			Raw value	Change	Raw value	Change
Supine Diastolic Blood Pressure (mmHg)	Day 5	n	0	0	159	159
		Mean (SD)			68.8 (6.38)	-0.7 (7.62)
		Median			68.0	-1.0
		Min, Max			54, 83	-20, 19
	Day 6/Discharge	n	2	2	162	162
		Mean (SD)	NC	NC	71.5 (7.27)	2.0 (8.00)
		Median	NC	NC	70.0	2.0
		Min, Max	NC	NC	54, 90	-19, 21
Supine Pulse Rate (beats/min)	Baseline	n	9		169	
		Mean (SD)	83.4 (9.06)		75.2 (9.86)	
		Median	81.0		75.0	
		Min, Max	75, 103		50, 103	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects who were enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Change is change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: NC = Not calculated

Appendix 15.3.6.7

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_vit.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.16 Summary of Supine Vital Signs - Safety Population**

Parameter (units)	Study Day	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)	
			Raw value	Change	Raw value	Change	Raw value	Change
Supine Pulse Rate (beats/min)	Day 1	n	80	80	41	41	39	39
		Mean (SD)	72.2 (9.72)	-2.2 (7.48)	72.6 (8.39)	-2.4 (8.44)	68.8 (10.10)	-6.5 (7.22)
		Median	72.0	-2.0	71.0	-3.0	68.0	-7.0
		Min, Max	51, 95	-21, 23	59, 91	-16, 17	51, 98	-26, 11
	Day 2	n	80	80	41	41	39	39
		Mean (SD)	74.4 (9.41)	0.0 (8.13)	76.3 (9.16)	1.3 (10.04)	69.4 (8.36)	-5.9 (7.48)
		Median	76.0	-1.0	76.0	1.0	71.0	-6.0
		Min, Max	48, 99	-23, 21	59, 98	-20, 30	54, 87	-27, 10
	Day 3	n	80	80	41	41	39	39
		Mean (SD)	76.3 (11.12)	2.0 (8.55)	78.5 (11.61)	3.5 (10.22)	69.4 (8.81)	-6.0 (8.35)
		Median	75.0	2.0	78.0	4.0	68.0	-5.0
		Min, Max	51, 109	-15, 30	60, 119	-14, 24	59, 97	-25, 10

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects who were enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Change is change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.7

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_vit.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.16 Summary of Supine Vital Signs - Safety Population**

Parameter (units)	Study Day	Statistic	Enrolled Not Randomized (N=9)		Overall Safety (N=169)	
			Raw value	Change	Raw value	Change
Supine Pulse Rate (beats/min)	Day 1	n	0	0	160	160
		Mean (SD)			71.5 (9.56)	-3.3 (7.84)
		Median			70.0	-4.0
		Min, Max			51, 98	-26, 23
	Day 2	n	0	0	160	160
		Mean (SD)			73.7 (9.40)	-1.1 (8.91)
		Median			74.0	-1.0
		Min, Max			48, 99	-27, 30
	Day 3	n	0	0	160	160
		Mean (SD)			75.2 (11.21)	0.4 (9.64)
		Median			74.0	-1.0
		Min, Max			51, 119	-25, 30

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects who were enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Change is change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.7

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_vit.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.16 Summary of Supine Vital Signs - Safety Population**

Parameter (units)	Study Day	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)	
			Raw value	Change	Raw value	Change	Raw value	Change
Supine Pulse Rate (beats/min)	Day 4	n	79	79	41	41	39	39
		Mean (SD)	74.2 (9.14)	-0.1 (8.29)	74.1 (8.03)	-0.9 (9.09)	69.3 (9.51)	-6.1 (9.06)
		Median	74.0	1.0	74.0	0.0	66.0	-7.0
		Min, Max	52, 93	-27, 21	61, 91	-15, 15	58, 93	-22, 23
	Day 5	n	79	79	41	41	39	39
		Mean (SD)	73.4 (8.99)	-0.9 (7.64)	73.9 (8.39)	-1.1 (7.99)	68.4 (7.87)	-7.0 (9.47)
		Median	72.0	-1.0	73.0	-2.0	67.0	-8.0
		Min, Max	53, 98	-21, 20	59, 93	-20, 13	55, 91	-28, 13
	Day 6/Discharge	n	80	80	41	41	39	39
		Mean (SD)	65.8 (7.61)	-8.6 (9.34)	66.6 (9.32)	-8.4 (9.43)	64.8 (7.65)	-10.6 (9.04)
		Median	64.0	-8.5	64.0	-6.0	63.0	-10.0
		Min, Max	51, 85	-32, 18	50, 90	-36, 12	51, 87	-38, 5

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects who were enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Change is change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.7

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_vit.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.16 Summary of Supine Vital Signs - Safety Population**

Parameter (units)	Study Day	Statistic	Enrolled Not Randomized (N=9)		Overall Safety (N=169)	
			Raw value	Change	Raw value	Change
Supine Pulse Rate (beats/min)	Day 4	n	0	0	159	159
		Mean (SD)			73.0 (9.15)	-1.8 (8.99)
		Median			73.0	-2.0
		Min, Max			52, 93	-27, 23
	Day 5	n	0	0	159	159
		Mean (SD)			72.3 (8.81)	-2.5 (8.55)
		Median			72.0	-2.0
		Min, Max			53, 98	-28, 20
	Day 6/Discharge	n	2	2	162	162
		Mean (SD)	NC	NC	65.7 (8.06)	-9.2 (9.39)
		Median	NC	NC	64.0	-9.0
		Min, Max	NC	NC	50, 90	-38, 18

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects who were enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Change is change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: NC = Not calculated

Appendix 15.3.6.7

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.16 Summary of Supine Vital Signs - Safety Population**

Parameter (units)	Study Day	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)	
			Raw value	Change	Raw value	Change	Raw value	Change
Supine Respiratory Rate (breaths/ min)	Baseline	n	80		41		39	
		Mean (SD)	16.1 (1.26)		15.9 (1.11)		16.1 (1.08)	
		Median	16.0		16.0		16.0	
		Min, Max	14, 18		14, 18		14, 18	
	Day 1	n	80	80	41	41	39	39
		Mean (SD)	15.2 (1.60)	-0.8 (2.08)	14.7 (2.13)	-1.1 (2.31)	15.2 (1.62)	-0.9 (2.00)
		Median	16.0	0.0	14.0	-1.0	16.0	0.0
		Min, Max	12, 18	-6, 4	12, 20	-6, 4	12, 18	-5, 3
	Day 2	n	80	80	41	41	39	39
		Mean (SD)	15.7 (1.60)	-0.4 (1.89)	16.0 (1.76)	0.1 (1.81)	15.0 (1.94)	-1.1 (2.12)
		Median	16.0	0.0	16.0	0.0	15.0	-2.0
		Min, Max	12, 20	-5, 4	12, 20	-4, 4	12, 20	-5, 4

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects who were enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Change is change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.16 Summary of Supine Vital Signs - Safety Population**

Parameter (units)	Study Day	Statistic	Enrolled Not Randomized (N=9)		Overall Safety (N=169)	
			Raw value	Change	Raw value	Change
Supine Respiratory Rate (breaths/ min)	Baseline	n	9		169	
		Mean (SD)	15.9 (1.62)		16.0 (1.20)	
		Median	16.0		16.0	
		Min, Max	14, 18		14, 18	
	Day 1	n	0	0	160	160
		Mean (SD)			15.1 (1.75)	-0.9 (2.11)
		Median			16.0	0.0
		Min, Max			12, 20	-6, 4
	Day 2	n	0	0	160	160
		Mean (SD)			15.6 (1.76)	-0.4 (1.96)
		Median			16.0	0.0
		Min, Max			12, 20	-5, 4

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects who were enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Change is change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.16 Summary of Supine Vital Signs - Safety Population**

Parameter (units)	Study Day	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)	
			Raw value	Change	Raw value	Change	Raw value	Change
Supine Respiratory Rate (breaths/ min)	Day 3	n	80	80	41	41	39	39
		Mean (SD)	16.0 (1.56)	0.0 (1.98)	16.0 (1.29)	0.2 (1.73)	16.5 (1.45)	0.4 (1.94)
		Median	16.0	0.0	16.0	0.0	16.0	0.0
		Min, Max	12, 20	-4, 4	14, 20	-2, 4	14, 18	-4, 4
	Day 4	n	79	79	41	41	39	39
		Mean (SD)	15.2 (1.63)	-0.9 (1.93)	15.6 (1.86)	-0.3 (2.27)	15.4 (1.93)	-0.7 (2.04)
		Median	16.0	-1.0	16.0	0.0	16.0	0.0
		Min, Max	12, 20	-5, 4	12, 20	-4, 5	10, 20	-6, 4
	Day 5	n	79	79	41	41	39	39
		Mean (SD)	15.3 (1.52)	-0.7 (1.90)	15.6 (1.58)	-0.2 (1.83)	16.1 (1.60)	0.0 (1.73)
		Median	16.0	0.0	16.0	0.0	16.0	0.0
		Min, Max	12, 18	-5, 4	12, 18	-4, 3	12, 18	-4, 2

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects who were enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Change is change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

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Path: /cvn/projects/prj/development/000000106324/dev/tables/t_vit.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.16 Summary of Supine Vital Signs - Safety Population**

Parameter (units)	Study Day	Statistic	Enrolled Not Randomized (N=9)		Overall Safety (N=169)	
			Raw value	Change	Raw value	Change
Supine Respiratory Rate (breaths/ min)	Day 3	n	0	0	160	160
		Mean (SD)			16.1 (1.47)	0.1 (1.90)
		Median			16.0	0.0
		Min, Max			12, 20	-4, 4
	Day 4	n	0	0	159	159
		Mean (SD)			15.3 (1.76)	-0.7 (2.05)
		Median			16.0	0.0
		Min, Max			10, 20	-6, 5
	Day 5	n	0	0	159	159
		Mean (SD)			15.6 (1.58)	-0.4 (1.86)
		Median			16.0	0.0
		Min, Max			12, 18	-5, 4

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects who were enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Change is change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

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Path: /cvn/projects/prj/development/000000106324/dev/tables/t_vit.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.16 Summary of Supine Vital Signs - Safety Population**

Parameter (units)	Study Day	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)	
			Raw value	Change	Raw value	Change	Raw value	Change
Supine Respiratory Rate (breaths/ min)	Day 6/Discharge	n	80	80	41	41	39	39
		Mean (SD)	15.0 (1.47)	-1.0 (1.99)	14.9 (1.18)	-0.9 (1.56)	15.1 (1.23)	-1.0 (1.73)
		Median	16.0	-1.0	14.0	-1.0	15.0	-1.0
		Min, Max	12, 20	-6, 6	12, 18	-4, 2	12, 18	-4, 4

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects who were enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Change is change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

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Path: /cvn/projects/prj/development/000000106324/dev/tables/t_vit.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.16 Summary of Supine Vital Signs - Safety Population**

Parameter (units)	Study Day	Statistic	Enrolled Not Randomized (N=9)		Overall Safety (N=169)	
			Raw value	Change	Raw value	Change
Supine Respiratory Rate (breaths/ min)	Day 6/Discharge	n	2	2	162	162
		Mean (SD)	NC	NC	15.0 (1.35)	-1.0 (1.86)
		Median	NC	NC	15.0	-1.0
		Min, Max	NC	NC	12, 20	-6, 6

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects who were enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Change is change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: NC = Not calculated

Appendix 15.3.6.7

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_vit.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.17 Summary of ECG Measurements - Safety Population**

Parameter (units)	Study Day	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)	
			Raw value	Change	Raw value	Change	Raw value	Change
Heart rate (beats/min)	Baseline	n	80		41		39	
		Mean (SD)	68.6 (9.39)		69.5 (10.05)		70.9 (11.27)	
		Median	67.5		70.0		71.0	
		Min, Max	52, 99		50, 90		48, 95	
	Day 6/ Discharge	n	79	79	41	41	39	39
		Mean (SD)	64.0 (8.38)	-4.4 (9.40)	65.7 (8.34)	-3.8 (9.59)	64.0 (8.78)	-6.9 (9.67)
		Median	63.0	-4.0	66.0	-6.0	63.0	-7.0
		Min, Max	50, 95	-28, 21	50, 85	-18, 15	50, 82	-24, 9

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Change is change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.9

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_ecg.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.17 Summary of ECG Measurements - Safety Population**

Parameter (units)	Study Day	Statistic	Enrolled not randomized (N=9)		Overall Safety (N=169)	
			Raw value	Change	Raw value	Change
Heart rate (beats/min) Baseline		n	9		169	
		Mean (SD)	70.0 (10.23)		69.4 (10.00)	
		Median	71.0		70.0	
		Min, Max	55, 86		48, 99	
Day 6/ Discharge		n	4	4	163	163
		Mean (SD)	82.8 (26.02)	11.3 (18.40)	64.9 (9.50)	-4.5 (10.06)
		Median	80.5	13.0	63.0	-4.0
		Min, Max	60, 110	-10, 29	50, 110	-28, 29

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Change is change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.9

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_ecg.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.17 Summary of ECG Measurements - Safety Population**

Parameter (units)	Study Day	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)	
			Raw value	Change	Raw value	Change	Raw value	Change
PR duration (msec)	Baseline	n	80		41		39	
		Mean (SD)	154.7 (16.78)		153.9 (23.19)		156.0 (18.99)	
		Median	160.0		150.0		158.0	
		Min, Max	120, 190		85, 200		120, 200	
	Day 6/ Discharge	n	79	79	41	41	39	39
		Mean (SD)	154.6 (18.09)	0.2 (11.68)	157.8 (18.24)	3.9 (19.42)	158.2 (18.91)	2.2 (14.33)
		Median	150.0	0.0	160.0	0.0	160.0	0.0
		Min, Max	120, 190	-35, 30	120, 200	-40, 85	130, 200	-30, 40

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Change is change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.9

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_ecg.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.17 Summary of ECG Measurements - Safety Population**

Parameter (units)	Study Day	Statistic	Enrolled not randomized (N=9)		Overall Safety (N=169)	
			Raw value	Change	Raw value	Change
PR duration (msec)	Baseline	n	9		169	
		Mean (SD)	157.8 (20.49)		155.0 (19.05)	
		Median	160.0		160.0	
		Min, Max	120, 190		85, 200	
	Day 6/ Discharge	n	4	4	163	163
		Mean (SD)	165.0 (52.60)	2.5 (33.05)	156.5 (19.48)	1.7 (15.15)
		Median	150.0	-10.0	150.0	0.0
		Min, Max	120, 240	-20, 50	120, 240	-40, 85

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Change is change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.9

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_ecg.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.17 Summary of ECG Measurements - Safety Population**

Parameter (units)	Study Day	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)	
			Raw value	Change	Raw value	Change	Raw value	Change
QT duration (msec)	Baseline	n	80		41		39	
		Mean (SD)	397.1 (25.08)		393.8 (25.63)		389.9 (25.70)	
		Median	400.0		400.0		390.0	
		Min, Max	340, 460		320, 440		340, 460	
	Day 6/ Discharge	n	79	79	41	41	39	39
		Mean (SD)	398.6 (20.64)	1.0 (22.19)	395.5 (22.97)	1.7 (22.49)	399.1 (23.17)	9.1 (22.99)
		Median	392.0	0.0	400.0	0.0	400.0	10.0
		Min, Max	360, 460	-50, 50	330, 440	-50, 40	340, 440	-40, 60

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Change is change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.9

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_ecg.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.17 Summary of ECG Measurements - Safety Population**

Parameter (units)	Study Day	Statistic	Enrolled not randomized (N=9)		Overall Safety (N=169)	
			Raw value	Change	Raw value	Change
QT duration (msec)	Baseline	n	9		169	
		Mean (SD)	392.4 (26.06)		394.4 (25.34)	
		Median	400.0		400.0	
		Min, Max	360, 430		320, 460	
	Day 6/ Discharge	n	4	4	163	163
		Mean (SD)	360.0 (0.00)	-25.5 (24.52)	397.0 (22.31)	2.5 (22.98)
		Median	360.0	-25.0	392.0	0.0
		Min, Max	360, 360	-52, 0	330, 460	-52, 60

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Change is change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.9

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_ecg.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.17 Summary of ECG Measurements - Safety Population**

Parameter (units)	Study Day	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)	
			Raw value	Change	Raw value	Change	Raw value	Change
QTcB - Bazett's correction formula (msec)	Baseline	n	80		41		39	
		Mean (SD)	422.3 (20.95)		418.9 (21.92)		417.2 (23.33)	
		Median	419.0		412.0		422.0	
		Min, Max	371, 461		383, 478		350, 458	
	Day 6/ Discharge	n	79	79	41	41	39	39
		Mean (SD)	414.6 (18.78)	-7.8 (22.97)	413.9 (17.33)	-5.0 (21.20)	414.9 (22.66)	-2.2 (26.13)
		Median	415.0	-7.0	416.0	-3.0	415.0	-5.0
		Min, Max	367, 475	-62, 60	375, 447	-55, 35	373, 478	-43, 80

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Change is change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.9

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.17 Summary of ECG Measurements - Safety Population**

Parameter (units)	Study Day	Statistic	Enrolled not randomized (N=9)		Overall Safety (N=169)	
			Raw value	Change	Raw value	Change
QTcB - Bazett's correction formula (msec)	Baseline	n	9		169	
		Mean (SD)	425.3 (16.94)		420.4 (21.53)	
		Median	430.0		420.0	
		Min, Max	400, 449		350, 478	
	Day 6/ Discharge	n	4	4	163	163
		Mean (SD)	400.3 (49.55)	-31.0 (41.97)	414.2 (20.37)	-6.3 (24.05)
		Median	388.0	-43.0	415.0	-5.0
		Min, Max	360, 465	-67, 29	360, 478	-67, 80

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Change is change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.9

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.17 Summary of ECG Measurements - Safety Population**

Parameter (units)	Study Day	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)	
			Raw value	Change	Raw value	Change	Raw value	Change
QTcF - Fridericia's correction formula (msec)	Baseline	n	80		41		39	
		Mean (SD)	413.6 (18.04)		411.7 (18.99)		410.0 (19.31)	
		Median	412.9		412.9		410.8	
		Min, Max	371, 459		365, 459		368, 440	
	Day 6/ Discharge	n	79	79	41	41	39	39
		Mean (SD)	405.9 (15.33)	-7.9 (18.99)	406.2 (14.41)	-5.5 (17.43)	406.2 (17.02)	-3.9 (17.34)
		Median	405.4	-7.8	410.6	-4.3	404.6	-4.0
		Min, Max	364, 443	-76, 37	371, 428	-59, 27	371, 439	-39, 33

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Change is change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.9

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_ecg.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.17 Summary of ECG Measurements - Safety Population**

Parameter (units)	Study Day	Statistic	Enrolled not randomized (N=9)		Overall Safety (N=169)	
			Raw value	Change	Raw value	Change
QTcF - Fridericia's correction formula (msec)	Baseline	n	9		169	
		Mean (SD)	411.3 (17.77)		412.2 (18.45)	
		Median	416.5		412.9	
		Min, Max	381, 442		365, 459	
	Day 6/ Discharge	n	4	4	163	163
		Mean (SD)	397.4 (42.37)	-9.7 (30.45)	405.9 (16.35)	-6.4 (18.42)
		Median	394.4	-7.5	405.5	-5.3
		Min, Max	360, 441	-47, 23	360, 443	-76, 37

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Change is change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.9

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.17 Summary of ECG Measurements - Safety Population**

Parameter (units)	Study Day	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)	
			Raw value	Change	Raw value	Change	Raw value	Change
QRS duration (msec)	Baseline	n	80		41		39	
		Mean (SD)	97.1 (9.71)		93.5 (9.56)		94.1 (9.80)	
		Median	100.0		95.0		95.0	
		Min, Max	80, 140		80, 120		80, 130	
	Day 6/ Discharge	n	79	79	41	41	39	39
		Mean (SD)	94.5 (7.46)	-2.8 (7.36)	94.1 (6.98)	0.6 (7.72)	93.7 (8.46)	-0.4 (5.09)
		Median	95.0	0.0	95.0	0.0	95.0	0.0
		Min, Max	80, 120	-35, 20	80, 110	-10, 30	75, 130	-10, 10

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Change is change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.9

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**Table 15.2.6.17 Summary of ECG Measurements - Safety Population**

Parameter (units)	Study Day	Statistic	Enrolled not randomized (N=9)		Overall Safety (N=169)	
			Raw value	Change	Raw value	Change
QRS duration (msec)	Baseline	n	9		169	
		Mean (SD)	96.0 (12.81)		95.5 (9.92)	
		Median	100.0		100.0	
		Min, Max	80, 110		80, 140	
	Day 6/ Discharge	n	4	4	163	163
		Mean (SD)	97.5 (17.08)	9.0 (18.00)	94.3 (7.83)	-1.0 (7.59)
		Median	95.0	0.0	95.0	0.0
		Min, Max	80, 120	0, 36	75, 130	-35, 36

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Change is change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.9

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.17 Summary of ECG Measurements - Safety Population**

Parameter (units)	Study Day	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)	
			Raw value	Change	Raw value	Change	Raw value	Change
Interpretation	Baseline	Normal - n (%)	70 (87.5%)		37 (90.2%)		36 (92.3%)	
		Abnormal non-clinically significant - n (%)	10 (12.5%)		4 (9.8%)		3 (7.7%)	
		Abnormal clinically significant - n (%)	0		0		0	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Change is change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

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**Table 15.2.6.17 Summary of ECG Measurements - Safety Population**

Parameter (units)	Study Day	Statistic	Enrolled not randomized (N=9)		Overall Safety (N=169)	
			Raw value	Change	Raw value	Change
Interpretation	Baseline	Normal - n (%)	8 (88.9%)		151 (89.3%)	
		Abnormal non-clinically significant - n (%)	1 (11.1%)		18 (10.7%)	
		Abnormal clinically significant - n (%)	0		0	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Change is change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

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**Table 15.2.6.17 Summary of ECG Measurements - Safety Population**

Parameter (units)	Study Day	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)	
			Raw value	Change	Raw value	Change	Raw value	Change
Interpretation	Day 6/ Discharge	Normal - n (%)	68 (85.0%)		39 (95.1%)		33 (84.6%)	
		Abnormal non-clinically significant - n (%)	10 (12.5%)		2 (4.9%)		6 (15.4%)	
		Abnormal clinically significant - n (%)	1 (1.3%)		0		0	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Change is change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.9

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**Table 15.2.6.17 Summary of ECG Measurements - Safety Population**

Parameter (units)	Study Day	Statistic	Enrolled not randomized (N=9)		Overall Safety (N=169)	
			Raw value	Change	Raw value	Change
Interpretation	Day 6/ Discharge	Normal - n (%)	2 (22.2%)		142 (84.0%)	
		Abnormal non-clinically significant - n (%)	1 (11.1%)		19 (11.2%)	
		Abnormal clinically significant - n (%)	1 (11.1%)		2 (1.2%)	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Change is change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.9

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_ecg.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.18 Summary of Spirometry Measurements - Safety Population**

Parameter (units)	Study Day	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)	
			Raw value	Change	Raw value	Change	Raw value	Change
Best measured FVC value (with bronchodilator) (L)	Baseline	n	80		41		39	
		Mean (SD)	4.796 (1.1134)		4.720 (1.1895)		4.735 (1.3255)	
		Median	4.730		4.560		5.090	
		Min, Max	2.81, 7.52		2.92, 7.82		2.81, 7.44	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Change is change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.8

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_spiro.sas

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**Table 15.2.6.18 Summary of Spirometry Measurements - Safety Population**

Parameter (units)	Study Day	Statistic	Enrolled Not Randomized (N=9)		Overall Safety (N=169)	
			Raw value	Change	Raw value	Change
Best measured FVC value (with bronchodilator) (L)	Baseline	n	9		169	
		Mean (SD)	4.599 (1.2452)		4.753 (1.1805)	
		Median	4.440		4.690	
		Min, Max	2.88, 6.73		2.81, 7.82	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Change is change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.8

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_spiro.sas

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**Table 15.2.6.18 Summary of Spirometry Measurements - Safety Population**

Parameter (units)	Study Day	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)	
			Raw value	Change	Raw value	Change	Raw value	Change
Percent of predicted FVC value (with bronchodilator) (%)	Baseline	n	80		41		39	
		Mean (SD)	103.0 (10.42)		101.0 (11.32)		101.2 (11.88)	
		Median	103.5		100.0		99.0	
		Min, Max	81, 132		81, 134		84, 127	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Change is change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.8

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.18 Summary of Spirometry Measurements - Safety Population**

Parameter (units)	Study Day	Statistic	Enrolled Not Randomized (N=9)		Overall Safety (N=169)	
			Raw value	Change	Raw value	Change
Percent of predicted FVC value (with bronchodilator) (%)	Baseline	n	9		169	
		Mean (SD)	106.6 (11.92)		102.3 (11.05)	
		Median	107.0		102.0	
		Min, Max	79, 118		79, 134	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Change is change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.18 Summary of Spirometry Measurements - Safety Population**

Parameter (units)	Study Day	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)	
			Raw value	Change	Raw value	Change	Raw value	Change
Best measured FEV1 value (with bronchodilator) (L)	Baseline	n	80		41		39	
		Mean (SD)	3.953 (0.8680)		3.906 (0.9499)		3.958 (1.0401)	
		Median	3.875		3.800		4.070	
		Min, Max	2.32, 5.98		2.40, 5.94		2.22, 6.26	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Change is change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

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**Table 15.2.6.18 Summary of Spirometry Measurements - Safety Population**

Parameter (units)	Study Day	Statistic	Enrolled Not Randomized (N=9)		Overall Safety (N=169)	
			Raw value	Change	Raw value	Change
Best measured FEV1 value (with bronchodilator) (L)	Baseline	n	9		169	
		Mean (SD)	3.796 (1.1121)		3.934 (0.9350)	
		Median	3.680		3.850	
		Min, Max	2.36, 5.80		2.22, 6.26	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Change is change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.8

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_spiro.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.18 Summary of Spirometry Measurements - Safety Population**

Parameter (units)	Study Day	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)	
			Raw value	Change	Raw value	Change	Raw value	Change
Percent of predicted FEV1 value (with bronchodilator) (%)	Baseline	n	80		41		39	
		Mean (SD)	104.1 (9.91)		102.0 (11.00)		103.9 (12.83)	
		Median	103.0		101.0		104.0	
		Min, Max	78, 129		82, 123		81, 127	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Change is change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.8

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_spiro.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.18 Summary of Spirometry Measurements - Safety Population**

Parameter (units)	Study Day	Statistic	Enrolled Not Randomized (N=9)		Overall Safety (N=169)	
			Raw value	Change	Raw value	Change
Percent of predicted FEV1 value (with bronchodilator) (%)	Baseline	n	9		169	
		Mean (SD)	110.1 (12.71)		103.8 (11.09)	
		Median	111.0		103.0	
		Min, Max	83, 126		78, 129	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Change is change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.8

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_spiro.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.18 Summary of Spirometry Measurements - Safety Population**

Parameter (units)	Study Day	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)	
			Raw value	Change	Raw value	Change	Raw value	Change
Calculated ratio between FEV1/FVC (with bronchodilator) (RATIO)	Baseline	n	80		41		39	
		Mean (SD)	0.828 (0.0544)		0.831 (0.0491)		0.842 (0.0564)	
		Median	0.825		0.830		0.850	
		Min, Max	0.71, 1.00		0.73, 0.91		0.71, 0.97	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Change is change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.8

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_spiro.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.18 Summary of Spirometry Measurements - Safety Population**

Parameter (units)	Study Day	Statistic	Enrolled Not Randomized (N=9)		Overall Safety (N=169)	
			Raw value	Change	Raw value	Change
Calculated ratio between FEV1/FVC (with bronchodilator) (RATIO)	Baseline	n	9		169	
		Mean (SD)	0.821 (0.0318)		0.831 (0.0526)	
		Median	0.830		0.830	
		Min, Max	0.77, 0.86		0.71, 1.00	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Change is change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.8

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.18 Summary of Spirometry Measurements - Safety Population**

Parameter (units)	Study Day	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)	
			Raw value	Change	Raw value	Change	Raw value	Change
Best measured FVC value (L)	Baseline	n	80		41		39	
		Mean (SD)	4.801 (1.0487)		4.693 (1.1393)		4.725 (1.2554)	
		Median	4.715		4.570		4.760	
		Min, Max	2.81, 7.64		3.01, 7.05		2.90, 7.45	
	Day 6/ Discharge	n	79	79	41	41	39	39
		Mean (SD)	4.857 (1.1283)	0.055 (0.3915)	4.765 (1.2047)	0.072 (0.3165)	4.814 (1.3593)	0.090 (0.3750)
		Median	4.680	0.030	4.520	0.030	4.940	0.050
		Min, Max	2.95, 7.45	-2.41, 0.85	2.84, 7.52	-0.56, 1.15	2.98, 8.32	-1.19, 1.15

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Change is change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

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**Table 15.2.6.18 Summary of Spirometry Measurements - Safety Population**

Parameter (units)	Study Day	Statistic	Enrolled Not Randomized (N=9)		Overall Safety (N=169)	
			Raw value	Change	Raw value	Change
Best measured FVC value (L)	Baseline	n	9		169	
		Mean (SD)	4.443 (0.9939)		4.738 (1.1121)	
		Median	4.500		4.710	
		Min, Max	2.85, 5.61		2.81, 7.64	
	Day 6/ Discharge	n	0		159	159
		Mean (SD)			4.823 (1.2007)	0.068 (0.3675)
		Median			4.680	0.030
		Min, Max			2.84, 8.32	-2.41, 1.15

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Change is change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.8

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.18 Summary of Spirometry Measurements - Safety Population**

Parameter (units)	Study Day	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)	
			Raw value	Change	Raw value	Change	Raw value	Change
Percent of predicted FVC value (%)	Baseline	n	80		41		39	
		Mean (SD)	103.4 (12.14)		100.3 (9.98)		100.7 (10.61)	
		Median	102.5		99.0		98.0	
		Min, Max	79, 167		84, 120		83, 123	
	Day 6/ Discharge	n	79	79	41	41	39	39
		Mean (SD)	104.0 (11.50)	0.8 (9.94)	102.4 (12.61)	2.0 (6.59)	103.5 (12.15)	2.9 (6.10)
		Median	103.0	0.0	103.0	0.0	100.0	1.0
		Min, Max	83, 140	-70, 19	81, 139	-8, 21	88, 135	-5, 19

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Change is change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

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**Table 15.2.6.18 Summary of Spirometry Measurements - Safety Population**

Parameter (units)	Study Day	Statistic	Enrolled Not Randomized (N=9)		Overall Safety (N=169)	
			Raw value	Change	Raw value	Change
Percent of predicted FVC value (%)	Baseline	n	9		169	
		Mean (SD)	104.0 (11.70)		102.0 (11.28)	
		Median	107.0		101.0	
		Min, Max	78, 115		78, 167	
	Day 6/ Discharge	n	0		159	159
		Mean (SD)			103.5 (11.90)	1.6 (8.34)
		Median			103.0	1.0
		Min, Max			81, 140	-70, 21

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Change is change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

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**Table 15.2.6.18 Summary of Spirometry Measurements - Safety Population**

Parameter (units)	Study Day	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)	
			Raw value	Change	Raw value	Change	Raw value	Change
Best measured FEV1 value (L)	Baseline	n	80		41		39	
		Mean (SD)	3.741 (0.7981)		3.729 (0.8666)		3.758 (0.9458)	
		Median	3.655		3.710		3.580	
		Min, Max	2.11, 5.67		2.37, 5.28		2.18, 5.82	
	Day 6/ Discharge	n	79	79	41	41	39	39
		Mean (SD)	3.817 (0.8374)	0.079 (0.2720)	3.749 (0.8919)	0.020 (0.2461)	3.852 (1.0146)	0.093 (0.3101)
		Median	3.740	0.080	3.570	0.030	3.810	0.060
		Min, Max	2.30, 5.80	-0.71, 0.84	2.10, 5.43	-0.81, 0.52	2.24, 6.25	-0.56, 0.96

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Change is change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

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**Table 15.2.6.18 Summary of Spirometry Measurements - Safety Population**

Parameter (units)	Study Day	Statistic	Enrolled Not Randomized (N=9)		Overall Safety (N=169)	
			Raw value	Change	Raw value	Change
Best measured FEV1 value (L)	Baseline	n	9		169	
		Mean (SD)	3.474 (0.7994)		3.728 (0.8455)	
		Median	3.710		3.650	
		Min, Max	2.06, 4.32		2.06, 5.82	
	Day 6/ Discharge	n	0		159	159
		Mean (SD)			3.808 (0.8925)	0.067 (0.2753)
		Median			3.740	0.070
		Min, Max			2.10, 6.25	-0.81, 0.96

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Change is change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.18 Summary of Spirometry Measurements - Safety Population**

Parameter (units)	Study Day	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)	
			Raw value	Change	Raw value	Change	Raw value	Change
Percent of predicted FEV1 value (%)	Baseline	n	80		41		39	
		Mean (SD)	98.9 (10.90)		97.3 (9.86)		98.7 (12.03)	
		Median	99.0		97.0		102.0	
		Min, Max	77, 126		81, 116		71, 119	
	Day 6/ Discharge	n	79	79	41	41	39	39
		Mean (SD)	101.0 (12.39)	2.3 (6.98)	98.8 (13.34)	1.5 (7.70)	101.4 (13.20)	2.6 (7.73)
		Median	100.0	2.0	99.0	1.0	103.0	2.0
		Min, Max	76, 130	-24, 21	75, 134	-23, 23	75, 125	-12, 25

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Change is change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

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**Table 15.2.6.18 Summary of Spirometry Measurements - Safety Population**

Parameter (units)	Study Day	Statistic	Enrolled Not Randomized (N=9)		Overall Safety (N=169)	
			Raw value	Change	Raw value	Change
Percent of predicted FEV1 value (%)	Baseline	n	9		169	
		Mean (SD)	102.3 (14.98)		98.7 (11.12)	
		Median	108.0		99.0	
		Min, Max	72, 117		71, 126	
	Day 6/ Discharge	n	0		159	159
		Mean (SD)			100.5 (12.80)	2.2 (7.32)
		Median			100.0	2.0
		Min, Max			75, 134	-24, 25

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Change is change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.18 Summary of Spirometry Measurements - Safety Population**

Parameter (units)	Study Day	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)	
			Raw value	Change	Raw value	Change	Raw value	Change
Calculated ratio between FEV1/FVC (RATIO)	Baseline	n	80		41		39	
		Mean (SD)	0.800 (0.0561)		0.811 (0.0537)		0.813 (0.0605)	
		Median	0.810		0.800		0.820	
		Min, Max	0.68, 0.92		0.68, 0.91		0.66, 0.95	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Change is change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

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**Table 15.2.6.18 Summary of Spirometry Measurements - Safety Population**

Parameter (units)	Study Day	Statistic	Enrolled Not Randomized (N=9)		Overall Safety (N=169)	
			Raw value	Change	Raw value	Change
Calculated ratio between FEV1/FVC (RATIO)	Baseline	n	9		169	
		Mean (SD)	0.793 (0.0381)		0.805 (0.0557)	
		Median	0.780		0.810	
		Min, Max	0.75, 0.85		0.66, 0.95	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Change is change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

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**Table 15.2.6.18 Summary of Spirometry Measurements - Safety Population**

Parameter (units)	Study Day	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)	
			Raw value	Change	Raw value	Change	Raw value	Change
Calculated ratio between FEV1/FVC ¹ (RATIO)	Baseline	n	80		41		39	
		Mean (SD)	0.783 (0.0657)		0.799 (0.0520)		0.802 (0.0734)	
		Median	0.785		0.790		0.810	
		Min, Max	0.60, 0.92		0.67, 0.89		0.64, 0.96	
	Day 6/ Discharge	n	79	79	41	41	39	39
		Mean (SD)	0.791 (0.0700)	0.009 (0.0464)	0.792 (0.0699)	-0.007 (0.0457)	0.806 (0.0678)	0.004 (0.0565)
		Median	0.780	0.010	0.800	0.000	0.810	0.000
		Min, Max	0.65, 1.00	-0.19, 0.24	0.63, 0.91	-0.19, 0.11	0.68, 0.95	-0.08, 0.30

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Change is change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: 1: Ratio is derived in analysis datasets.

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**Table 15.2.6.18 Summary of Spirometry Measurements - Safety Population**

Parameter (units)	Study Day	Statistic	Enrolled Not Randomized (N=9)		Overall Safety (N=169)	
			Raw value	Change	Raw value	Change
Calculated ratio between FEV1/FVC ¹ (RATIO)	Baseline	n	9		169	
		Mean (SD)	0.778 (0.0282)		0.791 (0.0633)	
		Median	0.780		0.790	
		Min, Max	0.72, 0.82		0.60, 0.96	
	Day 6/ Discharge	n	0		159	159
		Mean (SD)			0.795 (0.0693)	0.004 (0.0490)
		Median			0.790	0.000
		Min, Max			0.63, 1.00	-0.19, 0.30

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Change is change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: 1: Ratio is derived in analysis datasets.

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.18 Summary of Spirometry Measurements - Safety Population**

Parameter (units)	Study Day	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)	
			Raw value	Change	Raw value	Change	Raw value	Change
Interpretation (with bronchodilator)	Baseline	Normal - n (%)	80 (100 %)		41 (100 %)		39 (100 %)	
		Abnormal non-clinically significant - n (%)	0		0		0	
		Abnormal clinically significant - n (%)	0		0		0	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Change is change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

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**Table 15.2.6.18 Summary of Spirometry Measurements - Safety Population**

Parameter (units)	Study Day	Statistic	Enrolled Not Randomized (N=9)		Overall Safety (N=169)	
			Raw value	Change	Raw value	Change
Interpretation (with bronchodilator)	Baseline	Normal - n (%)	9 (100 %)		169 (100 %)	
		Abnormal non-clinically significant - n (%)	0		0	
		Abnormal clinically significant - n (%)	0		0	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Change is change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

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**Table 15.2.6.18 Summary of Spirometry Measurements - Safety Population**

Parameter (units)	Study Day	Statistic	THS 2.2 (N=80)		CC (N=41)		SA (N=39)	
			Raw value	Change	Raw value	Change	Raw value	Change
Interpretation	Baseline	Normal - n (%)	75 (93.8%)		39 (95.1%)		35 (89.7%)	
		Abnormal non-clinically significant - n (%)	4 (5.0%)		2 (4.9%)		4 (10.3%)	
		Abnormal clinically significant - n (%)	1 (1.3%)		0		0	
	Day 6/ Discharge	Normal - n (%)	76 (95.0%)		36 (87.8%)		35 (89.7%)	
		Abnormal non-clinically significant - n (%)	1 (1.3%)		1 (2.4%)		3 (7.7%)	
		Abnormal clinically significant - n (%)	2 (2.5%)		3 (7.3%)		1 (2.6%)	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Change is change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.8

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_spiro.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.18 Summary of Spirometry Measurements - Safety Population**

Parameter (units)	Study Day	Statistic	Enrolled Not Randomized (N=9)		Overall Safety (N=169)	
			Raw value	Change	Raw value	Change
Interpretation	Baseline	Normal - n (%)	6 (66.7%)		155 (91.7%)	
		Abnormal non-clinically significant - n (%)	0		10 (5.9%)	
		Abnormal clinically significant - n (%)	3 (33.3%)		4 (2.4%)	
	Day 6/ Discharge	Normal - n (%)	0		147 (87.0%)	
		Abnormal non-clinically significant - n (%)	0		5 (3.0%)	
		Abnormal clinically significant - n (%)	0		6 (3.6%)	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Change is change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Appendix 15.3.6.8

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_spiro.sas

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.19 Summary of Weight and BMI Measurements - Safety Population**

Parameter (units)	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Weight (kg)	Baseline	n	80	41	39	9	169
		Mean (SD)	72.72 (11.708)	76.05 (14.579)	72.52 (11.614)	74.24 (15.708)	73.56 (12.627)
		Median	70.65	75.40	72.90	73.90	72.90
		Min, Max	49.7, 103.4	48.2, 106.0	52.3, 99.0	52.0, 102.8	48.2, 106.0
	Day 6/ Discharge	n	79	41	39	0	159
		Mean (SD)	72.73 (11.578)	76.06 (14.285)	73.63 (11.861)		73.81 (12.392)
		Median	71.10	75.30	72.40		72.50
		Min, Max	49.5, 102.5	49.7, 105.5	53.5, 100.8		49.5, 105.5

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects who were enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Change is change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: 1: BMI is derived in analysis datasets.

Appendix 15.3.6.10

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_bmi.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.19 Summary of Weight and BMI Measurements - Safety Population**

Parameter (units)	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
BMI ¹ (kg/m ²)	Baseline	n	80	41	39	9	169
		Mean (SD)	24.46 (3.034)	25.80 (3.228)	24.81 (2.505)	26.18 (3.896)	24.96 (3.055)
		Median	24.35	26.00	24.60	25.80	24.70
		Min, Max	18.9, 31.6	18.8, 31.8	20.3, 31.6	21.4, 31.6	18.8, 31.8
	Day 6/ Discharge	n	79	41	39	0	159
		Mean (SD)	24.46 (3.053)	25.82 (3.145)	25.19 (2.558)		24.99 (3.001)
		Median	23.90	26.00	24.80		24.80
		Min, Max	18.8, 32.3	19.4, 32.4	20.8, 31.9		18.8, 32.4

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects who were enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Change is change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: 1: BMI is derived in analysis datasets.

Appendix 15.3.6.10

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_bmi.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.19 Summary of Weight and BMI Measurements - Safety Population**

Parameter (units)	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
BMI ¹ (kg/m ²)	Change from Baseline	n	79	41	39	0	159
		Mean (SD)	0.05 (0.384)	0.02 (0.377)	0.38 (0.412)		0.12 (0.415)
		Median	0.00	0.00	0.40		0.10
		Min, Max	-0.8, 0.9	-0.7, 0.6	-0.8, 1.4		-0.8, 1.4

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects who were enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Change is change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: 1: BMI is derived in analysis datasets.

Appendix 15.3.6.10

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_bmi.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.19 Summary of Weight and BMI Measurements - Safety Population**

Parameter (units)	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
BMI (kg/m ²)	Baseline	Underweight - n (%)	0	0	0	0	0
		Normal weight - n (%)	46 (57.5%)	17 (41.5%)	22 (56.4%)	4 (44.4%)	89 (52.7%)
		Overweight - n (%)	29 (36.3%)	19 (46.3%)	16 (41.0%)	3 (33.3%)	67 (39.6%)
		Obese - n (%)	5 (6.3%)	5 (12.2%)	1 (2.6%)	2 (22.2%)	13 (7.7%)
	Day 6/ Discharge	Underweight - n (%)	0	0	0	0	0
		Normal weight - n (%)	46 (57.5%)	17 (41.5%)	21 (53.8%)	0	84 (49.7%)
		Overweight - n (%)	29 (36.3%)	19 (46.3%)	17 (43.6%)	0	65 (38.5%)
		Obese - n (%)	4 (5.0%)	5 (12.2%)	1 (2.6%)	0	10 (5.9%)

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects who were enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Change is change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: 1: BMI is derived in analysis datasets.

Appendix 15.3.6.10

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_bmi.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.19 Summary of Weight and BMI Measurements - Safety Population**

Parameter (units)	Study Day	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled Not Randomized (N=9)	Overall Safety (N=169)
Height (m)	Screening	n	80	41	39	9	169
		Mean (SD)	1.722 (0.0918)	1.710 (0.0926)	1.707 (0.1060)	1.678 (0.0828)	1.713 (0.0948)
		Median	1.710	1.700	1.720	1.660	1.710
		Min, Max	1.57, 2.00	1.52, 1.87	1.51, 1.88	1.56, 1.81	1.51, 2.00

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects who were enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Change is change from baseline, where baseline is the last assessment prior to first product use in CC/THS 2.2 arms on Day 1 or last assessment prior to 06:29 AM in SA arm on Day 1.

Note: 1: BMI is derived in analysis datasets.

Appendix 15.3.6.10

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_bmi.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.20 Summary of Cough Assessments Over Study- Safety Population**

Variable	Sequence										Overall Safety (N=169) n (%) Events	
	THS 2.2 (N=80)		CC (N=41)		SA (N=39)		Enrolled Not Randomized (N=9)					
	n	(%) Events	n	(%) Events	n	(%) Events	n	(%) Events	n	(%) Events		
Has subject experienced cough in the study period assessed?												
Yes	59	(73.8%)	215	27 (65.9%)	110	21 (53.8%)	69	0			107 (63.3%)	394
No	21	(26.3%)	147	14 (34.1%)	98	18 (46.2%)	126	9 (100 %)	9		62 (36.7%)	380
Intensity of cough												
Very mild	13	(16.3%)	31	7 (17.1%)	19	8 (20.5%)	16	0			28 (16.6%)	66
Mild	26	(32.5%)	46	10 (24.4%)	22	6 (15.4%)	12	0			42 (24.9%)	80
Moderate	16	(20.0%)	41	8 (19.5%)	15	7 (17.9%)	16	0			31 (18.3%)	72
Severe	4	(5.0%)	7	2 (4.9%)	4	0		0			6 (3.6%)	11
Very severe	0			0		0		0			0	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Cough experienced in the previous 24h by the subjects is assessed in the morning of Day 0 to Day 6 (to assess Day -1 to Day 5).

Appendix 15.3.6.14

Path: /cvm/projects/prj/development/000000106324/dev/tables/t_cough.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.20 Summary of Cough Assessments Over Study- Safety Population**

Variable	Sequence										Overall Safety (N=169) n (%) Events
	THS 2.2 (N=80)		CC (N=41)		SA (N=39)		Enrolled Not Randomized (N=9)				
	n (%) Events		n (%) Events		n (%) Events		n (%) Events				
Frequency of cough											
Rarely	21 (26.3%)	60	12 (29.3%)	33	11 (28.2%)	25	0			44 (26.0%)	118
Sometimes	28 (35.0%)	57	12 (29.3%)	39	6 (15.4%)	15	0			46 (27.2%)	111
Fairly often	9 (11.3%)	16	2 (4.9%)	5	4 (10.3%)	8	0			15 (8.9%)	29
Often	1 (1.3%)	1	1 (2.4%)	1	0		0			2 (1.2%)	2
Almost always	0		0		0		0			0	
Amount of sputum											
No sputum	24 (30.0%)	79	8 (19.5%)	22	9 (23.1%)	25	0			41 (24.3%)	126
A moderate amount of sputum	34 (42.5%)	101	18 (43.9%)	49	11 (28.2%)	28	0			63 (37.3%)	178
A large amount of sputum	1 (1.3%)	1	1 (2.4%)	1	1 (2.6%)	1	0			3 (1.8%)	3
A very large amount of sputum	0		0		0		0			0	

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages are based on the number of subjects indicated in the column header (N).

Note: Cough experienced in the previous 24h by the subjects is assessed in the morning of Day 0 to Day 6 (to assess Day -1 to Day 5).

Note: If subject has answered question more than once then the most severe intensity is presented.

Appendix 15.3.6.14

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_cough.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.20.1 Summary of Cough Assessments by Study Day - Safety Population**

Study Day/ Variable	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled not randomized (N=9)	Overall Safety (N=169)
Day -1						
Has subject experienced cough in last 24 hours						
Yes	n (%)	24 (30.0%)	15 (36.6%)	9 (23.1%)	0	48 (28.4%)
No	n (%)	56 (70.0%)	26 (63.4%)	30 (76.9%)	9 (100 %)	121 (71.6%)
VAS						
	n	24	15	9		48
	Mean (SD)	17.1 (12.74)	19.1 (13.12)	25.7 (19.13)		19.3 (14.25)
	Median	17.0	18.0	22.0		18.5
	Min, Max	0, 50	2, 49	3, 63		0, 63

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages for number of subjects with a cough are calculated using N in the column header.

Note: The assessments performed at Day 0 to Day 6 are used to evaluate cough at Day -1 to Day 5, respectively.

Appendix 15.3.6.14

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_cough2.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.20.1 Summary of Cough Assessments by Study Day - Safety Population**

Study Day/ Variable	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled not randomized (N=9)	Overall Safety (N=169)
Intensity of cough						
Very mild	n (%)	8 (10.0%)	5 (12.2%)	4 (10.3%)		17 (10.1%)
Mild	n (%)	14 (17.5%)	8 (19.5%)	3 (7.7%)		25 (14.8%)
Moderate	n (%)	2 (2.5%)	2 (4.9%)	2 (5.1%)		6 (3.6%)
Severe	n (%)	0	0	0		0
Very severe	n (%)	0	0	0		0
Frequency of cough						
Rarely	n (%)	15 (18.8%)	10 (24.4%)	5 (12.8%)		30 (17.8%)
Sometimes	n (%)	7 (8.8%)	5 (12.2%)	4 (10.3%)		16 (9.5%)
Fairly often	n (%)	2 (2.5%)	0	0		2 (1.2%)
Often	n (%)	0	0	0		0
Almost always	n (%)	0	0	0		0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages for number of subjects with a cough are calculated using N in the column header.

Note: The assessments performed at Day 0 to Day 6 are used to evaluate cough at Day -1 to Day 5, respectively.

Appendix 15.3.6.14

Path: /cvm/projects/prj/development/000000106324/dev/tables/t_cough2.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.20.1 Summary of Cough Assessments by Study Day - Safety Population**

Study Day/ Variable	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled not randomized (N=9)	Overall Safety (N=169)
Amount of sputum produced						
No sputum	n (%)	12 (15.0%)	9 (22.0%)	4 (10.3%)		25 (14.8%)
A moderate amount of sputum	n (%)	12 (15.0%)	6 (14.6%)	5 (12.8%)		23 (13.6%)
A large amount of sputum	n (%)	0	0	0		0
A very large amount of sputum	n (%)	0	0	0		0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages for number of subjects with a cough are calculated using N in the column header.

Note: The assessments performed at Day 0 to Day 6 are used to evaluate cough at Day -1 to Day 5, respectively.

Appendix 15.3.6.14

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Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

**Table 15.2.6.20.1 Summary of Cough Assessments by Study Day - Safety Population**

Study Day/ Variable	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled not randomized (N=9)	Overall Safety (N=169)
Day 0						
Has subject experienced cough in last 24 hours						
Yes	n (%)	23 (28.8%)	16 (39.0%)	9 (23.1%)	0	48 (28.4%)
No	n (%)	57 (71.3%)	25 (61.0%)	30 (76.9%)	0	112 (66.3%)
VAS						
	n	23	16	9		48
	Mean (SD)	20.2 (15.38)	22.3 (16.99)	26.0 (22.68)		22.0 (17.18)
	Median	17.0	20.5	18.0		18.5
	Min, Max	0, 52	0, 61	2, 74		0, 74

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages for number of subjects with a cough are calculated using N in the column header.

Note: The assessments performed at Day 0 to Day 6 are used to evaluate cough at Day -1 to Day 5, respectively.

Appendix 15.3.6.14

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_cough2.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.20.1 Summary of Cough Assessments by Study Day - Safety Population**

Study Day/ Variable	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled not randomized (N=9)	Overall Safety (N=169)
Intensity of cough						
Very mild	n (%)	7 (8.8%)	10 (24.4%)	2 (5.1%)		19 (11.2%)
Mild	n (%)	7 (8.8%)	3 (7.3%)	3 (7.7%)		13 (7.7%)
Moderate	n (%)	9 (11.3%)	3 (7.3%)	4 (10.3%)		16 (9.5%)
Severe	n (%)	0	0	0		0
Very severe	n (%)	0	0	0		0
Frequency of cough						
Rarely	n (%)	10 (12.5%)	9 (22.0%)	5 (12.8%)		24 (14.2%)
Sometimes	n (%)	12 (15.0%)	7 (17.1%)	2 (5.1%)		21 (12.4%)
Fairly often	n (%)	1 (1.3%)	0	2 (5.1%)		3 (1.8%)
Often	n (%)	0	0	0		0
Almost always	n (%)	0	0	0		0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages for number of subjects with a cough are calculated using N in the column header.

Note: The assessments performed at Day 0 to Day 6 are used to evaluate cough at Day -1 to Day 5, respectively.

Appendix 15.3.6.14

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_cough2.sas

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**Table 15.2.6.20.1 Summary of Cough Assessments by Study Day - Safety Population**

Study Day/ Variable	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled not randomized (N=9)	Overall Safety (N=169)
Amount of sputum produced						
No sputum	n (%)	11 (13.8%)	10 (24.4%)	3 (7.7%)		24 (14.2%)
A moderate amount of sputum	n (%)	12 (15.0%)	6 (14.6%)	5 (12.8%)		23 (13.6%)
A large amount of sputum	n (%)	0	0	1 (2.6%)		1 (0.6%)
A very large amount of sputum	n (%)	0	0	0		0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages for number of subjects with a cough are calculated using N in the column header.

Note: The assessments performed at Day 0 to Day 6 are used to evaluate cough at Day -1 to Day 5, respectively.

Appendix 15.3.6.14

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**Table 15.2.6.20.1 Summary of Cough Assessments by Study Day - Safety Population**

Study Day/ Variable	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled not randomized (N=9)	Overall Safety (N=169)
Day 1						
Has subject experienced cough in last 24 hours						
Yes	n (%)	35 (43.8%)	13 (31.7%)	8 (20.5%)	0	56 (33.1%)
No	n (%)	45 (56.3%)	28 (68.3%)	31 (79.5%)	0	104 (61.5%)
VAS						
	n	35	13	8		56
	Mean (SD)	28.5 (22.27)	28.3 (19.33)	29.1 (26.00)		28.6 (21.78)
	Median	24.0	22.0	21.5		22.5
	Min, Max	0, 90	1, 67	2, 74		0, 90

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages for number of subjects with a cough are calculated using N in the column header.

Note: The assessments performed at Day 0 to Day 6 are used to evaluate cough at Day -1 to Day 5, respectively.

Appendix 15.3.6.14

Path: /cvn/projects/prj/development/000000106324/dev/tables/t_cough2.sas

Program Run: 27OCT14 cvn_kbooth Program Status: FINAL

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**Table 15.2.6.20.1 Summary of Cough Assessments by Study Day - Safety Population**

Study Day/ Variable	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled not randomized (N=9)	Overall Safety (N=169)
Intensity of cough						
Very mild	n (%)	11 (13.8%)	4 (9.8%)	3 (7.7%)		18 (10.7%)
Mild	n (%)	10 (12.5%)	7 (17.1%)	2 (5.1%)		19 (11.2%)
Moderate	n (%)	12 (15.0%)	2 (4.9%)	3 (7.7%)		17 (10.1%)
Severe	n (%)	2 (2.5%)	0	0		2 (1.2%)
Very severe	n (%)	0	0	0		0
Frequency of cough						
Rarely	n (%)	14 (17.5%)	5 (12.2%)	5 (12.8%)		24 (14.2%)
Sometimes	n (%)	16 (20.0%)	6 (14.6%)	1 (2.6%)		23 (13.6%)
Fairly often	n (%)	5 (6.3%)	2 (4.9%)	2 (5.1%)		9 (5.3%)
Often	n (%)	0	0	0		0
Almost always	n (%)	0	0	0		0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages for number of subjects with a cough are calculated using N in the column header.

Note: The assessments performed at Day 0 to Day 6 are used to evaluate cough at Day -1 to Day 5, respectively.

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**Table 15.2.6.20.1 Summary of Cough Assessments by Study Day - Safety Population**

Study Day/ Variable	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled not randomized (N=9)	Overall Safety (N=169)
Amount of sputum produced						
No sputum	n (%)	18 (22.5%)	7 (17.1%)	3 (7.7%)		28 (16.6%)
A moderate amount of sputum	n (%)	17 (21.3%)	6 (14.6%)	5 (12.8%)		28 (16.6%)
A large amount of sputum	n (%)	0	0	0		0
A very large amount of sputum	n (%)	0	0	0		0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages for number of subjects with a cough are calculated using N in the column header.

Note: The assessments performed at Day 0 to Day 6 are used to evaluate cough at Day -1 to Day 5, respectively.

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**Table 15.2.6.20.1 Summary of Cough Assessments by Study Day - Safety Population**

Study Day/ Variable	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled not randomized (N=9)	Overall Safety (N=169)
Day 2						
Has subject experienced cough in last 24 hours						
Yes	n (%)	38 (47.5%)	15 (36.6%)	14 (35.9%)	0	67 (39.6%)
No	n (%)	42 (52.5%)	26 (63.4%)	25 (64.1%)	0	93 (55.0%)
VAS						
	n	38	15	14		67
	Mean (SD)	30.1 (23.19)	26.5 (16.05)	28.7 (28.26)		29.0 (22.71)
	Median	24.5	24.0	16.5		24.0
	Min, Max	0, 100	10, 69	0, 84		0, 100

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages for number of subjects with a cough are calculated using N in the column header.

Note: The assessments performed at Day 0 to Day 6 are used to evaluate cough at Day -1 to Day 5, respectively.

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**Table 15.2.6.20.1 Summary of Cough Assessments by Study Day - Safety Population**

Study Day/ Variable	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled not randomized (N=9)	Overall Safety (N=169)
Intensity of cough						
Very mild	n (%)	11 (13.8%)	7 (17.1%)	5 (12.8%)		23 (13.6%)
Mild	n (%)	18 (22.5%)	6 (14.6%)	5 (12.8%)		29 (17.2%)
Moderate	n (%)	7 (8.8%)	2 (4.9%)	4 (10.3%)		13 (7.7%)
Severe	n (%)	2 (2.5%)	0	0		2 (1.2%)
Very severe	n (%)	0	0	0		0
Frequency of cough						
Rarely	n (%)	19 (23.8%)	4 (9.8%)	7 (17.9%)		30 (17.8%)
Sometimes	n (%)	17 (21.3%)	10 (24.4%)	5 (12.8%)		32 (18.9%)
Fairly often	n (%)	1 (1.3%)	1 (2.4%)	2 (5.1%)		4 (2.4%)
Often	n (%)	1 (1.3%)	0	0		1 (0.6%)
Almost always	n (%)	0	0	0		0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages for number of subjects with a cough are calculated using N in the column header.

Note: The assessments performed at Day 0 to Day 6 are used to evaluate cough at Day -1 to Day 5, respectively.

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**Table 15.2.6.20.1 Summary of Cough Assessments by Study Day - Safety Population**

Study Day/ Variable	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled not randomized (N=9)	Overall Safety (N=169)
Amount of sputum produced						
No sputum	n (%)	22 (27.5%)	6 (14.6%)	9 (23.1%)		37 (21.9%)
A moderate amount of sputum	n (%)	16 (20.0%)	9 (22.0%)	5 (12.8%)		30 (17.8%)
A large amount of sputum	n (%)	0	0	0		0
A very large amount of sputum	n (%)	0	0	0		0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages for number of subjects with a cough are calculated using N in the column header.

Note: The assessments performed at Day 0 to Day 6 are used to evaluate cough at Day -1 to Day 5, respectively.

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**Table 15.2.6.20.1 Summary of Cough Assessments by Study Day - Safety Population**

Study Day/ Variable	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled not randomized (N=9)	Overall Safety (N=169)
Day 3						
Has subject experienced cough in last 24 hours						
Yes	n (%)	38 (47.5%)	16 (39.0%)	10 (25.6%)	0	64 (37.9%)
No	n (%)	41 (51.3%)	25 (61.0%)	29 (74.4%)	0	95 (56.2%)
VAS	n	38	16	10		64
	Mean (SD)	28.2 (22.80)	30.3 (17.79)	23.0 (21.47)		27.9 (21.26)
	Median	22.5	25.0	16.5		23.0
	Min, Max	0, 91	8, 63	2, 62		0, 91

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages for number of subjects with a cough are calculated using N in the column header.

Note: The assessments performed at Day 0 to Day 6 are used to evaluate cough at Day -1 to Day 5, respectively.

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**Table 15.2.6.20.1 Summary of Cough Assessments by Study Day - Safety Population**

Study Day/ Variable	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled not randomized (N=9)	Overall Safety (N=169)
Intensity of cough						
Very mild	n (%)	18 (22.5%)	5 (12.2%)	5 (12.8%)		28 (16.6%)
Mild	n (%)	8 (10.0%)	9 (22.0%)	3 (7.7%)		20 (11.8%)
Moderate	n (%)	11 (13.8%)	1 (2.4%)	2 (5.1%)		14 (8.3%)
Severe	n (%)	1 (1.3%)	1 (2.4%)	0		2 (1.2%)
Very severe	n (%)	0	0	0		0
Frequency of cough						
Rarely	n (%)	21 (26.3%)	9 (22.0%)	5 (12.8%)		35 (20.7%)
Sometimes	n (%)	12 (15.0%)	5 (12.2%)	4 (10.3%)		21 (12.4%)
Fairly often	n (%)	5 (6.3%)	2 (4.9%)	1 (2.6%)		8 (4.7%)
Often	n (%)	0	0	0		0
Almost always	n (%)	0	0	0		0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages for number of subjects with a cough are calculated using N in the column header.

Note: The assessments performed at Day 0 to Day 6 are used to evaluate cough at Day -1 to Day 5, respectively.

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**Table 15.2.6.20.1 Summary of Cough Assessments by Study Day - Safety Population**

Study Day/ Variable	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled not randomized (N=9)	Overall Safety (N=169)
Amount of sputum produced						
No sputum	n (%)	18 (22.5%)	11 (26.8%)	6 (15.4%)		35 (20.7%)
A moderate amount of sputum	n (%)	20 (25.0%)	5 (12.2%)	4 (10.3%)		29 (17.2%)
A large amount of sputum	n (%)	0	0	0		0
A very large amount of sputum	n (%)	0	0	0		0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages for number of subjects with a cough are calculated using N in the column header.

Note: The assessments performed at Day 0 to Day 6 are used to evaluate cough at Day -1 to Day 5, respectively.

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**Table 15.2.6.20.1 Summary of Cough Assessments by Study Day - Safety Population**

Study Day/ Variable	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled not randomized (N=9)	Overall Safety (N=169)
Day 4						
Has subject experienced cough in last 24 hours						
Yes	n (%)	29 (36.3%)	17 (41.5%)	10 (25.6%)	0	56 (33.1%)
No	n (%)	50 (62.5%)	24 (58.5%)	29 (74.4%)	0	103 (60.9%)
VAS						
	n	29	17	10		56
	Mean (SD)	24.6 (18.96)	26.6 (17.75)	26.3 (24.70)		25.5 (19.38)
	Median	21.0	21.0	15.5		20.5
	Min, Max	0, 76	9, 68	3, 79		0, 79

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages for number of subjects with a cough are calculated using N in the column header.

Note: The assessments performed at Day 0 to Day 6 are used to evaluate cough at Day -1 to Day 5, respectively.

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**Table 15.2.6.20.1 Summary of Cough Assessments by Study Day - Safety Population**

Study Day/ Variable	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled not randomized (N=9)	Overall Safety (N=169)
Intensity of cough						
Very mild	n (%)	11 (13.8%)	8 (19.5%)	4 (10.3%)		23 (13.6%)
Mild	n (%)	13 (16.3%)	4 (9.8%)	5 (12.8%)		22 (13.0%)
Moderate	n (%)	4 (5.0%)	4 (9.8%)	1 (2.6%)		9 (5.3%)
Severe	n (%)	1 (1.3%)	1 (2.4%)	0		2 (1.2%)
Very severe	n (%)	0	0	0		0
Frequency of cough						
Rarely	n (%)	14 (17.5%)	10 (24.4%)	4 (10.3%)		28 (16.6%)
Sometimes	n (%)	13 (16.3%)	6 (14.6%)	6 (15.4%)		25 (14.8%)
Fairly often	n (%)	2 (2.5%)	1 (2.4%)	0		3 (1.8%)
Often	n (%)	0	0	0		0
Almost always	n (%)	0	0	0		0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages for number of subjects with a cough are calculated using N in the column header.

Note: The assessments performed at Day 0 to Day 6 are used to evaluate cough at Day -1 to Day 5, respectively.

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**Table 15.2.6.20.1 Summary of Cough Assessments by Study Day - Safety Population**

Study Day/ Variable	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled not randomized (N=9)	Overall Safety (N=169)
Amount of sputum produced						
No sputum	n (%)	16 (20.0%)	9 (22.0%)	6 (15.4%)		31 (18.3%)
A moderate amount of sputum	n (%)	13 (16.3%)	8 (19.5%)	4 (10.3%)		25 (14.8%)
A large amount of sputum	n (%)	0	0	0		0
A very large amount of sputum	n (%)	0	0	0		0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages for number of subjects with a cough are calculated using N in the column header.

Note: The assessments performed at Day 0 to Day 6 are used to evaluate cough at Day -1 to Day 5, respectively.

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**Table 15.2.6.20.1 Summary of Cough Assessments by Study Day - Safety Population**

Study Day/ Variable	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled not randomized (N=9)	Overall Safety (N=169)
Day 5						
Has subject experienced cough in last 24 hours						
Yes	n (%)	28 (35.0%)	18 (43.9%)	9 (23.1%)	0	55 (32.5%)
No	n (%)	51 (63.8%)	23 (56.1%)	30 (76.9%)	0	104 (61.5%)
VAS						
	n	28	18	9		55
	Mean (SD)	33.4 (31.03)	30.6 (22.84)	30.7 (23.03)		32.1 (26.95)
	Median	24.5	20.0	32.0		22.0
	Min, Max	0, 100	3, 75	2, 68		0, 100

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages for number of subjects with a cough are calculated using N in the column header.

Note: The assessments performed at Day 0 to Day 6 are used to evaluate cough at Day -1 to Day 5, respectively.

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**Table 15.2.6.20.1 Summary of Cough Assessments by Study Day - Safety Population**

Study Day/ Variable	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled not randomized (N=9)	Overall Safety (N=169)
Intensity of cough						
Very mild	n (%)	10 (12.5%)	7 (17.1%)	3 (7.7%)		20 (11.8%)
Mild	n (%)	13 (16.3%)	6 (14.6%)	6 (15.4%)		25 (14.8%)
Moderate	n (%)	4 (5.0%)	3 (7.3%)	0		7 (4.1%)
Severe	n (%)	1 (1.3%)	2 (4.9%)	0		3 (1.8%)
Very severe	n (%)	0	0	0		0
Frequency of cough						
Rarely	n (%)	11 (13.8%)	11 (26.8%)	3 (7.7%)		25 (14.8%)
Sometimes	n (%)	15 (18.8%)	5 (12.2%)	5 (12.8%)		25 (14.8%)
Fairly often	n (%)	2 (2.5%)	1 (2.4%)	1 (2.6%)		4 (2.4%)
Often	n (%)	0	1 (2.4%)	0		1 (0.6%)
Almost always	n (%)	0	0	0		0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages for number of subjects with a cough are calculated using N in the column header.

Note: The assessments performed at Day 0 to Day 6 are used to evaluate cough at Day -1 to Day 5, respectively.

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**Table 15.2.6.20.1 Summary of Cough Assessments by Study Day - Safety Population**

Study Day/ Variable	Statistic	THS 2.2 (N=80)	CC (N=41)	SA (N=39)	Enrolled not randomized (N=9)	Overall Safety (N=169)
Amount of sputum produced						
No sputum	n (%)	14 (17.5%)	7 (17.1%)	3 (7.7%)		24 (14.2%)
A moderate amount of sputum	n (%)	13 (16.3%)	10 (24.4%)	6 (15.4%)		29 (17.2%)
A large amount of sputum	n (%)	1 (1.3%)	1 (2.4%)	0		2 (1.2%)
A very large amount of sputum	n (%)	0	0	0		0

Note: CC = Conventional cigarettes; SA = Smoking abstinence; THS = Tobacco Heating System.

Note: Enrolled Not Randomized refers to all subjects enrolled but not randomized. Overall Safety refers to enrolled subjects exposed to THS 2.2.

Note: Percentages for number of subjects with a cough are calculated using N in the column header.

Note: The assessments performed at Day 0 to Day 6 are used to evaluate cough at Day -1 to Day 5, respectively.

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