



## PMI RESEARCH & DEVELOPMENT

### **Study ZRHM-PK-05-JP Clinical Study Report Appendix 15.2 Tables**

<b>Study Title:</b>	A single-center, open-label, randomized, controlled, crossover study to investigate the nicotine pharmacokinetic profile and safety of Tobacco Heating System 2.2 Menthol (THS 2.2 Menthol) following single use in smoking, healthy subjects compared to menthol conventional cigarettes and nicotine gum
<b>Short Title:</b>	Nicotine pharmacokinetic profile and safety of Tobacco Heating System 2.2 Menthol (THS 2.2 Menthol)
<b>Study Number:</b>	ZRHM-PK-05-JP
<b>Product Name:</b>	Tobacco Heating System 2.2 Menthol (THS 2.2 Menthol)
<b>Study Initiated (first subject screened):</b>	01 August 2013
<b>Study Completed (last subject last visit):</b>	16 November 2013
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<b>Version:</b>	1.0
<b>Date:</b>	12 May 2015

This study was conducted in accordance with Good Clinical Practice.

#### **Confidentiality Statement**

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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	Sequence								Overall Screened
	THS 2.2 Menthol - mCC (N=22)		mCC - THS 2.2 Menthol (N=22)		THS 2.2 Menthol - NRT gum (N=9)		NRT gum - THS 2.2 Menthol (N=9)		
Total screened - n									147
Screen failures - n (%)									74 ( 50.3%)
Screening failures, without product test - n (%)									74 ( 50.3%)
Screening failures, with product test - n (%)									0
Enrolled - n (%)									73 ( 49.7%)
Enrolled, not randomized - n (%)									11 ( 7.5%)
Randomized - n (%)	22	( 100 %)	22	( 100 %)	9	( 100 %)	9	( 100 %)	62 ( 42.2%)
Total randomized - n	22		22		9		9		62
Completed - n (%)	21	( 95.5%)	22	( 100 %)	9	( 100 %)	9	( 100 %)	61 ( 98.4%)
Discontinued - n (%)	1	( 4.5%)	0		0		0		1 ( 1.6%)
Discontinued, with randomized product use	1	( 4.5%)							1 ( 1.6%)

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Note: Enrolled subjects are those who are eligible on Day -1; Discontinued refers to randomized subjects who discontinued the study before the planned discharge at Day 4; Completed refers to randomized subjects who did not discontinue the study before the planned discharge at Day 4.

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/000000106326/dev/tables/t\_sdisp.sas

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

	Sequence				Overall Randomized (N=62)
	THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2. Menthol (N=9)	
Total no. of discontinuations - n (%)	1 ( 4.5%)	0	0	0	1 ( 1.6%)
Reason for discontinuation					
Adverse events - n (%)	0				0
Protocol violation - n (%)	0				0
Withdrawal by subject - n (%)	1 ( 4.5%)				1 ( 1.6%)
Lost to follow-up - n (%)	0				0
Other - n (%)	0				0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Appendix 15.3.1.7

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_rsndis.sas

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

Significance/ Classification of Deviation	Sequence										Overall Safety (N=73)
	THS 2.2 Menthol - mCC (N=22)		mCC - THS 2.2 Menthol (N=22)		THS 2.2 Menthol - NRT gum (N=9)		NRT gum - THS 2.2 Menthol (N=9)		Exposed Not Randomized (N=11)		
	n	(%)	n	(%)	n	(%)	n	(%)	n	(%)	
	n	(%)	n	(%)	n	(%)	n	(%)	n	(%)	
Major	0		0		0		0		1 ( 9.1%)		1 ( 1.4%)
Protocol violation									1 ( 9.1%)		1 ( 1.4%)
Minor	22 (100 %)		22 (100 %)		9 (100 %)		9 (100 %)		11 (100 %)		73 (100 %)
Time schedule deviation	21 ( 95.5%)		22 (100 %)		9 (100 %)		9 (100 %)		0		61 ( 83.6%)
Procedure violation	5 ( 22.7%)		3 ( 13.6%)		2 ( 22.2%)		2 ( 22.2%)		11 (100 %)		23 ( 31.5%)

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Note: Exposed not randomized refers to all subjects exposed to THS 2.2 Menthol or NRT gum but not randomized. Overall Safety refers to all subjects exposed to THS 2.2 Menthol or NRT gum.

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

Appendix 15.3.1.11

Path: /cvn/projects/prj/development/000000106326/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	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
		THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Sex							
Male	n (%)	11 ( 50.0%)	12 ( 54.5%)	5 ( 55.6%)	4 ( 44.4%)	5 ( 45.5%)	37 ( 50.7%)
Female	n (%)	11 ( 50.0%)	10 ( 45.5%)	4 ( 44.4%)	5 ( 55.6%)	6 ( 54.5%)	36 ( 49.3%)
ISO nicotine level							
<= 0.6 mg	n (%)	14 ( 63.6%)	12 ( 54.5%)	5 ( 55.6%)	5 ( 55.6%)	8 ( 72.7%)	44 ( 60.3%)
> 0.6 - 1.0 mg	n (%)	8 ( 36.4%)	10 ( 45.5%)	4 ( 44.4%)	4 ( 44.4%)	3 ( 27.3%)	29 ( 39.7%)
Age (years)	n	22	22	9	9	11	73
	Mean (SD)	35.9 (11.93)	31.4 (7.38)	29.3 (7.50)	32.1 (8.29)	34.8 (10.68)	33.1 (9.63)
	Median	31.5	31.0	28.0	29.0	33.0	30.0
	Min, Max	23, 61	23, 44	23, 44	23, 44	24, 52	23, 61

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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, 15.3.1.10

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_demog.sas

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

Variable	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
		THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
BMI (kg/m <sup>2</sup> )	n	22	22	9	9	11	73
	Mean (SD)	22.21 (1.410)	22.46 (2.962)	23.84 (2.608)	21.87 (2.120)	21.78 (2.332)	22.38 (2.349)
	Median	22.18	21.50	23.38	21.91	21.24	22.21
	Min, Max	19.3, 24.5	18.7, 30.4	19.1, 28.4	18.8, 26.0	19.5, 26.7	18.7, 30.4
BMI classifications							
Underweight	n (%)	0	0	0	0	0	0
Normal weight	n (%)	22 (100 %)	18 ( 81.8%)	7 ( 77.8%)	8 ( 88.9%)	10 ( 90.9%)	65 ( 89.0%)
Overweight	n (%)	0	3 ( 13.6%)	2 ( 22.2%)	1 ( 11.1%)	1 ( 9.1%)	7 ( 9.6%)
Obese	n (%)	0	1 ( 4.5%)	0	0	0	1 ( 1.4%)

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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, 15.3.1.10

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

Variable	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
		THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Height (m)	n	22	22	9	9	11	73
	Mean (SD)	1.668 (0.0773)	1.661 (0.0715)	1.661 (0.1129)	1.640 (0.0764)	1.624 (0.0870)	1.655 (0.0812)
	Median	1.655	1.675	1.670	1.650	1.590	1.650
	Min, Max	1.54, 1.80	1.53, 1.76	1.50, 1.86	1.54, 1.76	1.50, 1.76	1.50, 1.86
Weight (kg)	n	22	22	9	9	11	73
	Mean (SD)	61.85 (6.450)	62.08 (9.310)	66.11 (11.702)	59.07 (9.221)	57.50 (8.329)	61.44 (8.814)
	Median	60.95	63.35	67.50	55.10	53.80	60.80
	Min, Max	49.4, 77.5	46.0, 78.7	51.9, 82.2	48.2, 75.0	47.4, 75.5	46.0, 82.2

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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, 15.3.1.10

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

Variable	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
		THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Ethnicity							
Japanese	n (%)	22 (100 %)	22 (100 %)	9 (100 %)	9 (100 %)	11 (100 %)	73 (100 %)
Not Japanese	n (%)	0	0	0	0	0	0
Daily mCC consumption at Admission							
10-19 cig/day	n (%)	14 ( 63.6%)	11 ( 50.0%)	5 ( 55.6%)	6 ( 66.7%)	8 ( 72.7%)	44 ( 60.3%)
>19 cig/day	n (%)	8 ( 36.4%)	11 ( 50.0%)	4 ( 44.4%)	3 ( 33.3%)	3 ( 27.3%)	29 ( 39.7%)
mCC nicotine yield at Admission (mg)	n	22	22	9	9	11	73
	Mean (SD)	0.41 (0.317)	0.51 (0.261)	0.50 (0.250)	0.42 (0.282)	0.41 (0.321)	0.45 (0.286)
	Median	0.40	0.60	0.60	0.40	0.40	0.60
	Min, Max	0.1, 0.9	0.1, 0.8	0.1, 0.7	0.1, 0.7	0.1, 0.9	0.1, 0.9

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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, 15.3.1.10

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

Variable	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
		THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
mCC ISO tar yield at Admission (mg)	n	22	22	9	9	11	73
	Mean (SD)	5.3 (4.58)	6.5 (3.57)	6.0 (3.25)	5.1 (3.56)	5.3 (4.52)	5.7 (3.95)
	Median	5.0	8.0	8.0	5.0	5.0	8.0
	Min, Max	1, 12	1, 12	1, 10	1, 10	1, 12	1, 12
mCC ISO tar yield at Admission							
1-5 mg	n (%)	13 ( 59.1%)	7 ( 31.8%)	4 ( 44.4%)	5 ( 55.6%)	6 ( 54.5%)	35 ( 47.9%)
6-8 mg	n (%)	4 ( 18.2%)	11 ( 50.0%)	4 ( 44.4%)	3 ( 33.3%)	3 ( 27.3%)	25 ( 34.2%)
9-10 mg	n (%)	0	2 ( 9.1%)	1 ( 11.1%)	1 ( 11.1%)	0	4 ( 5.5%)
>10 mg	n (%)	5 ( 22.7%)	2 ( 9.1%)	0	0	2 ( 18.2%)	9 ( 12.3%)

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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, 15.3.1.10

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

Variable	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
		THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
CYP2A6 activity (%)	n	22	22	9	9	10	72
	Mean (SD)	26.903 (21.0556)	26.688 (19.7770)	32.447 (25.1195)	21.271 (16.4927)	36.050 (32.5216)	28.097 (22.3808)
	Median	19.642	23.467	22.597	14.166	31.790	20.631
	Min, Max	4.19, 76.88	4.55, 79.41	4.72, 68.99	7.88, 57.96	8.72, 120.90	4.19, 120.90
FTND total score classifications							
Mild	n (%)	8 ( 36.4%)	10 ( 45.5%)	4 ( 44.4%)	5 ( 55.6%)	4 ( 36.4%)	31 ( 42.5%)
Moderate	n (%)	12 ( 54.5%)	11 ( 50.0%)	4 ( 44.4%)	3 ( 33.3%)	6 ( 54.5%)	36 ( 49.3%)
Severe	n (%)	2 ( 9.1%)	1 ( 4.5%)	1 ( 11.1%)	1 ( 11.1%)	1 ( 9.1%)	6 ( 8.2%)

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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, 15.3.1.10

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

Variable	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
		THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
FTND total score	n	22	22	9	9	11	73
	Mean (SD)	4.0 (2.06)	4.1 (1.45)	3.9 (1.70)	3.3 (2.13)	3.9 (1.65)	3.9 (1.76)
	Median	4.0	4.0	4.0	3.0	4.0	4.0
	Min, Max	1, 10	1, 7	2, 7	1, 8	1, 7	1, 10

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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, 15.3.1.10

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**Table 15.2.1.4.2 Summary of Demographics and Other Baseline Characteristics - Group-1 PK Population**

Variable	Statistic	THS 2.2 Menthol (N=43)	mCC (N=43)	Group-1 PK (N=43)	Overall PK (N=61)
Sex					
Male	n (%)	23 ( 53.5%)	23 ( 53.5%)	23 ( 53.5%)	32 ( 52.5%)
Female	n (%)	20 ( 46.5%)	20 ( 46.5%)	20 ( 46.5%)	29 ( 47.5%)
ISO nicotine level					
<= 0.6 mg	n (%)	25 ( 58.1%)	25 ( 58.1%)	25 ( 58.1%)	35 ( 57.4%)
> 0.6 - 1.0 mg	n (%)	18 ( 41.9%)	18 ( 41.9%)	18 ( 41.9%)	26 ( 42.6%)
Age (years)	n	43	43	43	61
	Mean (SD)	33.4 (10.03)	33.4 (10.03)	33.4 (10.03)	32.6 (9.44)
	Median	31.0	31.0	31.0	29.0
	Min, Max	23, 61	23, 61	23, 61	23, 61

Note: mCC = menthol conventional cigarettes; 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, 15.3.1.10

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_demog2.sas

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**Table 15.2.1.4.2 Summary of Demographics and Other Baseline Characteristics - Group-1 PK Population**

Variable	Statistic	THS 2.2 Menthol (N=43)	mCC (N=43)	Group-1 PK (N=43)	Overall PK (N=61)
BMI (kg/m <sup>2</sup> )	n	43	43	43	61
	Mean (SD)	22.34 (2.324)	22.34 (2.324)	22.34 (2.324)	22.49 (2.374)
	Median	22.10	22.10	22.10	22.28
	Min, Max	18.7, 30.4	18.7, 30.4	18.7, 30.4	18.7, 30.4
BMI classifications					
Underweight	n (%)	0	0	0	0
Normal weight	n (%)	39 ( 90.7%)	39 ( 90.7%)	39 ( 90.7%)	54 ( 88.5%)
Overweight	n (%)	3 ( 7.0%)	3 ( 7.0%)	3 ( 7.0%)	6 ( 9.8%)
Obese	n (%)	1 ( 2.3%)	1 ( 2.3%)	1 ( 2.3%)	1 ( 1.6%)

Note: mCC = menthol conventional cigarettes; 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, 15.3.1.10

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**Table 15.2.1.4.2 Summary of Demographics and Other Baseline Characteristics - Group-1 PK Population**

Variable	Statistic	THS 2.2 Menthol (N=43)	mCC (N=43)	Group-1 PK (N=43)	Overall PK (N=61)
Height (m)	n	43	43	43	61
	Mean (SD)	1.666 (0.0742)	1.666 (0.0742)	1.666 (0.0742)	1.661 (0.0801)
	Median	1.670	1.670	1.670	1.670
	Min, Max	1.53, 1.80	1.53, 1.80	1.53, 1.80	1.50, 1.86
Weight (kg)	n	43	43	43	61
	Mean (SD)	62.05 (7.989)	62.05 (7.989)	62.05 (7.989)	62.21 (8.835)
	Median	61.80	61.80	61.80	61.80
	Min, Max	46.0, 78.7	46.0, 78.7	46.0, 78.7	46.0, 82.2
Ethnicity					
Japanese	n (%)	43 (100 %)	43 (100 %)	43 (100 %)	61 (100 %)
Not Japanese	n (%)	0	0	0	0

Note: mCC = menthol conventional cigarettes; 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, 15.3.1.10

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**Table 15.2.1.4.2 Summary of Demographics and Other Baseline Characteristics - Group-1 PK Population**

Variable	Statistic	THS 2.2 Menthol (N=43)	mCC (N=43)	Group-1 PK (N=43)	Overall PK (N=61)
Daily mCC consumption at Admission					
10-19 cig/day	n (%)	25 ( 58.1%)	25 ( 58.1%)	25 ( 58.1%)	36 ( 59.0%)
>19 cig/day	n (%)	18 ( 41.9%)	18 ( 41.9%)	18 ( 41.9%)	25 ( 41.0%)
mCC nicotine yield at Admission (mg)	n	43	43	43	61
	Mean (SD)	0.47 (0.290)	0.47 (0.290)	0.47 (0.290)	0.47 (0.280)
	Median	0.60	0.60	0.60	0.60
	Min, Max	0.1, 0.9	0.1, 0.9	0.1, 0.9	0.1, 0.9

Note: mCC = menthol conventional cigarettes; 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, 15.3.1.10

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_demog2.sas

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**Table 15.2.1.4.2 Summary of Demographics and Other Baseline Characteristics - Group-1 PK Population**

Variable	Statistic	THS 2.2 Menthol (N=43)	mCC (N=43)	Group-1 PK (N=43)	Overall PK (N=61)
mCC ISO tar yield at Admission (mg)	n	43	43	43	61
	Mean (SD)	6.0 (4.08)	6.0 (4.08)	6.0 (4.08)	5.9 (3.86)
	Median	8.0	8.0	8.0	8.0
	Min, Max	1, 12	1, 12	1, 12	1, 12
mCC ISO tar yield at Admission					
1-5 mg	n (%)	19 ( 44.2%)	19 ( 44.2%)	19 ( 44.2%)	28 ( 45.9%)
6-8 mg	n (%)	15 ( 34.9%)	15 ( 34.9%)	15 ( 34.9%)	22 ( 36.1%)
9-10 mg	n (%)	2 ( 4.7%)	2 ( 4.7%)	2 ( 4.7%)	4 ( 6.6%)
>10 mg	n (%)	7 ( 16.3%)	7 ( 16.3%)	7 ( 16.3%)	7 ( 11.5%)

Note: mCC = menthol conventional cigarettes; 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, 15.3.1.10

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**Table 15.2.1.4.2 Summary of Demographics and Other Baseline Characteristics - Group-1 PK Population**

Variable	Statistic	THS 2.2 Menthol (N=43)	mCC (N=43)	Group-1 PK (N=43)	Overall PK (N=61)
CYP2A6 activity (%)	n	43	43	43	61
	Mean (SD)	26.980 (20.3891)	26.980 (20.3891)	26.980 (20.3891)	26.944 (20.5127)
	Median	21.019	21.019	21.019	20.530
	Min, Max	4.19, 79.41	4.19, 79.41	4.19, 79.41	4.19, 79.41
FTND total score classifications					
Mild	n (%)	18 ( 41.9%)	18 ( 41.9%)	18 ( 41.9%)	27 ( 44.3%)
Moderate	n (%)	23 ( 53.5%)	23 ( 53.5%)	23 ( 53.5%)	30 ( 49.2%)
Severe	n (%)	2 ( 4.7%)	2 ( 4.7%)	2 ( 4.7%)	4 ( 6.6%)
FTND total score					
	n	43	43	43	61
	Mean (SD)	3.9 (1.52)	3.9 (1.52)	3.9 (1.52)	3.8 (1.63)
	Median	4.0	4.0	4.0	4.0
	Min, Max	1, 7	1, 7	1, 7	1, 8

Note: mCC = menthol conventional cigarettes; 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, 15.3.1.10

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**Table 15.2.1.4.2.1 Summary of Demographics and Other Baseline Characteristics by Sex - Group-1 PK Population**

Variable	Statistic	THS 2.2 Menthol		mCC		Group-1 PK	
		Males (N=23)	Females (N=20)	Males (N=23)	Females (N=20)	Males (N=23)	Females (N=20)
ISO nicotine level							
<= 0.6 mg	n (%)	12 ( 52.2%)	13 ( 65.0%)	12 ( 52.2%)	13 ( 65.0%)	12 ( 52.2%)	13 ( 65.0%)
> 0.6 - 1.0 mg	n (%)	11 ( 47.8%)	7 ( 35.0%)	11 ( 47.8%)	7 ( 35.0%)	11 ( 47.8%)	7 ( 35.0%)
Age (years)	n	23	20	23	20	23	20
	Mean (SD)	32.3 (10.39)	34.6 (9.72)	32.3 (10.39)	34.6 (9.72)	32.3 (10.39)	34.6 (9.72)
	Median	30.0	32.5	30.0	32.5	30.0	32.5
	Min, Max	23, 61	23, 61	23, 61	23, 61	23, 61	23, 61
BMI (kg/m <sup>2</sup> )	n	23	20	23	20	23	20
	Mean (SD)	22.07 (1.637)	22.65 (2.939)	22.07 (1.637)	22.65 (2.939)	22.07 (1.637)	22.65 (2.939)
	Median	21.96	22.66	21.96	22.66	21.96	22.66
	Min, Max	19.3, 25.1	18.7, 30.4	19.3, 25.1	18.7, 30.4	19.3, 25.1	18.7, 30.4

Note: mCC = menthol conventional cigarettes; 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, 15.3.1.10

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**Table 15.2.1.4.2.1 Summary of Demographics and Other Baseline Characteristics by Sex - Group-1 PK Population**

Variable	Statistic	THS 2.2 Menthol		mCC		Group-1 PK	
		Males (N=23)	Females (N=20)	Males (N=23)	Females (N=20)	Males (N=23)	Females (N=20)
BMI classifications							
Underweight	n (%)	0	0	0	0	0	0
Normal weight	n (%)	22 ( 95.7%)	17 ( 85.0%)	22 ( 95.7%)	17 ( 85.0%)	22 ( 95.7%)	17 ( 85.0%)
Overweight	n (%)	1 ( 4.3%)	2 ( 10.0%)	1 ( 4.3%)	2 ( 10.0%)	1 ( 4.3%)	2 ( 10.0%)
Obese	n (%)	0	1 ( 5.0%)	0	1 ( 5.0%)	0	1 ( 5.0%)
Height (m)	n	23	20	23	20	23	20
	Mean (SD)	1.716 (0.0440)	1.608 (0.0581)	1.716 (0.0440)	1.608 (0.0581)	1.716 (0.0440)	1.608 (0.0581)
	Median	1.710	1.600	1.710	1.600	1.710	1.600
	Min, Max	1.64, 1.80	1.53, 1.75	1.64, 1.80	1.53, 1.75	1.64, 1.80	1.53, 1.75

Note: mCC = menthol conventional cigarettes; 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, 15.3.1.10

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**Table 15.2.1.4.2.1 Summary of Demographics and Other Baseline Characteristics by Sex - Group-1 PK Population**

Variable	Statistic	THS 2.2 Menthol		mCC		Group-1 PK	
		Males (N=23)	Females (N=20)	Males (N=23)	Females (N=20)	Males (N=23)	Females (N=20)
Weight (kg)	n	23	20	23	20	23	20
	Mean (SD)	64.94 (5.272)	58.72 (9.330)	64.94 (5.272)	58.72 (9.330)	64.94 (5.272)	58.72 (9.330)
	Median	64.20	55.65	64.20	55.65	64.20	55.65
	Min, Max	54.4, 77.5	46.0, 78.7	54.4, 77.5	46.0, 78.7	54.4, 77.5	46.0, 78.7
Ethnicity							
Japanese	n (%)	23 (100 %)	20 (100 %)	23 (100 %)	20 (100 %)	23 (100 %)	20 (100 %)
Not Japanese	n (%)	0	0	0	0	0	0
Daily mCC consumption at Admission							
10-19 cig/day	n (%)	13 ( 56.5%)	12 ( 60.0%)	13 ( 56.5%)	12 ( 60.0%)	13 ( 56.5%)	12 ( 60.0%)
>19 cig/day	n (%)	10 ( 43.5%)	8 ( 40.0%)	10 ( 43.5%)	8 ( 40.0%)	10 ( 43.5%)	8 ( 40.0%)

Note: mCC = menthol conventional cigarettes; 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, 15.3.1.10

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**Table 15.2.1.4.2.1 Summary of Demographics and Other Baseline Characteristics by Sex - Group-1 PK Population**

Variable	Statistic	THS 2.2 Menthol		mCC		Group-1 PK	
		Males (N=23)	Females (N=20)	Males (N=23)	Females (N=20)	Males (N=23)	Females (N=20)
mCC nicotine yield at Admission (mg)	n	23	20	23	20	23	20
	Mean (SD)	0.55 (0.261)	0.38 (0.298)	0.55 (0.261)	0.38 (0.298)	0.55 (0.261)	0.38 (0.298)
	Median	0.60	0.30	0.60	0.30	0.60	0.30
	Min, Max	0.1, 0.8	0.1, 0.9	0.1, 0.8	0.1, 0.9	0.1, 0.8	0.1, 0.9
mCC ISO tar yield at Admission (mg)	n	23	20	23	20	23	20
	Mean (SD)	7.2 (3.87)	4.7 (3.99)	7.2 (3.87)	4.7 (3.99)	7.2 (3.87)	4.7 (3.99)
	Median	8.0	4.5	8.0	4.5	8.0	4.5
	Min, Max	1, 12	1, 12	1, 12	1, 12	1, 12	1, 12
mCC ISO tar yield at Admission							
1-5 mg	n (%)	6 ( 26.1%)	13 ( 65.0%)	6 ( 26.1%)	13 ( 65.0%)	6 ( 26.1%)	13 ( 65.0%)
6-8 mg	n (%)	11 ( 47.8%)	4 ( 20.0%)	11 ( 47.8%)	4 ( 20.0%)	11 ( 47.8%)	4 ( 20.0%)
9-10 mg	n (%)	1 ( 4.3%)	1 ( 5.0%)	1 ( 4.3%)	1 ( 5.0%)	1 ( 4.3%)	1 ( 5.0%)
>10 mg	n (%)	5 ( 21.7%)	2 ( 10.0%)	5 ( 21.7%)	2 ( 10.0%)	5 ( 21.7%)	2 ( 10.0%)

Note: mCC = menthol conventional cigarettes; 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, 15.3.1.10

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**Table 15.2.1.4.2.1 Summary of Demographics and Other Baseline Characteristics by Sex - Group-1 PK Population**

Variable	Statistic	THS 2.2 Menthol		mCC		Group-1 PK	
		Males (N=23)	Females (N=20)	Males (N=23)	Females (N=20)	Males (N=23)	Females (N=20)
CYP2A6 activity (%)	n	23	20	23	20	23	20
	Mean (SD)	23.590 (19.3795)	30.877 (21.3116)	23.590 (19.3795)	30.877 (21.3116)	23.590 (19.3795)	30.877 (21.3116)
	Median	16.752	32.858	16.752	32.858	16.752	32.858
	Min, Max	4.19, 79.41	4.55, 76.88	4.19, 79.41	4.55, 76.88	4.19, 79.41	4.55, 76.88
FTND total score classifications							
Mild	n (%)	11 ( 47.8%)	7 ( 35.0%)	11 ( 47.8%)	7 ( 35.0%)	11 ( 47.8%)	7 ( 35.0%)
Moderate	n (%)	12 ( 52.2%)	11 ( 55.0%)	12 ( 52.2%)	11 ( 55.0%)	12 ( 52.2%)	11 ( 55.0%)
Severe	n (%)	0	2 ( 10.0%)	0	2 ( 10.0%)	0	2 ( 10.0%)
FTND total score	n	23	20	23	20	23	20
	Mean (SD)	3.8 (1.39)	4.1 (1.69)	3.8 (1.39)	4.1 (1.69)	3.8 (1.39)	4.1 (1.69)
	Median	4.0	4.0	4.0	4.0	4.0	4.0
	Min, Max	1, 6	1, 7	1, 6	1, 7	1, 6	1, 7

Note: mCC = menthol conventional cigarettes; 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, 15.3.1.10

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**Table 15.2.1.4.2.2 Summary of Demographics and Other Baseline Characteristics by Nicotine Level - Group-1 PK Population**

Variable	Statistic	THS 2.2 Menthol		mCC		Group-1 PK	
		<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)	<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)	<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)
Sex							
Male	n (%)	12 ( 48.0%)	11 ( 61.1%)	12 ( 48.0%)	11 ( 61.1%)	12 ( 48.0%)	11 ( 61.1%)
Female	n (%)	13 ( 52.0%)	7 ( 38.9%)	13 ( 52.0%)	7 ( 38.9%)	13 ( 52.0%)	7 ( 38.9%)
Age (years)	n	25	18	25	18	25	18
	Mean (SD)	34.8 (11.19)	31.4 (8.05)	34.8 (11.19)	31.4 (8.05)	34.8 (11.19)	31.4 (8.05)
	Median	31.0	28.5	31.0	28.5	31.0	28.5
	Min, Max	23, 61	23, 45	23, 61	23, 45	23, 61	23, 45
BMI (kg/m <sup>2</sup> )	n	25	18	25	18	25	18
	Mean (SD)	22.08 (2.032)	22.70 (2.696)	22.08 (2.032)	22.70 (2.696)	22.08 (2.032)	22.70 (2.696)
	Median	22.06	22.55	22.06	22.55	22.06	22.55
	Min, Max	19.3, 26.9	18.7, 30.4	19.3, 26.9	18.7, 30.4	19.3, 26.9	18.7, 30.4

Note: mCC = menthol conventional cigarettes; 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, 15.3.1.10

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**Table 15.2.1.4.2.2 Summary of Demographics and Other Baseline Characteristics by Nicotine Level - Group-1 PK Population**

Variable	Statistic	THS 2.2 Menthol		mCC		Group-1 PK	
		<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)	<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)	<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)
BMI classifications							
Underweight	n (%)	0	0	0	0	0	0
Normal weight	n (%)	23 ( 92.0%)	16 ( 88.9%)	23 ( 92.0%)	16 ( 88.9%)	23 ( 92.0%)	16 ( 88.9%)
Overweight	n (%)	2 ( 8.0%)	1 ( 5.6%)	2 ( 8.0%)	1 ( 5.6%)	2 ( 8.0%)	1 ( 5.6%)
Obese	n (%)	0	1 ( 5.6%)	0	1 ( 5.6%)	0	1 ( 5.6%)
Height (m)	n	25	18	25	18	25	18
	Mean (SD)	1.662 (0.0789)	1.671 (0.0689)	1.662 (0.0789)	1.671 (0.0689)	1.662 (0.0789)	1.671 (0.0689)
	Median	1.660	1.675	1.660	1.675	1.660	1.675
	Min, Max	1.53, 1.80	1.57, 1.79	1.53, 1.80	1.57, 1.79	1.53, 1.80	1.57, 1.79

Note: mCC = menthol conventional cigarettes; 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, 15.3.1.10

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**Table 15.2.1.4.2.2 Summary of Demographics and Other Baseline Characteristics by Nicotine Level - Group-1 PK Population**

Variable	Statistic	THS 2.2 Menthol		mCC		Group-1 PK	
		<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)	<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)	<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)
Weight (kg)	n	25	18	25	18	25	18
	Mean (SD)	61.02 (7.234)	63.48 (8.948)	61.02 (7.234)	63.48 (8.948)	61.02 (7.234)	63.48 (8.948)
	Median	60.60	64.15	60.60	64.15	60.60	64.15
	Min, Max	49.4, 78.1	46.0, 78.7	49.4, 78.1	46.0, 78.7	49.4, 78.1	46.0, 78.7
Ethnicity							
Japanese	n (%)	25 (100 %)	18 (100 %)	25 (100 %)	18 (100 %)	25 (100 %)	18 (100 %)
Not Japanese	n (%)	0	0	0	0	0	0
Daily mCC consumption at Admission							
10-19 cig/day	n (%)	15 ( 60.0%)	10 ( 55.6%)	15 ( 60.0%)	10 ( 55.6%)	15 ( 60.0%)	10 ( 55.6%)
>19 cig/day	n (%)	10 ( 40.0%)	8 ( 44.4%)	10 ( 40.0%)	8 ( 44.4%)	10 ( 40.0%)	8 ( 44.4%)

Note: mCC = menthol conventional cigarettes; 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, 15.3.1.10

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**Table 15.2.1.4.2.2 Summary of Demographics and Other Baseline Characteristics by Nicotine Level - Group-1 PK Population**

Variable	Statistic	THS 2.2 Menthol		mCC		Group-1 PK	
		<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)	<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)	<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)
mCC nicotine yield at Admission (mg)	n	25	18	25	18	25	18
	Mean (SD)	0.27 (0.216)	0.74 (0.062)	0.27 (0.216)	0.74 (0.062)	0.27 (0.216)	0.74 (0.062)
	Median	0.10	0.70	0.10	0.70	0.10	0.70
	Min, Max	0.1, 0.6	0.7, 0.9	0.1, 0.6	0.7, 0.9	0.1, 0.6	0.7, 0.9
mCC ISO tar yield at Admission (mg)	n	25	18	25	18	25	18
	Mean (SD)	3.4 (2.92)	9.8 (1.93)	3.4 (2.92)	9.8 (1.93)	3.4 (2.92)	9.8 (1.93)
	Median	1.0	9.0	1.0	9.0	1.0	9.0
	Min, Max	1, 8	8, 12	1, 8	8, 12	1, 8	8, 12

Note: mCC = menthol conventional cigarettes; 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, 15.3.1.10

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**Table 15.2.1.4.2.2 Summary of Demographics and Other Baseline Characteristics by Nicotine Level - Group-1 PK Population**

Variable	Statistic	THS 2.2 Menthol		mCC		Group-1 PK	
		<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)	<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)	<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)
mCC ISO tar yield at Admission							
1-5 mg	n (%)	19 ( 76.0%)	0	19 ( 76.0%)	0	19 ( 76.0%)	0
6-8 mg	n (%)	6 ( 24.0%)	9 ( 50.0%)	6 ( 24.0%)	9 ( 50.0%)	6 ( 24.0%)	9 ( 50.0%)
9-10 mg	n (%)	0	2 ( 11.1%)	0	2 ( 11.1%)	0	2 ( 11.1%)
>10 mg	n (%)	0	7 ( 38.9%)	0	7 ( 38.9%)	0	7 ( 38.9%)
CYP2A6 activity (%)	n	25	18	25	18	25	18
	Mean (SD)	32.076 (21.1849)	19.901 (17.3840)	32.076 (21.1849)	19.901 (17.3840)	32.076 (21.1849)	19.901 (17.3840)
	Median	33.574	15.566	33.574	15.566	33.574	15.566
	Min, Max	6.14, 76.88	4.19, 79.41	6.14, 76.88	4.19, 79.41	6.14, 76.88	4.19, 79.41

Note: mCC = menthol conventional cigarettes; 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, 15.3.1.10

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**Table 15.2.1.4.2.2 Summary of Demographics and Other Baseline Characteristics by Nicotine Level - Group-1 PK Population**

Variable	Statistic	THS 2.2 Menthol		mCC		Group-1 PK	
		<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)	<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)	<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)
FTND total score classifications							
Mild	n (%)	10 ( 40.0%)	8 ( 44.4%)	10 ( 40.0%)	8 ( 44.4%)	10 ( 40.0%)	8 ( 44.4%)
Moderate	n (%)	14 ( 56.0%)	9 ( 50.0%)	14 ( 56.0%)	9 ( 50.0%)	14 ( 56.0%)	9 ( 50.0%)
Severe	n (%)	1 ( 4.0%)	1 ( 5.6%)	1 ( 4.0%)	1 ( 5.6%)	1 ( 4.0%)	1 ( 5.6%)
FTND total score	n	25	18	25	18	25	18
	Mean (SD)	3.8 (1.73)	4.1 (1.22)	3.8 (1.73)	4.1 (1.22)	3.8 (1.73)	4.1 (1.22)
	Median	4.0	4.0	4.0	4.0	4.0	4.0
	Min, Max	1, 7	3, 7	1, 7	3, 7	1, 7	3, 7

Note: mCC = menthol conventional cigarettes; 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, 15.3.1.10

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_demog2c.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.1.4.3 Summary of Demographics and Other Baseline Characteristics - Group-2 PK Population**

Variable	Statistic	THS 2.2 Menthol (N=18)	NRT gum (N=18)	Group-2 PK (N=18)	Overall PK (N=61)
Sex					
Male	n (%)	9 ( 50.0%)	9 ( 50.0%)	9 ( 50.0%)	32 ( 52.5%)
Female	n (%)	9 ( 50.0%)	9 ( 50.0%)	9 ( 50.0%)	29 ( 47.5%)
ISO nicotine level					
<= 0.6 mg	n (%)	10 ( 55.6%)	10 ( 55.6%)	10 ( 55.6%)	35 ( 57.4%)
> 0.6 - 1.0 mg	n (%)	8 ( 44.4%)	8 ( 44.4%)	8 ( 44.4%)	26 ( 42.6%)
Age (years)	n	18	18	18	61
	Mean (SD)	30.7 (7.80)	30.7 (7.80)	30.7 (7.80)	32.6 (9.44)
	Median	28.0	28.0	28.0	29.0
	Min, Max	23, 44	23, 44	23, 44	23, 61

Note: NRT gum = Nicotine Replacement Therapy gum; 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, 15.3.1.10

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_demog3.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.1.4.3 Summary of Demographics and Other Baseline Characteristics - Group-2 PK Population**

Variable	Statistic	THS 2.2 Menthol (N=18)	NRT gum (N=18)	Group-2 PK (N=18)	Overall PK (N=61)
BMI (kg/m <sup>2</sup> )	n	18	18	18	61
	Mean (SD)	22.86 (2.520)	22.86 (2.520)	22.86 (2.520)	22.49 (2.374)
	Median	23.13	23.13	23.13	22.28
	Min, Max	18.8, 28.4	18.8, 28.4	18.8, 28.4	18.7, 30.4
BMI classifications					
Underweight	n (%)	0	0	0	0
Normal weight	n (%)	15 ( 83.3%)	15 ( 83.3%)	15 ( 83.3%)	54 ( 88.5%)
Overweight	n (%)	3 ( 16.7%)	3 ( 16.7%)	3 ( 16.7%)	6 ( 9.8%)
Obese	n (%)	0	0	0	1 ( 1.6%)

Note: NRT gum = Nicotine Replacement Therapy gum; 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, 15.3.1.10

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_demog3.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.1.4.3 Summary of Demographics and Other Baseline Characteristics - Group-2 PK Population**

Variable	Statistic	THS 2.2 Menthol (N=18)	NRT gum (N=18)	Group-2 PK (N=18)	Overall PK (N=61)
Height (m)	n	18	18	18	61
	Mean (SD)	1.651 (0.0941)	1.651 (0.0941)	1.651 (0.0941)	1.661 (0.0801)
	Median	1.660	1.660	1.660	1.670
	Min, Max	1.50, 1.86	1.50, 1.86	1.50, 1.86	1.50, 1.86
Weight (kg)	n	18	18	18	61
	Mean (SD)	62.59 (10.843)	62.59 (10.843)	62.59 (10.843)	62.21 (8.835)
	Median	60.95	60.95	60.95	61.80
	Min, Max	48.2, 82.2	48.2, 82.2	48.2, 82.2	46.0, 82.2
Ethnicity					
Japanese	n (%)	18 (100 %)	18 (100 %)	18 (100 %)	61 (100 %)
Not Japanese	n (%)	0	0	0	0

Note: NRT gum = Nicotine Replacement Therapy gum; 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, 15.3.1.10

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_demog3.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.1.4.3 Summary of Demographics and Other Baseline Characteristics - Group-2 PK Population**

Variable	Statistic	THS 2.2 Menthol (N=18)	NRT gum (N=18)	Group-2 PK (N=18)	Overall PK (N=61)
Daily mCC consumption at Admission					
10-19 cig/day	n (%)	11 ( 61.1%)	11 ( 61.1%)	11 ( 61.1%)	36 ( 59.0%)
>19 cig/day	n (%)	7 ( 38.9%)	7 ( 38.9%)	7 ( 38.9%)	25 ( 41.0%)
mCC nicotine yield at Admission (mg)	n	18	18	18	61
	Mean (SD)	0.46 (0.262)	0.46 (0.262)	0.46 (0.262)	0.47 (0.280)
	Median	0.55	0.55	0.55	0.60
	Min, Max	0.1, 0.7	0.1, 0.7	0.1, 0.7	0.1, 0.9
mCC ISO tar yield at Admission (mg)	n	18	18	18	61
	Mean (SD)	5.6 (3.33)	5.6 (3.33)	5.6 (3.33)	5.9 (3.86)
	Median	6.5	6.5	6.5	8.0
	Min, Max	1, 10	1, 10	1, 10	1, 12

Note: NRT gum = Nicotine Replacement Therapy gum; 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, 15.3.1.10

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_demog3.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.1.4.3 Summary of Demographics and Other Baseline Characteristics - Group-2 PK Population**

Variable	Statistic	THS 2.2 Menthol (N=18)	NRT gum (N=18)	Group-2 PK (N=18)	Overall PK (N=61)
mCC ISO tar yield at Admission					
1-5 mg	n (%)	9 ( 50.0%)	9 ( 50.0%)	9 ( 50.0%)	28 ( 45.9%)
6-8 mg	n (%)	7 ( 38.9%)	7 ( 38.9%)	7 ( 38.9%)	22 ( 36.1%)
9-10 mg	n (%)	2 ( 11.1%)	2 ( 11.1%)	2 ( 11.1%)	4 ( 6.6%)
>10 mg	n (%)	0	0	0	7 ( 11.5%)
CYP2A6 activity (%)					
	n	18	18	18	61
	Mean (SD)	26.859 (21.4010)	26.859 (21.4010)	26.859 (21.4010)	26.944 (20.5127)
	Median	18.147	18.147	18.147	20.530
	Min, Max	4.72, 68.99	4.72, 68.99	4.72, 68.99	4.19, 79.41
FTND total score classifications					
Mild	n (%)	9 ( 50.0%)	9 ( 50.0%)	9 ( 50.0%)	27 ( 44.3%)
Moderate	n (%)	7 ( 38.9%)	7 ( 38.9%)	7 ( 38.9%)	30 ( 49.2%)
Severe	n (%)	2 ( 11.1%)	2 ( 11.1%)	2 ( 11.1%)	4 ( 6.6%)

Note: NRT gum = Nicotine Replacement Therapy gum; 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, 15.3.1.10

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_demog3.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.1.4.3 Summary of Demographics and Other Baseline Characteristics - Group-2 PK Population**

Variable	Statistic	THS 2.2 Menthol (N=18)	NRT gum (N=18)	Group-2 PK (N=18)	Overall PK (N=61)
FTND total score	n	18	18	18	61
	Mean (SD)	3.6 (1.89)	3.6 (1.89)	3.6 (1.89)	3.8 (1.63)
	Median	3.5	3.5	3.5	4.0
	Min, Max	1, 8	1, 8	1, 8	1, 8

Note: NRT gum = Nicotine Replacement Therapy gum; 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, 15.3.1.10

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_demog3.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.1.4.3.1 Summary of Demographics and Other Baseline Characteristics by Sex - Group-2 PK Population**

Variable	Statistic	THS 2.2 Menthol		NRT gum		Group-2 PK	
		Males (N=9)	Females (N=9)	Males (N=9)	Females (N=9)	Males (N=9)	Females (N=9)
ISO nicotine level							
<= 0.6 mg	n (%)	5 ( 55.6%)	5 ( 55.6%)	5 ( 55.6%)	5 ( 55.6%)	5 ( 55.6%)	5 ( 55.6%)
> 0.6 - 1.0 mg	n (%)	4 ( 44.4%)	4 ( 44.4%)	4 ( 44.4%)	4 ( 44.4%)	4 ( 44.4%)	4 ( 44.4%)
Age (years)							
	n	9	9	9	9	9	9
	Mean (SD)	28.6 (8.37)	32.9 (6.98)	28.6 (8.37)	32.9 (6.98)	28.6 (8.37)	32.9 (6.98)
	Median	24.0	29.0	24.0	29.0	24.0	29.0
	Min, Max	23, 44	25, 44	23, 44	25, 44	23, 44	25, 44
BMI (kg/m <sup>2</sup> )							
	n	9	9	9	9	9	9
	Mean (SD)	23.87 (2.175)	21.84 (2.541)	23.87 (2.175)	21.84 (2.541)	23.87 (2.175)	21.84 (2.541)
	Median	23.15	21.91	23.15	21.91	23.15	21.91
	Min, Max	21.1, 28.4	18.8, 26.6	21.1, 28.4	18.8, 26.6	21.1, 28.4	18.8, 26.6

Note: NRT gum = Nicotine Replacement Therapy gum; 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, 15.3.1.10

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_demog3s.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.1.4.3.1 Summary of Demographics and Other Baseline Characteristics by Sex - Group-2 PK Population**

Variable	Statistic	THS 2.2 Menthol		NRT gum		Group-2 PK	
		Males (N=9)	Females (N=9)	Males (N=9)	Females (N=9)	Males (N=9)	Females (N=9)
BMI classifications							
Underweight	n (%)	0	0	0	0	0	0
Normal weight	n (%)	7 ( 77.8%)	8 ( 88.9%)	7 ( 77.8%)	8 ( 88.9%)	7 ( 77.8%)	8 ( 88.9%)
Overweight	n (%)	2 ( 22.2%)	1 ( 11.1%)	2 ( 22.2%)	1 ( 11.1%)	2 ( 22.2%)	1 ( 11.1%)
Obese	n (%)	0	0	0	0	0	0
Height (m)	n	9	9	9	9	9	9
	Mean (SD)	1.720 (0.0619)	1.581 (0.0644)	1.720 (0.0619)	1.581 (0.0644)	1.720 (0.0619)	1.581 (0.0644)
	Median	1.700	1.580	1.700	1.580	1.700	1.580
	Min, Max	1.65, 1.86	1.50, 1.68	1.65, 1.86	1.50, 1.68	1.65, 1.86	1.50, 1.68

Note: NRT gum = Nicotine Replacement Therapy gum; 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, 15.3.1.10

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_demog3s.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.1.4.3.1 Summary of Demographics and Other Baseline Characteristics by Sex - Group-2 PK Population**

Variable	Statistic	THS 2.2 Menthol		NRT gum		Group-2 PK	
		Males (N=9)	Females (N=9)	Males (N=9)	Females (N=9)	Males (N=9)	Females (N=9)
Weight (kg)	n	9	9	9	9	9	9
	Mean (SD)	70.68 (7.835)	54.50 (6.422)	70.68 (7.835)	54.50 (6.422)	70.68 (7.835)	54.50 (6.422)
	Median	71.60	52.60	71.60	52.60	71.60	52.60
	Min, Max	57.5, 82.2	48.2, 70.8	57.5, 82.2	48.2, 70.8	57.5, 82.2	48.2, 70.8
Ethnicity							
Japanese	n (%)	9 (100 %)	9 (100 %)	9 (100 %)	9 (100 %)	9 (100 %)	9 (100 %)
Not Japanese	n (%)	0	0	0	0	0	0
Daily mCC consumption at Admission							
10-19 cig/day	n (%)	5 ( 55.6%)	6 ( 66.7%)	5 ( 55.6%)	6 ( 66.7%)	5 ( 55.6%)	6 ( 66.7%)
>19 cig/day	n (%)	4 ( 44.4%)	3 ( 33.3%)	4 ( 44.4%)	3 ( 33.3%)	4 ( 44.4%)	3 ( 33.3%)

Note: NRT gum = Nicotine Replacement Therapy gum; 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, 15.3.1.10

Path: /cvm/projects/prj/development/000000106326/dev/tables/t\_demog3s.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.1.4.3.1 Summary of Demographics and Other Baseline Characteristics by Sex - Group-2 PK Population**

Variable	Statistic	THS 2.2 Menthol		NRT gum		Group-2 PK	
		Males (N=9)	Females (N=9)	Males (N=9)	Females (N=9)	Males (N=9)	Females (N=9)
mCC nicotine yield at Admission (mg)	n	9	9	9	9	9	9
	Mean (SD)	0.51 (0.221)	0.41 (0.302)	0.51 (0.221)	0.41 (0.302)	0.51 (0.221)	0.41 (0.302)
	Median	0.60	0.50	0.60	0.50	0.60	0.50
	Min, Max	0.1, 0.7	0.1, 0.7	0.1, 0.7	0.1, 0.7	0.1, 0.7	0.1, 0.7
mCC ISO tar yield at Admission (mg)	n	9	9	9	9	9	9
	Mean (SD)	6.3 (2.79)	4.8 (3.81)	6.3 (2.79)	4.8 (3.81)	6.3 (2.79)	4.8 (3.81)
	Median	8.0	5.0	8.0	5.0	8.0	5.0
	Min, Max	1, 10	1, 10	1, 10	1, 10	1, 10	1, 10
mCC ISO tar yield at Admission							
1-5 mg	n (%)	4 ( 44.4%)	5 ( 55.6%)	4 ( 44.4%)	5 ( 55.6%)	4 ( 44.4%)	5 ( 55.6%)
6-8 mg	n (%)	4 ( 44.4%)	3 ( 33.3%)	4 ( 44.4%)	3 ( 33.3%)	4 ( 44.4%)	3 ( 33.3%)
9-10 mg	n (%)	1 ( 11.1%)	1 ( 11.1%)	1 ( 11.1%)	1 ( 11.1%)	1 ( 11.1%)	1 ( 11.1%)
>10 mg	n (%)	0	0	0	0	0	0

Note: NRT gum = Nicotine Replacement Therapy gum; 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, 15.3.1.10

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_demog3s.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.1.4.3.1 Summary of Demographics and Other Baseline Characteristics by Sex - Group-2 PK Population**

Variable	Statistic	THS 2.2 Menthol		NRT gum		Group-2 PK	
		Males (N=9)	Females (N=9)	Males (N=9)	Females (N=9)	Males (N=9)	Females (N=9)
CYP2A6 activity (%)	n	9	9	9	9	9	9
	Mean (SD)	23.822 (19.5464)	29.896 (23.8840)	23.822 (19.5464)	29.896 (23.8840)	23.822 (19.5464)	29.896 (23.8840)
	Median	17.881	18.413	17.881	18.413	17.881	18.413
	Min, Max	4.72, 57.96	7.88, 68.99	4.72, 57.96	7.88, 68.99	4.72, 57.96	7.88, 68.99
FTND total score classifications							
Mild	n (%)	5 ( 55.6%)	4 ( 44.4%)	5 ( 55.6%)	4 ( 44.4%)	5 ( 55.6%)	4 ( 44.4%)
Moderate	n (%)	2 ( 22.2%)	5 ( 55.6%)	2 ( 22.2%)	5 ( 55.6%)	2 ( 22.2%)	5 ( 55.6%)
Severe	n (%)	2 ( 22.2%)	0	2 ( 22.2%)	0	2 ( 22.2%)	0
FTND total score	n	9	9	9	9	9	9
	Mean (SD)	3.8 (2.34)	3.4 (1.43)	3.8 (2.34)	3.4 (1.43)	3.8 (2.34)	3.4 (1.43)
	Median	3.0	4.0	3.0	4.0	3.0	4.0
	Min, Max	1, 8	1, 6	1, 8	1, 6	1, 8	1, 6

Note: NRT gum = Nicotine Replacement Therapy gum; 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, 15.3.1.10

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_demog3s.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.1.4.3.2 Summary of Demographics and Other Baseline Characteristics by Nicotine Level - Group-2 PK Population**

Variable	Statistic	THS 2.2 Menthol		NRT gum		Group-2 PK	
		<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)	<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)	<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)
Sex							
Male	n (%)	5 ( 50.0%)	4 ( 50.0%)	5 ( 50.0%)	4 ( 50.0%)	5 ( 50.0%)	4 ( 50.0%)
Female	n (%)	5 ( 50.0%)	4 ( 50.0%)	5 ( 50.0%)	4 ( 50.0%)	5 ( 50.0%)	4 ( 50.0%)
Age (years)	n	10	8	10	8	10	8
	Mean (SD)	32.8 (8.34)	28.1 (6.67)	32.8 (8.34)	28.1 (6.67)	32.8 (8.34)	28.1 (6.67)
	Median	32.0	26.5	32.0	26.5	32.0	26.5
	Min, Max	23, 44	23, 44	23, 44	23, 44	23, 44	23, 44
BMI (kg/m <sup>2</sup> )	n	10	8	10	8	10	8
	Mean (SD)	22.72 (2.469)	23.03 (2.743)	22.72 (2.469)	23.03 (2.743)	22.72 (2.469)	23.03 (2.743)
	Median	22.92	23.60	22.92	23.60	22.92	23.60
	Min, Max	19.1, 28.4	18.8, 26.6	19.1, 28.4	18.8, 26.6	19.1, 28.4	18.8, 26.6

Note: NRT gum = Nicotine Replacement Therapy gum; 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, 15.3.1.10

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_demog3c.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.1.4.3.2 Summary of Demographics and Other Baseline Characteristics by Nicotine Level - Group-2 PK Population**

Variable	Statistic	THS 2.2 Menthol		NRT gum		Group-2 PK	
		<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)	<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)	<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)
BMI classifications							
Underweight	n (%)	0	0	0	0	0	0
Normal weight	n (%)	9 ( 90.0%)	6 ( 75.0%)	9 ( 90.0%)	6 ( 75.0%)	9 ( 90.0%)	6 ( 75.0%)
Overweight	n (%)	1 ( 10.0%)	2 ( 25.0%)	1 ( 10.0%)	2 ( 25.0%)	1 ( 10.0%)	2 ( 25.0%)
Obese	n (%)	0	0	0	0	0	0
Height (m)							
	n	10	8	10	8	10	8
	Mean (SD)	1.654 (0.1121)	1.646 (0.0729)	1.654 (0.1121)	1.646 (0.0729)	1.654 (0.1121)	1.646 (0.0729)
	Median	1.675	1.660	1.675	1.660	1.675	1.660
	Min, Max	1.50, 1.86	1.50, 1.74	1.50, 1.86	1.50, 1.74	1.50, 1.86	1.50, 1.74

Note: NRT gum = Nicotine Replacement Therapy gum; 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, 15.3.1.10

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_demog3c.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.1.4.3.2 Summary of Demographics and Other Baseline Characteristics by Nicotine Level - Group-2 PK Population**

Variable	Statistic	THS 2.2 Menthol		NRT gum		Group-2 PK	
		<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)	<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)	<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)
Weight (kg)	n	10	8	10	8	10	8
	Mean (SD)	62.60 (12.180)	62.58 (9.738)	62.60 (12.180)	62.58 (9.738)	62.60 (12.180)	62.58 (9.738)
	Median	59.75	62.50	59.75	62.50	59.75	62.50
	Min, Max	48.2, 82.2	52.1, 75.0	48.2, 82.2	52.1, 75.0	48.2, 82.2	52.1, 75.0
Ethnicity							
Japanese	n (%)	10 (100 %)	8 (100 %)	10 (100 %)	8 (100 %)	10 (100 %)	8 (100 %)
Not Japanese	n (%)	0	0	0	0	0	0
Daily mCC consumption at Admission							
10-19 cig/day	n (%)	7 ( 70.0%)	4 ( 50.0%)	7 ( 70.0%)	4 ( 50.0%)	7 ( 70.0%)	4 ( 50.0%)
>19 cig/day	n (%)	3 ( 30.0%)	4 ( 50.0%)	3 ( 30.0%)	4 ( 50.0%)	3 ( 30.0%)	4 ( 50.0%)

Note: NRT gum = Nicotine Replacement Therapy gum; 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, 15.3.1.10

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_demog3c.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.1.4.3.2 Summary of Demographics and Other Baseline Characteristics by Nicotine Level - Group-2 PK Population**

Variable	Statistic	THS 2.2 Menthol		NRT gum		Group-2 PK	
		<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)	<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)	<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)
mCC nicotine yield at Admission (mg)	n	10	8	10	8	10	8
	Mean (SD)	0.27 (0.195)	0.70 (0.000)	0.27 (0.195)	0.70 (0.000)	0.27 (0.195)	0.70 (0.000)
	Median	0.20	0.70	0.20	0.70	0.20	0.70
	Min, Max	0.1, 0.6	0.7, 0.7	0.1, 0.6	0.7, 0.7	0.1, 0.6	0.7, 0.7
mCC ISO tar yield at Admission (mg)	n	10	8	10	8	10	8
	Mean (SD)	3.2 (2.53)	8.5 (0.93)	3.2 (2.53)	8.5 (0.93)	3.2 (2.53)	8.5 (0.93)
	Median	2.5	8.0	2.5	8.0	2.5	8.0
	Min, Max	1, 8	8, 10	1, 8	8, 10	1, 8	8, 10

Note: NRT gum = Nicotine Replacement Therapy gum; 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, 15.3.1.10

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_demog3c.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.1.4.3.2 Summary of Demographics and Other Baseline Characteristics by Nicotine Level - Group-2 PK Population**

Variable	Statistic	THS 2.2 Menthol		NRT gum		Group-2 PK	
		<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)	<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)	<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)
mCC ISO tar yield at Admission							
1-5 mg	n (%)	9 ( 90.0%)	0	9 ( 90.0%)	0	9 ( 90.0%)	0
6-8 mg	n (%)	1 ( 10.0%)	6 ( 75.0%)	1 ( 10.0%)	6 ( 75.0%)	1 ( 10.0%)	6 ( 75.0%)
9-10 mg	n (%)	0	2 ( 25.0%)	0	2 ( 25.0%)	0	2 ( 25.0%)
>10 mg	n (%)	0	0	0	0	0	0
CYP2A6 activity (%)							
	n	10	8	10	8	10	8
	Mean (SD)	25.724 (24.2615)	28.277 (18.7447)	25.724 (24.2615)	28.277 (18.7447)	25.724 (24.2615)	28.277 (18.7447)
	Median	15.470	20.505	15.470	20.505	15.470	20.505
	Min, Max	4.72, 68.99	10.21, 57.96	4.72, 68.99	10.21, 57.96	4.72, 68.99	10.21, 57.96

Note: NRT gum = Nicotine Replacement Therapy gum; 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, 15.3.1.10

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_demog3c.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.1.4.3.2 Summary of Demographics and Other Baseline Characteristics by Nicotine Level - Group-2 PK Population**

Variable	Statistic	THS 2.2 Menthol		NRT gum		Group-2 PK	
		<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)	<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)	<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)
FTND total score classifications							
Mild	n (%)	6 ( 60.0%)	3 ( 37.5%)	6 ( 60.0%)	3 ( 37.5%)	6 ( 60.0%)	3 ( 37.5%)
Moderate	n (%)	2 ( 20.0%)	5 ( 62.5%)	2 ( 20.0%)	5 ( 62.5%)	2 ( 20.0%)	5 ( 62.5%)
Severe	n (%)	2 ( 20.0%)	0	2 ( 20.0%)	0	2 ( 20.0%)	0
FTND total score	n	10	8	10	8	10	8
	Mean (SD)	3.8 (2.15)	3.4 (1.60)	3.8 (2.15)	3.4 (1.60)	3.8 (2.15)	3.4 (1.60)
	Median	3.0	4.0	3.0	4.0	3.0	4.0
	Min, Max	1, 8	1, 6	1, 8	1, 6	1, 8	1, 6

Note: NRT gum = Nicotine Replacement Therapy gum; 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, 15.3.1.10

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_demog3c.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)	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9) n	NRT gum - THS 2.2 Menthol (N=9) (%)		
Kool milds 8	0.7	8	1 ( 4.5%)	2 ( 9.1%)	1 ( 11.1%)	2 ( 22.2%)	1 ( 9.1%)	7 ( 9.6%)
Marlboro menthol	0.8	12	3 ( 13.6%)	2 ( 9.1%)	0	0	1 ( 9.1%)	6 ( 8.2%)
Hi-lite menthol	0.7	10	0	2 ( 9.1%)	1 ( 11.1%)	1 ( 11.1%)	0	4 ( 5.5%)
Pianissimo aria menthol	0.1	1	1 ( 4.5%)	1 ( 4.5%)	0	1 ( 11.1%)	1 ( 9.1%)	4 ( 5.5%)
Marlboro ice blast 8	0.6	8	0	1 ( 4.5%)	1 ( 11.1%)	0	1 ( 9.1%)	3 ( 4.1%)
Mevius premium menthol 8	0.7	8	0	3 ( 13.6%)	0	0	0	3 ( 4.1%)
Mevius premium menthol one 100's	0.1	1	3 ( 13.6%)	0	0	0	0	3 ( 4.1%)
Mevius premium menthol option 8	0.7	8	1 ( 4.5%)	0	1 ( 11.1%)	1 ( 11.1%)	0	3 ( 4.1%)
Pianissimo petil menthol one	0.1	1	0	2 ( 9.1%)	0	0	1 ( 9.1%)	3 ( 4.1%)
Kent iboost 1	0.1	1	1 ( 4.5%)	0	0	0	1 ( 9.1%)	2 ( 2.7%)
Kool mixx 5	0.4	5	1 ( 4.5%)	0	0	0	1 ( 9.1%)	2 ( 2.7%)

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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/000000106326/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)	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9) n	NRT gum - THS 2.2 Menthol (N=9) (%)		
Marlboro ice blast 1	0.1	1	1 ( 4.5%)	0	0	1 ( 11.1%)	0	2 ( 2.7%)
Marlboro lights menthol box	0.6	8	0	1 ( 4.5%)	0	0	1 ( 9.1%)	2 ( 2.7%)
Virginia s. ice pearl	0.1	1	1 ( 4.5%)	1 ( 4.5%)	0	0	0	2 ( 2.7%)
Kent charged mintek 1 100's	0.1	1	0	0	1 ( 11.1%)	0	0	1 ( 1.4%)
Kent nanotek2.0 8	0.7	8	1 ( 4.5%)	0	0	0	0	1 ( 1.4%)
Kent s-series 100's	0.1	1	0	0	0	0	1 ( 9.1%)	1 ( 1.4%)
Kool boost 5 box	0.4	5	0	0	0	1 ( 11.1%)	0	1 ( 1.4%)
Kool kings 12	0.9	12	0	0	0	0	1 ( 9.1%)	1 ( 1.4%)
Kool maxx 8	0.7	8	0	0	1 ( 11.1%)	0	0	1 ( 1.4%)
Kool milds box	0.7	8	0	1 ( 4.5%)	0	0	0	1 ( 1.4%)
Kool natural 8	0.6	8	1 ( 4.5%)	0	0	0	0	1 ( 1.4%)

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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/000000106326/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)	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9) n	NRT gum - THS 2.2 Menthol (N=9) (%)		
Kool natural 8 box	0.6	8	0	1 ( 4.5%)	0	0	0	1 ( 1.4%)
Lark ultra menthol 100's	0.1	1	0	0	0	0	1 ( 9.1%)	1 ( 1.4%)
Lucky strike menthol lights	0.6	6	0	1 ( 4.5%)	0	0	0	1 ( 1.4%)
Marlboro black menthol one	0.1	1	1 ( 4.5%)	0	0	0	0	1 ( 1.4%)
Marlboro menthol box	0.8	12	1 ( 4.5%)	0	0	0	0	1 ( 1.4%)
Marlboro menthol lights	0.6	8	0	1 ( 4.5%)	0	0	0	1 ( 1.4%)
Marlboro ultra lights menthol	0.3	4	0	1 ( 4.5%)	0	0	0	1 ( 1.4%)
Marlboro ultra lights menthol box	0.3	4	0	0	0	1 ( 11.1%)	0	1 ( 1.4%)
Mevius premium menthol 5	0.5	5	0	0	1 ( 11.1%)	0	0	1 ( 1.4%)
Philip morris super menthol	0.4	5	1 ( 4.5%)	0	0	0	0	1 ( 1.4%)
Philip morris super menthol box	0.4	5	1 ( 4.5%)	0	0	0	0	1 ( 1.4%)

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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/000000106326/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)	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9) n	NRT gum - THS 2.2 Menthol (N=9) (%)		
Philip morris super menthol ks box	0.4	5	0	0	1 ( 11.1%)	0	0	1 ( 1.4%)
Pianissimo icene menthol 1	0.1	1	1 ( 4.5%)	0	0	0	0	1 ( 1.4%)
Pianissimo lucia menthol	0.3	5	0	1 ( 4.5%)	0	0	0	1 ( 1.4%)
Pianissimo precia super slims	0.1	1	1 ( 4.5%)	0	0	0	0	1 ( 1.4%)
Seven stars deep menthol box	0.9	12	1 ( 4.5%)	0	0	0	0	1 ( 1.4%)
Virginia s. duo menthol	0.1	1	0	0	1 ( 11.1%)	0	0	1 ( 1.4%)
Virginia s. one	0.1	1	0	1 ( 4.5%)	0	0	0	1 ( 1.4%)
Virginia s. rose menthol	0.1	1	0	0	0	1 ( 11.1%)	0	1 ( 1.4%)

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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/000000106326/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	Sequence					Overall Safety (N=73)
	THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)	Enrolled Not Randomized (N=11)	
	n (%) Events	n (%) Events	n (%) Events	n (%) Events	n (%) Events	

No medical history data recorded

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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/000000106326/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	Sequence					Overall Safety (N=73)
	THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)	Enrolled Not Randomized (N=11)	
	n (%) Events	n (%) Events	n (%) Events	n (%) Events	n (%) Events	n (%) Events

No concomitant disease data recorded

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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/000000106326/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 Product Use - Safety Population**

Visit (Study Day)	Product	Product use	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Day -1	THS 2.2 Menthol	1 n (%)	22 (100 %)	22 (100 %)	9 (100 %)	9 (100 %)	11 (100 %)	73 (100 %)
		2 n (%)	0	0	0	0	0	0
		3 n (%)	0	0	0	0	0	0
	NRT gum	35 +/- 5 min n (%)	22 (100 %)	22 (100 %)	9 (100 %)	9 (100 %)	11 (100 %)	73 (100 %)

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Product use for NRT gum = chewing duration; Product use for mCC/THS = number of menthol cigarettes/THS Tobacco Sticks.

Appendix 15.3.2.1.1, 15.3.2.1.2, 15.3.2.1.3

Path: /cvn/projects/prj/development/000000106326/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 Product Use - Safety Population**

Visit (Study Day)	Product	Product use	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Day 1	THS 2.2 Menthol	0 n (%)	0	22 (100 %)	0	9 (100 %)		31 ( 42.5%)
		1 n (%)	22 (100 %)	0	9 (100 %)	0		31 ( 42.5%)
	NRT gum	0 n (%)			9 (100 %)	0		9 ( 12.3%)
		35 +/- 5 min n (%)			0	9 (100 %)		9 ( 12.3%)
	mCC	0 n (%)	22 (100 %)	0				22 ( 30.1%)
		1 n (%)	0	22 (100 %)				22 ( 30.1%)

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Product use for NRT gum = chewing duration; Product use for mCC/THS = number of menthol cigarettes/THS Tobacco Sticks.

Appendix 15.3.2.1.1, 15.3.2.1.2, 15.3.2.1.3

Path: /cvn/projects/prj/development/000000106326/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 Product Use - Safety Population**

Visit (Study Day)	Product	Product use	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Day 3	THS 2.2 Menthol	0 n (%)	21 ( 95.5%)	0	9 (100 %)	0		30 ( 41.1%)
		1 n (%)	0	22 (100 %)	0	9 (100 %)		31 ( 42.5%)
	NRT gum	0 n (%)			0	9 (100 %)		9 ( 12.3%)
		35 +/- 5 min n (%)			9 (100 %)	0		9 ( 12.3%)
	mCC	0 n (%)	0	22 (100 %)				22 ( 30.1%)
		1 n (%)	21 ( 95.5%)	0				21 ( 28.8%)

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Product use for NRT gum = chewing duration; Product use for mCC/THS = number of menthol cigarettes/THS Tobacco Sticks.

Appendix 15.3.2.1.1, 15.3.2.1.2, 15.3.2.1.3

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_ustatss.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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### **15.2.3 Primary Endpoints Summary Tables**

**Table 15.2.3.1 Analysis of Primary Pharmacokinetic Parameters of Nicotine - Group-1 PK Population**

Variable	Statistic	THS 2.2 Menthol (N=43)	mCC (N=43)	THS 2.2 Menthol:mCC Ratio (%)
C <sub>max</sub> (ng/mL)	n	43	43	
	Geometric LS Mean (CV%)	10.70	12.09	88.47 (63.54)
	95% CI	8.94, 12.80	10.10, 14.47	68.64, 114.03
	Precision			25.55
AUC <sub>(0-last)</sub> (h*ng/mL)	n	43	43	
	Geometric LS Mean (CV%)	23.99	24.45	98.13 (47.55)
	95% CI	20.87, 27.57	21.27, 28.10	80.61, 119.46
	Precision			21.33

Note: mCC = menthol conventional cigarettes; THS = Tobacco Heating System.

Note: Geometric LS Mean and 95% CI are the adjusted geometric least squares means based on an ANOVA model. Geometrical CV% of the ratio is estimated only for the ratio. Precision is the largest difference between the 95% CI bounds and the mean

Appendix 15.4.3.1 and 15.3.3.1

Path: /cvn/projects/prj/development/000000106326/dev/tables/tl\_anlpk31.sas

Program Run: 28OCT14 cvn\_ahedge Program Status: FINAL

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**Table 15.2.3.2 Analysis of Primary Pharmacokinetic Parameters of Nicotine by Sex and Nicotine Level - Group-1 PK Population**

Variable	Statistic	THS 2.2 Menthol		mCC		THS 2.2 Menthol:mCC Ratio (%)	
		Male (N=23)	Female (N=20)	Male (N=23)	Female (N=20)	Male	Female
AUC <sub>(0-last)</sub> (h*ng/mL)	n	23	20	23	20		
	Geometric LS Mean (CV%)	26.26	21.81	28.85	20.44	91.01 (40.11)	106.71 (56.68)
	95% CI	22.20, 31.05	17.02, 27.95	24.39, 34.12	15.95, 26.19	71.79, 115.36	75.15, 151.53
	Precision					24.35	44.82
C <sub>max</sub> (ng/mL)	n	23	20	23	20		
	Geometric LS Mean (CV%)	10.70	10.75	11.01	13.55	97.20 (56.20)	79.35 (74.31)
	95% CI	8.52, 13.44	7.87, 14.69	8.77, 13.83	9.92, 18.51	70.47, 134.08	51.07, 123.28
	Precision					36.88	43.93

Note: mCC = menthol conventional cigarettes; THS = Tobacco Heating System.

Note: Geometric LS Mean and 95% CI are the adjusted geometric least squares means based on an ANOVA model. Geometrical CV% of the ratio is estimated only for the ratio. Precision is the largest difference between the 95% CI bounds and the mean

Appendix 15.4.3.2 and 15.3.3.1

Path: /cvn/projects/prj/development/000000106326/dev/tables/tl\_anlpk32.sas

Program Run: 28OCT14 cvn\_ahedge Program Status: FINAL

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**Table 15.2.3.2 Analysis of Primary Pharmacokinetic Parameters of Nicotine by Sex and Nicotine Level - Group-1 PK Population**

Variable	Statistic	THS 2.2 Menthol:mCC Ratio (%)					
		THS 2.2 Menthol		mCC			
		<= 0.6mg (N=25)	> 0.6 mg - 1 mg (N=18)	<= 0.6mg (N=25)	> 0.6 mg - 1 mg (N=18)	<= 0.6mg	> 0.6 mg - 1 mg
AUC <sub>(0-last)</sub> (h*ng/mL)	n	25	18	25	18		
	Geometric LS Mean (CV%)	27.87	19.72	21.58	28.78	129.13 (33.08)	68.53 (48.82)
	95% CI	24.38, 31.85	15.62, 24.89	18.88, 24.67	22.80, 36.31	106.92, 155.95	49.32, 95.21
	Precision					26.82	26.68
C <sub>max</sub> (ng/mL)	n	25	18	25	18		
	Geometric LS Mean (CV%)	14.13	7.35	11.95	11.98	118.30 (37.83)	61.36 (76.79)
	95% CI	12.14, 16.45	5.22, 10.36	10.26, 13.91	8.50, 16.88	95.49, 146.56	37.81, 99.57
	Precision					28.26	38.21

Note: mCC = menthol conventional cigarettes; THS = Tobacco Heating System.

Note: Geometric LS Mean and 95% CI are the adjusted geometric least squares means based on an ANOVA model. Geometrical CV% of the ratio is estimated only for the ratio. Precision is the largest difference between the 95% CI bounds and the mean

Appendix 15.4.3.2 and 15.3.3.1

Path: /cvm/projects/prj/development/000000106326/dev/tables/tl\_anlpk32.sas

Program Run: 28OCT14 cvn\_ahedge Program Status: FINAL

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**Table 15.2.3.3 Analysis of Pharmacokinetic Parameters of Nicotine  $C_{\max}$ ,  $AUC_{(0-\text{last})}$ ,  $t_{\max}$  - PK Population**

Variable	Statistic	Group-1 PK		
		THS 2.2 Menthol (N=43)	mCC (N=43)	THS 2.2 Menthol:mCC Ratio (%)
$C_{\max}$ (ng/mL)	n	43	43	
	Geometric LS Mean (CV%) <sup>1</sup>	10.70	12.09	88.47 (63.54)
	95% CI	8.94, 12.80	10.10, 14.47	68.64, 114.03
	Precision			25.55
$AUC_{(0-\text{last})}$ (h*ng/mL)	n	43	43	
	Geometric LS Mean (CV%) <sup>1</sup>	23.99	24.45	98.13 (47.55)
	95% CI	20.87, 27.57	21.27, 28.10	80.61, 119.46
	Precision			21.33

Note: mCC = menthol 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. Geometrical CV% of the ratio is estimated only for the ratio. Precision is the largest difference between the 95% CI bounds and the mean

Note: 2: 95% CI are estimated only for the median difference based on the Hodges-Lehmann estimate.

Appendix 15.2.3.1, 15.2.4.1 and 15.2.4.2

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_anlpk33.sas

Program Run: 28OCT14 cvn\_ahedge Program Status: FINAL

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**Table 15.2.3.3 Analysis of Pharmacokinetic Parameters of Nicotine  $C_{\max}$ ,  $AUC_{(0-\text{last})}$ ,  $t_{\max}$  - PK Population**

Variable	Statistic	Group-1 PK		
		THS 2.2 Menthol (N=43)	mCC (N=43)	THS 2.2 Menthol - mCC (min)
$t_{\max}$ (min)	n	43	43	
	Median	6.00	6.00	1.00
	95% CI <sup>2</sup>			0.00, 2.50

Note: mCC = menthol 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. Geometrical CV% of the ratio is estimated only for the ratio. Precision is the largest difference between the 95% CI bounds and the mean

Note: 2: 95% CI are estimated only for the median difference based on the Hodges-Lehmann estimate.

Appendix 15.2.3.1, 15.2.4.1 and 15.2.4.2

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_anlpk33.sas

Program Run: 28OCT14 cvn\_ahedge Program Status: FINAL

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**Table 15.2.3.3 Analysis of Pharmacokinetic Parameters of Nicotine  $C_{\max}$ ,  $AUC_{(0-\text{last})}$ ,  $t_{\max}$  - PK Population**

Variable	Statistic	Group-2 PK		
		THS 2.2 Menthol (N=18)	NRT gum (N=18)	THS 2.2 Menthol :NRT gum Ratio (%)
$C_{\max}$ (ng/mL)	n	18	18	
	Geometric LS Mean (CV%) <sup>1</sup>	7.64	7.52	101.63 (78.75)
	95% CI	5.39, 10.81	5.31, 10.64	62.21, 166.04
	P-value (one-sided)			0.473
$AUC_{(0-\text{last})}$ (h*ng/mL)	n	18	18	
	Geometric LS Mean (CV%) <sup>1</sup>	15.61	27.94	55.87 (57.19)
	95% CI	11.96, 20.37	21.41, 36.45	38.36, 81.36
	P-value (one-sided)			0.998

Note: NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Note: 1: Geometric LS Mean and 95% CI are the adjusted geometric least squares means based on an ANOVA model. Geometrical CV% of the ratio is estimated only for the ratio. Precision is the largest difference between the 95% CI bounds and the mean

Note: 2: 95% CI are estimated only for the median difference based on the Hodges-Lehmann estimate.

Appendix 15.2.3.1, 15.2.4.1 and 15.2.4.2

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_anlpk33.sas

Program Run: 28OCT14 cvn\_ahedge Program Status: FINAL

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**Table 15.2.3.3 Analysis of Pharmacokinetic Parameters of Nicotine  $C_{max}$ ,  $AUC_{(0-last)}$ ,  $t_{max}$  - PK Population**

Variable	Statistic	Group-2 PK		
		THS 2.2 Menthol (N=18)	NRT gum (N=18)	THS 2.2 Menthol - NRT gum (min)
$t_{max}$ (min)	n	18	18	
	Median	8.00	45.00	-37.50
	95% CI <sup>2</sup>			-45.00, -31.50
	P-value (one-sided)			<0.001

Note: NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Note: 1: Geometric LS Mean and 95% CI are the adjusted geometric least squares means based on an ANOVA model. Geometrical CV% of the ratio is estimated only for the ratio. Precision is the largest difference between the 95% CI bounds and the mean

Note: 2: 95% CI are estimated only for the median difference based on the Hodges-Lehmann estimate.

Appendix 15.2.3.1, 15.2.4.1 and 15.2.4.2

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_anlpk33.sas

Program Run: 28OCT14 cvn\_ahedge Program Status: FINAL

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## **15.2.4 Secondary Endpoints Summary Tables**

**Table 15.2.4.1 Analysis of Secondary Pharmacokinetic Parameters of Nicotine - Group-1 PK Population**

Variable	Statistic	THS 2.2 Menthol (N=43)	mCC (N=43)	THS 2.2 Menthol:mCC Ratio (%)
AUC <sub>(0-inf)</sub> (h*ng/mL)	n	34	34	
	Geometric LS Mean (CV%) <sup>1</sup>	26.33	27.73	94.97 (42.31)
	95% CI	22.84, 30.35	24.05, 31.96	77.69, 116.09
AUC <sub>(0-t)</sub> (h*ng/mL)	n	43	43	
	Geometric LS Mean (CV%) <sup>1</sup>	0.59	0.80	74.46 (66.67)
	95% CI	0.49, 0.72	0.66, 0.96	57.17, 96.98
t <sub>1/2</sub> (h)	n	34	34	
	Geometric LS Mean (CV%) <sup>1</sup>	4.11	4.02	102.30 (37.94)
	95% CI	3.61, 4.68	3.53, 4.57	85.31, 122.66

Note: mCC = menthol 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. Geometrical CV% of the ratio is estimated only for the ratio

Note: 2: 95% CI are estimated only for the median difference based on the Hodges-Lehmann estimate.

Appendix 15.4.4.1 and 15.3.3.1

Path: /cvn/projects/prj/development/000000106326/dev/tables/tl\_anlpk41.sas

Program Run: 28OCT14 cvn\_ahedge Program Status: FINAL

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**Table 15.2.4.1 Analysis of Secondary Pharmacokinetic Parameters of Nicotine - Group-1 PK Population**

Variable	Statistic	THS 2.2 Menthol (N=43)	mCC (N=43)	THS 2.2 Menthol - mCC (min)
t <sub>max</sub> (min)	n	43	43	
	Median	6.00	6.00	1.00
	95% CI <sup>2</sup>			0.00, 2.50

Note: mCC = menthol 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. Geometrical CV% of the ratio is estimated only for the ratio

Note: 2: 95% CI are estimated only for the median difference based on the Hodges-Lehmann estimate.

Appendix 15.4.4.1 and 15.3.3.1

Path: /cvn/projects/prj/development/000000106326/dev/tables/tl\_anl\_pk41.sas

Program Run: 28OCT14 cvn\_ahedge Program Status: FINAL

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**Table 15.2.4.2 Analysis of Pharmacokinetic Parameters of Nicotine - Group-2 PK Population**

Variable	Statistic	THS 2.2 Menthol (N=18)	NRT gum (N=18)	THS 2.2 Menthol :NRT gum Ratio (%)
C <sub>max</sub> (ng/mL)	n	18	18	
	Geometric LS Mean (CV%) <sup>1</sup>	7.64	7.52	101.63 (78.75)
	95% CI	5.39, 10.81	5.31, 10.64	62.21, 166.04
	P-value (one-sided)			0.473
AUC <sub>(0-last)</sub> (h*ng/mL)	n	18	18	
	Geometric LS Mean (CV%) <sup>1</sup>	15.61	27.94	55.87 (57.19)
	95% CI	11.96, 20.37	21.41, 36.45	38.36, 81.36
	P-value (one-sided)			0.998
AUC <sub>(0-inf)</sub> (h*ng/mL)	n	15	15	
	Geometric LS Mean (CV%) <sup>1</sup>	15.77	31.09	50.72 (51.05)
	95% CI	12.04, 20.64	23.75, 40.69	34.66, 74.21
	P-value (one-sided)			0.999

Note: NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Note: 1: Geometric LS Mean and 95% CI are the adjusted geometric least squares means based on an ANOVA model. Geometrical CV% of the ratio is estimated only for the ratio

Note: 2: 95% CI are estimated only for the median difference based on the Hodges-Lehmann estimate.

Appendix 15.4.4.2 and 15.3.3.1

Path: /cvn/projects/prj/development/000000106326/dev/tables/tl\_anlpk42.sas

Program Run: 28OCT14 cvn\_ahedge Program Status: FINAL

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**Table 15.2.4.2 Analysis of Pharmacokinetic Parameters of Nicotine - Group-2 PK Population**

Variable	Statistic	THS 2.2 Menthol (N=18)	NRT gum (N=18)	THS 2.2 Menthol :NRT gum Ratio (%)
AUC <sub>(0-t)</sub> (h*ng/mL)	n	18	18	
	Geometric LS Mean (CV%) <sup>1</sup>	3.38	2.97	113.63 (54.10)
	95% CI	2.62, 4.36	2.30, 3.83	79.43, 162.55
	P-value (one-sided)			0.230
t <sub>1/2</sub> (h)	n	15	15	
	Geometric LS Mean (CV%) <sup>1</sup>	3.20	3.47	92.06 (28.97)
	95% CI	2.72, 3.75	2.96, 4.08	73.55, 115.22

Note: NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Note: 1: Geometric LS Mean and 95% CI are the adjusted geometric least squares means based on an ANOVA model. Geometrical CV% of the ratio is estimated only for the ratio

Note: 2: 95% CI are estimated only for the median difference based on the Hodges-Lehmann estimate.

Appendix 15.4.4.2 and 15.3.3.1

Path: /cvn/projects/prj/development/000000106326/dev/tables/tl\_anl\_pk42.sas

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Program Run: 28OCT14 cvn\_ahedge Program Status: FINAL

**Table 15.2.4.2 Analysis of Pharmacokinetic Parameters of Nicotine - Group-2 PK Population**

Variable	Statistic	THS 2.2 Menthol (N=18)	NRT gum (N=18)	THS 2.2 Menthol - NRT gum (min)
t <sub>max</sub> (min)	n	18	18	
	Median	8.00	45.00	-37.50
	95% CI <sup>2</sup>			-45.00, -31.50
	P-value (one-sided)			<0.001

Note: NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Note: 1: Geometric LS Mean and 95% CI are the adjusted geometric least squares means based on an ANOVA model. Geometrical CV% of the ratio is estimated only for the ratio

Note: 2: 95% CI are estimated only for the median difference based on the Hodges-Lehmann estimate.

Appendix 15.4.4.2 and 15.3.3.1

Path: /cvn/projects/prj/development/000000106326/dev/tables/tl\_anlphk42.sas

Program Run: 28OCT14 cvn\_ahedge Program Status: FINAL

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**Table 15.2.4.3.1 Analysis of Pharmacokinetic Parameters of Nicotine by Bootstrapping Techniques  
- PK Population**

Variable	Statistic	Group-1 PK		
		THS 2.2 Menthol (N=43)	mCC (N=43)	THS 2.2 Menthol:mCC Ratio (%)
t <sub>max</sub> (min)	n	43	43	
	Geometric LS Mean (CV%)	10.14	7.24	140.05 (92.36)
	95% CI	7.30, 14.63	6.25, 8.52	101.17, 199.83
	Median	10.04	7.22	139.30

Note: mCC = menthol conventional cigarettes; THS = Tobacco Heating System.

Note: Point and 95% interval estimates by means of the percentile bootstrap technique of the adjusted geometric least squares means from an ANOVA model.

Appendix 15.4.4.3.1 and 15.3.3.1

Path: /cvn/projects/prj/development/000000106326/dev/tables/tl\_anlpk431.sas

Program Run: 28OCT14 cvn\_ahedge Program Status: FINAL

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**Table 15.2.4.3.1 Analysis of Pharmacokinetic Parameters of Nicotine by Bootstrapping Techniques  
- PK Population**

Variable	Statistic	Group-2 PK		
		THS 2.2 Menthol (N=18)	NRT gum (N=18)	THS 2.2 Menthol :NRT gum Ratio (%)
t <sub>max</sub> (min)	n	18	18	
	Geometric LS Mean (CV%)	8.07	46.07	17.51 (33.43)
	95% CI	6.53, 10.02	40.60, 51.82	14.12, 21.77
	Median	8.10	46.15	17.58

Note: NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Note: Point and 95% interval estimates by means of the percentile bootstrap technique of the adjusted geometric least squares means from an ANOVA model.

Appendix 15.4.4.3.1 and 15.3.3.1

Path: /cvn/projects/prj/development/000000106326/dev/tables/tl\_anlpk431.sas

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Program Run: 28OCT14 cvn\_ahedge Program Status: FINAL

**Table 15.2.4.3.2 Supportive Analysis of Pharmacokinetic Parameters of Nicotine Excluding Subject with T<sub>0</sub> Value >5% of Their C<sub>max</sub> Value - PK Population**

Variable	Statistic	Group-1 PK		
		THS 2.2 Menthol (N=43)	mCC (N=43)	THS 2.2 Menthol:mCC Ratio (%)
Number of subjects excluded		7	7	
C <sub>max</sub> (ng/mL)	n	36	36	
	Geometric LS Mean (CV%) <sup>1</sup>	11.58	13.33	86.88 (65.98)
	95% CI	9.44, 14.21	10.87, 16.35	65.11, 115.93
	Precision			29.05
AUC <sub>(0-last)</sub> (h*ng/mL)	n	36	36	
	Geometric LS Mean (CV%) <sup>1</sup>	22.65	22.83	99.22 (48.01)
	95% CI	19.40, 26.44	19.55, 26.64	79.74, 123.45
	Precision			24.23

Note: mCC = menthol 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. Geometrical CV% of the ratio is estimated only for the ratio. Precision is the largest difference between the 95% CI bounds and the mean

Note: 2: 95% CI are estimated only for the median difference based on the Hodges-Lehmann estimate.

Appendix 15.4.4.3.2 and 15.3.3.1

Path: /cvm/projects/prj/development/000000106326/dev/tables/tl\_anl\_pk432.sas

Program Run: 28OCT14 cvn\_ahedge Program Status: FINAL

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**Table 15.2.4.3.2 Supportive Analysis of Pharmacokinetic Parameters of Nicotine Excluding Subject with T<sub>0</sub> Value >5% of Their C<sub>max</sub> Value - PK Population**

Variable	Statistic	Group-1 PK		
		THS 2.2 Menthol (N=43)	mCC (N=43)	THS 2.2 Menthol:mCC Ratio (%)
AUC <sub>(0-inf)</sub> (h*ng/mL)	n	29	29	
	Geometric LS Mean (CV%) <sup>1</sup>	25.22	25.04	100.69 (43.45)
	95% CI	21.51, 29.55	21.37, 29.35	80.47, 126.00
AUC <sub>(0-t')</sub> (h*ng/mL)	n	36	36	
	Geometric LS Mean (CV%) <sup>1</sup>	0.67	0.88	75.54 (70.82)
	95% CI	0.53, 0.83	0.71, 1.10	55.63, 102.57
t <sub>1/2</sub> (h)	n	29	29	
	Geometric LS Mean (CV%) <sup>1</sup>	3.78	3.61	104.67 (39.48)
	95% CI	3.26, 4.37	3.12, 4.18	85.25, 128.52

Note: mCC = menthol 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. Geometrical CV% of the ratio is estimated only for the ratio. Precision is the largest difference between the 95% CI bounds and the mean

Note: 2: 95% CI are estimated only for the median difference based on the Hodges-Lehmann estimate.

Appendix 15.4.4.3.2 and 15.3.3.1

Path: /cvm/projects/prj/development/000000106326/dev/tables/tl\_anl\_pk432.sas

Program Run: 28OCT14 cvn\_ahedge Program Status: FINAL

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**Table 15.2.4.3.2 Supportive Analysis of Pharmacokinetic Parameters of Nicotine Excluding Subject with T<sub>0</sub> Value >5% of Their C<sub>max</sub> Value - PK Population**

Variable	Statistic	Group-1 PK		
		THS 2.2 Menthol (N=43)	mCC (N=43)	THS 2.2 Menthol - mCC (min)
t <sub>max</sub> (min)	n	36	36	
	Median	6.00	6.00	0.00
	95% CI <sup>2</sup>			-1.00, 1.00

Note: mCC = menthol 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. Geometrical CV% of the ratio is estimated only for the ratio. Precision is the largest difference between the 95% CI bounds and the mean

Note: 2: 95% CI are estimated only for the median difference based on the Hodges-Lehmann estimate.

Appendix 15.4.4.3.2 and 15.3.3.1

Path: /cvn/projects/prj/development/000000106326/dev/tables/tl\_anlpk432.sas

Program Run: 28OCT14 cvn\_ahedge Program Status: FINAL

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**Table 15.2.4.3.2 Supportive Analysis of Pharmacokinetic Parameters of Nicotine Excluding Subject with T<sub>0</sub> Value >5% of Their C<sub>max</sub> Value - PK Population**

Variable	Statistic	Group-2 PK		
		THS 2.2 Menthol (N=18)	NRT gum (N=18)	THS 2.2 Menthol :NRT gum Ratio (%)
Number of subjects excluded		5	5	
C <sub>max</sub> (ng/mL)	n	13	13	
	Geometric LS Mean (CV%) <sup>1</sup>	7.94	8.02	99.08 (93.34)
	95% CI	4.83, 13.06	4.87, 13.18	49.08, 199.99
	P-value (one-sided)			0.511
AUC <sub>(0-last)</sub> (h*ng/mL)	n	13	13	
	Geometric LS Mean (CV%) <sup>1</sup>	13.99	25.94	53.94 (63.25)
	95% CI	9.72, 20.14	18.02, 37.33	32.23, 90.24
	P-value (one-sided)			0.989

Note: NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Note: 1: Geometric LS Mean and 95% CI are the adjusted geometric least squares means based on an ANOVA model. Geometrical CV% of the ratio is estimated only for the ratio. Precision is the largest difference between the 95% CI bounds and the mean

Note: 2: 95% CI are estimated only for the median difference based on the Hodges-Lehmann estimate.

Appendix 15.4.4.3.2 and 15.3.3.1

Path: /cvn/projects/prj/development/000000106326/dev/tables/tl\_anl\_pk432.sas

Program Run: 28OCT14 cvn\_ahedge Program Status: FINAL

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**Table 15.2.4.3.2 Supportive Analysis of Pharmacokinetic Parameters of Nicotine Excluding Subject with T<sub>0</sub> Value >5% of Their C<sub>max</sub> Value - PK Population**

Variable	Statistic	Group-2 PK		
		THS 2.2 Menthol (N=18)	NRT gum (N=18)	THS 2.2 Menthol :NRT gum Ratio (%)
AUC <sub>(0-inf)</sub> (h*ng/mL)	n	11	11	
	Geometric LS Mean (CV%) <sup>1</sup>	13.17	27.17	48.48 (56.57)
	95% CI	9.06, 19.14	18.69, 39.47	28.58, 82.23
	P-value (one-sided)			0.994
AUC <sub>(0-t')</sub> (h*ng/mL)	n	13	13	
	Geometric LS Mean (CV%) <sup>1</sup>	3.09	2.87	107.80 (59.82)
	95% CI	2.18, 4.38	2.02, 4.06	65.99, 176.09
	P-value (one-sided)			0.371
t <sub>1/2</sub> (h)	n	11	11	
	Geometric LS Mean (CV%) <sup>1</sup>	2.84	3.14	90.24 (26.71)
	95% CI	2.35, 3.42	2.60, 3.79	69.35, 117.40

Note: NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Note: 1: Geometric LS Mean and 95% CI are the adjusted geometric least squares means based on an ANOVA model. Geometrical CV% of the ratio is estimated only for the ratio. Precision is the largest difference between the 95% CI bounds and the mean

Note: 2: 95% CI are estimated only for the median difference based on the Hodges-Lehmann estimate.

Appendix 15.4.4.3.2 and 15.3.3.1

Path: /cvn/projects/prj/development/000000106326/dev/tables/tl\_anl\_pk432.sas

Program Run: 28OCT14 cvn\_ahedge Program Status: FINAL

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**Table 15.2.4.3.2 Supportive Analysis of Pharmacokinetic Parameters of Nicotine Excluding Subject with  $T_0$  Value >5% of Their  $C_{max}$  Value - PK Population**

Variable	Statistic	Group-2 PK		
		THS 2.2 Menthol (N=18)	NRT gum (N=18)	THS 2.2 Menthol - NRT gum (min)
$t_{max}$ (min)	n	13	13	
	Median	6.00	45.00	-37.50
	95% CI <sup>2</sup>			-45.50, -27.50
	P-value (one-sided)			<0.001

Note: NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Note: 1: Geometric LS Mean and 95% CI are the adjusted geometric least squares means based on an ANOVA model. Geometrical CV% of the ratio is estimated only for the ratio. Precision is the largest difference between the 95% CI bounds and the mean

Note: 2: 95% CI are estimated only for the median difference based on the Hodges-Lehmann estimate.

Appendix 15.4.4.3.2 and 15.3.3.1

Path: /cvn/projects/prj/development/000000106326/dev/tables/tl\_anlpk432.sas

Program Run: 28OCT14 cvn\_ahedge Program Status: FINAL

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**Table 15.2.4.4.1 Analysis of Pharmacokinetic Parameters of Nicotine by Zero Second Imputation  
- PK Population**

Variable	Statistic	Group-1 PK		
		THS 2.2 Menthol (N=43)	mCC (N=43)	THS 2.2 Menthol:mCC Ratio (%)

No analysis necessary

Note: mCC = menthol conventional cigarettes; THS = Tobacco Heating System.

Appendix 15.4.4.4.1 and 15.3.3.1

Path: /cvn/projects/prj/development/000000106326/dev/tables/tl\_anl\_pk441.sas

Program Run: 28OCT14 cvn\_ahedge Program Status: FINAL

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**Table 15.2.4.4.1 Analysis of Pharmacokinetic Parameters of Nicotine by Zero Second Imputation  
- PK Population**

Variable	Statistic	Group-2 PK		
		THS 2.2 Menthol (N=18)	NRT gum (N=18)	THS 2.2 Menthol :NRT gum Ratio (%)

No analysis necessary

Note: NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Appendix 15.4.4.4.1 and 15.3.3.1

Path: /cvn/projects/prj/development/000000106326/dev/tables/tl\_anlpk441.sas

Program Run: 28OCT14 cvn\_ahedge Program Status: FINAL

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**Table 15.2.4.4.2 Analysis of Pharmacokinetic Parameters of Nicotine Where >20% Missing Observations  
- PK Population**

Variable	Statistic	Group-1 PK		
		THS 2.2 Menthol (N=43)	mCC (N=43)	THS 2.2 Menthol:mCC Ratio (%)

No analysis necessary

Note: mCC = menthol conventional cigarettes; THS = Tobacco Heating System.

Appendix 15.4.4.4.2 and 15.3.3.1

Path: /cvn/projects/prj/development/000000106326/dev/tables/tl\_anlpk442.sas

Program Run: 28OCT14 cvn\_ahedge Program Status: FINAL

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**Table 15.2.4.4.2 Analysis of Pharmacokinetic Parameters of Nicotine Where >20% Missing Observations  
- PK Population**

Variable	Statistic	Group-2 PK		
		THS 2.2 Menthol (N=18)	NRT gum (N=18)	THS 2.2 Menthol :NRT gum Ratio (%)

No analysis necessary

Note: NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Appendix 15.4.4.4.2 and 15.3.3.1

Path: /cvn/projects/prj/development/000000106326/dev/tables/tl\_anlpk442.sas

Program Run: 28OCT14 cvn\_ahedge Program Status: FINAL

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**Table 15.2.4.5 Descriptive Statistics of Pharmacokinetic Parameters of Nicotine - PK Population**

Variable	Statistic	Group-1 PK		Group-2 PK	
		THS 2.2 Menthol (N=43)	mCC (N=43)	THS 2.2 Menthol (N=18)	NRT gum (N=18)
C <sub>max</sub> (ng/mL)	n	43	43	18	18
	Geometric Mean (CV%)	10.65 (117.10%)	12.12 (87.71%)	7.64 (108.56%)	7.52 (55.24%)
	95% CI	7.99, 14.17	9.60, 15.30	4.92, 11.85	5.81, 9.72
	Median	12.70	12.20	8.29	7.36
	Q25, Q75	7.23, 19.70	7.97, 19.70	3.71, 11.50	5.80, 9.37
	Min, Max	0.4, 43.8	2.3, 58.1	1.3, 45.8	2.7, 34.1
	Mean (SD)	14.38 (9.996)	15.63 (11.349)	10.92 (10.568)	8.76 (6.702)

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_nic.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.5 Descriptive Statistics of Pharmacokinetic Parameters of Nicotine - PK Population**

Variable	Statistic	Group-1 PK		Group-2 PK	
		THS 2.2 Menthol (N=43)	mCC (N=43)	THS 2.2 Menthol (N=18)	NRT gum (N=18)
AUC <sub>(0-last)</sub> (h*ng/mL)	n	43	43	18	18
	Geometric Mean (CV%)	23.84 (110.43%)	24.50 (82.63%)	15.61 (77.53%)	27.94 (72.31%)
	95% CI	18.11, 31.39	19.61, 30.59	11.09, 21.96	20.23, 38.57
	Median	21.53	24.46	16.02	25.44
	Q25, Q75	15.97, 47.04	15.19, 37.58	9.90, 26.22	16.77, 42.87
	Min, Max	1.2, 117.7	4.9, 100.9	3.9, 48.3	9.6, 86.4
	Mean (SD)	33.12 (26.797)	31.21 (22.622)	19.41 (13.557)	34.15 (23.041)

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_nic.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.5 Descriptive Statistics of Pharmacokinetic Parameters of Nicotine - PK Population**

Variable	Statistic	Group-1 PK		Group-2 PK	
		THS 2.2 Menthol (N=43)	mCC (N=43)	THS 2.2 Menthol (N=18)	NRT gum (N=18)
AUC <sub>(0-∞)</sub> (h*ng/mL)	n	37	40	16	17
	Geometric Mean (CV%)	26.03 (85.72%)	26.97 (85.42%)	16.78 (76.36%)	29.71 (74.88%)
	95% CI	20.32, 33.34	21.28, 34.17	11.69, 24.08	21.08, 41.87
	Median	22.91	27.31	15.50	27.55
	Q25, Q75	16.85, 50.30	17.43, 41.15	10.97, 25.54	17.72, 47.35
	Min, Max	5.7, 111.0	5.4, 117.1	4.9, 53.3	11.0, 91.7
	Mean (SD)	33.81 (25.607)	34.82 (25.973)	20.94 (15.235)	36.78 (25.574)

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_nic.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.5 Descriptive Statistics of Pharmacokinetic Parameters of Nicotine - PK Population**

Variable	Statistic	Group-1 PK		Group-2 PK	
		THS 2.2 Menthol (N=43)	mCC (N=43)	THS 2.2 Menthol (N=18)	NRT gum (N=18)
%AUC <sub>extrap</sub> (%)	n	37	40	16	17
	Geometric Mean (CV%)	7.31 (59.84%)	7.16 (48.32%)	8.64 (63.78%)	6.71 (42.55%)
	95% CI	6.07, 8.79	6.18, 8.30	6.32, 11.80	5.43, 8.28
	Median	7.95	7.25	8.62	6.18
	Q25, Q75	4.55, 10.05	5.03, 10.21	5.84, 13.47	5.10, 9.11
	Min, Max	2.7, 24.3	2.7, 20.3	2.7, 20.1	3.4, 15.4
	Mean (SD)	8.48 (4.872)	7.93 (3.738)	10.06 (5.640)	7.28 (3.197)

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_nic.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.5 Descriptive Statistics of Pharmacokinetic Parameters of Nicotine - PK Population**

Variable	Statistic	Group-1 PK		Group-2 PK	
		THS 2.2 Menthol (N=43)	mCC (N=43)	THS 2.2 Menthol (N=18)	NRT gum (N=18)
AUC <sub>(0-t)</sub> (h*ng/mL)	n	43	43	18	18
	Geometric Mean (CV%)	0.59 (159.51%)	0.80 (97.91%)	3.38 (75.49%)	2.97 (50.66%)
	95% CI	0.41, 0.84	0.61, 1.03	2.41, 4.72	2.34, 3.78
	Median	0.67	0.78	3.54	3.08
	Q25, Q75	0.28, 1.29	0.47, 1.38	2.11, 5.56	2.44, 3.81
	Min, Max	0.0, 4.7	0.1, 4.2	1.0, 11.2	1.1, 7.6
	Mean (SD)	0.99 (1.047)	1.08 (0.896)	4.14 (2.707)	3.30 (1.559)

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_nic.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.5 Descriptive Statistics of Pharmacokinetic Parameters of Nicotine - PK Population**

Variable	Statistic	Group-1 PK		Group-2 PK	
		THS 2.2 Menthol (N=43)	mCC (N=43)	THS 2.2 Menthol (N=18)	NRT gum (N=18)
t <sub>max</sub> (min)	n	43	43	18	18
	Median	6.00	6.00	8.00	45.00
	Q25, Q75	6.00, 15.00	4.00, 10.00	4.00, 15.00	40.00, 60.00
	Min, Max	4.0, 720.0	4.0, 30.0	4.0, 30.0	25.0, 60.0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_nic.sas

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**Table 15.2.4.5 Descriptive Statistics of Pharmacokinetic Parameters of Nicotine - PK Population**

Variable	Statistic	Group-1 PK		Group-2 PK	
		THS 2.2 Menthol (N=43)	mCC (N=43)	THS 2.2 Menthol (N=18)	NRT gum (N=18)
t <sub>last</sub> (h)	n	43	43	18	18
	Median	12.00	24.00	12.00	12.00
	Q25, Q75	12.00, 24.00	9.00, 24.00	9.00, 24.00	12.00, 24.00
	Min, Max	6.0, 24.0	6.0, 24.0	4.0, 24.0	6.0, 24.0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_nic.sas

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**Table 15.2.4.5 Descriptive Statistics of Pharmacokinetic Parameters of Nicotine - PK Population**

Variable	Statistic	Group-1 PK		Group-2 PK	
		THS 2.2 Menthol (N=43)	mCC (N=43)	THS 2.2 Menthol (N=18)	NRT gum (N=18)
t <sub>1/2</sub> (h)	n	37	40	16	17
	Geometric Mean (CV%)	3.97 (65.91%)	3.88 (67.53%)	3.34 (63.71%)	3.82 (66.59%)
	95% CI	3.25, 4.86	3.18, 4.72	2.45, 4.57	2.79, 5.22
	Median	3.90	3.42	3.37	3.29
	Q25, Q75	2.47, 6.62	2.01, 6.80	1.93, 5.62	2.68, 5.90
	Min, Max	1.3, 12.0	1.6, 11.5	1.3, 7.7	1.3, 11.6
	Mean (SD)	4.69 (2.636)	4.61 (2.643)	3.89 (2.098)	4.52 (2.719)

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_nic.sas

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**Table 15.2.4.5 Descriptive Statistics of Pharmacokinetic Parameters of Nicotine - PK Population**

Variable	Statistic	Group-1 PK		Group-2 PK	
		THS 2.2 Menthol (N=43)	mCC (N=43)	THS 2.2 Menthol (N=18)	NRT gum (N=18)
$\lambda_z$ (1/h)	n	37	40	16	17
	Geometric Mean (CV%)	0.17 (65.91%)	0.18 (67.53%)	0.21 (63.71%)	0.18 (66.59%)
	95% CI	0.14, 0.22	0.14, 0.22	0.15, 0.29	0.13, 0.25
	Median	0.18	0.20	0.21	0.21
	Q25, Q75	0.10, 0.29	0.10, 0.35	0.12, 0.37	0.11, 0.26
	Min, Max	0.1, 0.5	0.1, 0.4	0.1, 0.6	0.1, 0.6
	Mean (SD)	0.21 (0.124)	0.21 (0.124)	0.24 (0.142)	0.22 (0.134)

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_nic.sas

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**Table 15.2.4.5.1 Descriptive Statistics of Pharmacokinetic Parameters of Nicotine by Sex - PK Population**

		Group-1 PK			
		THS 2.2 Menthol (N=43)		mCC (N=43)	
Variable	Statistic	Males (N=23)	Females (N=20)	Males (N=23)	Females (N=20)
C <sub>max</sub> (ng/mL)	n	23	20	23	20
	Geometric Mean (CV%)	10.55 (141.69%)	10.75 (94.01%)	11.00 (63.65%)	13.55 (115.02%)
	95% CI	6.70, 16.62	7.40, 15.61	8.54, 14.16	8.81, 20.83
	Median	14.50	9.29	11.80	16.80
	Q25, Q75	6.44, 19.70	7.27, 18.10	7.97, 15.90	7.66, 29.70
	Min, Max	0.4, 31.3	1.4, 43.8	2.6, 31.5	2.3, 58.1
	Mean (SD)	14.44 (8.483)	14.32 (11.727)	12.81 (7.290)	18.88 (14.226)

Note: mCC = menthol conventional cigarettes; THS = Tobacco Heating System.

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

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_nic2.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.5.1 Descriptive Statistics of Pharmacokinetic Parameters of Nicotine by Sex - PK Population**

		Group-1 PK			
		THS 2.2 Menthol (N=43)		mCC (N=43)	
		Males (N=23)	Females (N=20)	Males (N=23)	Females (N=20)
Variable	Statistic				
AUC <sub>(0-last)</sub> (h*ng/mL)	n	23	20	23	20
	Geometric Mean (CV%)	25.76 (116.16%)	21.81 (106.47%)	28.67 (64.10%)	20.44 (99.42%)
	95% CI	17.27, 38.43	14.51, 32.78	22.24, 36.96	13.86, 30.13
	Median	28.86	18.73	31.29	21.04
	Q25, Q75	18.81, 47.04	13.31, 40.48	18.06, 39.61	12.58, 29.58
	Min, Max	1.2, 83.9	6.3, 117.7	8.2, 100.9	4.9, 88.3
	Mean (SD)	33.90 (21.381)	32.23 (32.503)	33.78 (20.951)	28.25 (24.613)

Note: mCC = menthol conventional cigarettes; THS = Tobacco Heating System.

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

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_nic2.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.5.1 Descriptive Statistics of Pharmacokinetic Parameters of Nicotine by Sex - PK Population**

Variable	Statistic	Group-1 PK			
		THS 2.2 Menthol (N=43)		mCC (N=43)	
		Males (N=23)	Females (N=20)	Males (N=23)	Females (N=20)
AUC <sub>(0-∞)</sub> (h*ng/mL)	n	19	18	22	18
	Geometric Mean (CV%)	31.40 (72.96%)	21.35 (94.05%)	31.86 (64.72%)	21.99 (105.16%)
	95% CI	22.92, 43.03	14.36, 31.72	24.51, 41.42	14.31, 33.78
	Median	32.65	17.97	33.80	21.85
	Q25, Q75	19.54, 52.86	14.03, 24.73	19.24, 42.10	12.14, 31.41
	Min, Max	5.7, 91.7	7.3, 111.0	9.6, 117.1	5.4, 93.2
	Mean (SD)	37.55 (22.106)	29.87 (28.966)	37.72 (24.220)	31.28 (28.264)

Note: mCC = menthol conventional cigarettes; THS = Tobacco Heating System.

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

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_nic2.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.5.1 Descriptive Statistics of Pharmacokinetic Parameters of Nicotine by Sex - PK Population**

Variable	Statistic	Group-1 PK			
		THS 2.2 Menthol (N=43)		mCC (N=43)	
		Males (N=23)	Females (N=20)	Males (N=23)	Females (N=20)
%AUC <sub>extrap</sub> (%)	n	19	18	22	18
	Geometric Mean (CV%)	7.84 (55.57%)	6.79 (64.95%)	7.92 (41.39%)	6.34 (54.20%)
	95% CI	6.10, 10.07	5.05, 9.12	6.63, 9.45	4.92, 8.17
	Median	8.55	6.53	7.57	6.19
	Q25, Q75	4.84, 11.01	4.32, 10.05	5.89, 9.57	4.33, 10.36
	Min, Max	2.8, 24.3	2.7, 19.9	4.3, 20.3	2.7, 15.5
	Mean (SD)	8.90 (4.893)	8.03 (4.952)	8.58 (3.827)	7.14 (3.573)

Note: mCC = menthol conventional cigarettes; THS = Tobacco Heating System.

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

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_nic2.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.5.1 Descriptive Statistics of Pharmacokinetic Parameters of Nicotine by Sex - PK Population**

Variable	Statistic	Group-1 PK			
		THS 2.2 Menthol (N=43)		mCC (N=43)	
		Males (N=23)	Females (N=20)	Males (N=23)	Females (N=20)
AUC <sub>(0-t)</sub> (h*ng/mL)	n	23	20	23	20
	Geometric Mean (CV%)	0.75 (197.15%)	0.45 (112.08%)	0.89 (91.58%)	0.70 (105.73%)
	95% CI	0.43, 1.29	0.29, 0.69	0.63, 1.25	0.46, 1.06
	Median	0.86	0.47	0.78	0.79
	Q25, Q75	0.28, 1.87	0.30, 0.73	0.49, 1.54	0.39, 1.29
	Min, Max	0.0, 4.7	0.0, 2.1	0.2, 4.2	0.1, 3.8
	Mean (SD)	1.32 (1.274)	0.62 (0.522)	1.18 (0.952)	0.97 (0.839)

Note: mCC = menthol conventional cigarettes; THS = Tobacco Heating System.

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

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_nic2.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.5.1 Descriptive Statistics of Pharmacokinetic Parameters of Nicotine by Sex - PK Population**

Variable	Statistic	Group-1 PK			
		THS 2.2 Menthol (N=43)		mCC (N=43)	
		Males (N=23)	Females (N=20)	Males (N=23)	Females (N=20)
t <sub>max</sub> (min)	n	23	20	23	20
	Median	8.00	6.00	8.00	5.00
	Q25, Q75	6.00, 15.00	4.00, 9.00	6.00, 10.00	4.00, 8.00
	Min, Max	4.0, 720.0	4.0, 720.0	4.0, 30.0	4.0, 15.0

Note: mCC = menthol conventional cigarettes; THS = Tobacco Heating System.

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

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_nic2.sas

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**Table 15.2.4.5.1 Descriptive Statistics of Pharmacokinetic Parameters of Nicotine by Sex - PK Population**

		Group-1 PK			
		THS 2.2 Menthol (N=43)		mCC (N=43)	
		Males (N=23)	Females (N=20)	Males (N=23)	Females (N=20)
Variable	Statistic				
t <sub>last</sub> (h)	n	23	20	23	20
	Median	24.00	12.00	24.00	9.00
	Q25, Q75	12.00, 24.00	9.00, 24.00	12.00, 24.00	9.00, 24.00
	Min, Max	6.0, 24.0	6.0, 24.0	9.0, 24.0	6.0, 24.0

Note: mCC = menthol conventional cigarettes; THS = Tobacco Heating System.

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

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_nic2.sas

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**Table 15.2.4.5.1 Descriptive Statistics of Pharmacokinetic Parameters of Nicotine by Sex - PK Population**

		Group-1 PK			
		THS 2.2 Menthol (N=43)		mCC (N=43)	
Variable	Statistic	Males (N=23)	Females (N=20)	Males (N=23)	Females (N=20)
t <sub>1/2</sub> (h)	n	19	18	22	18
	Geometric Mean (CV%)	4.63 (56.71%)	3.39 (71.79%)	5.06 (55.22%)	2.80 (62.38%)
	95% CI	3.58, 5.97	2.45, 4.67	4.02, 6.37	2.10, 3.73
	Median	6.18	2.99	6.00	2.02
	Q25, Q75	2.59, 7.33	1.98, 5.06	3.09, 7.30	1.83, 5.80
	Min, Max	1.9, 8.2	1.3, 12.0	1.9, 11.5	1.6, 7.5
	Mean (SD)	5.20 (2.307)	4.15 (2.910)	5.68 (2.583)	3.31 (2.125)

Note: mCC = menthol conventional cigarettes; THS = Tobacco Heating System.

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

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_nic2.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.5.1 Descriptive Statistics of Pharmacokinetic Parameters of Nicotine by Sex - PK Population**

		Group-1 PK			
		THS 2.2 Menthol (N=43)		mCC (N=43)	
Variable	Statistic	Males (N=23)	Females (N=20)	Males (N=23)	Females (N=20)
$\lambda_z$ (1/h)	n	19	18	22	18
	Geometric Mean (CV%)	0.15 (56.71%)	0.20 (71.79%)	0.14 (55.22%)	0.25 (62.38%)
	95% CI	0.11, 0.20	0.14, 0.29	0.10, 0.18	0.18, 0.33
	Median	0.11	0.23	0.12	0.34
	Q25, Q75	0.09, 0.27	0.13, 0.35	0.09, 0.23	0.11, 0.38
	Min, Max	0.1, 0.4	0.1, 0.5	0.1, 0.4	0.1, 0.4
	Mean (SD)	0.17 (0.097)	0.24 (0.141)	0.16 (0.088)	0.28 (0.127)

Note: mCC = menthol conventional cigarettes; THS = Tobacco Heating System.

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

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_nic2.sas

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**Table 15.2.4.5.1 Descriptive Statistics of Pharmacokinetic Parameters of Nicotine by Sex - PK Population**

		Group-2 PK			
		THS 2.2 Menthol (N=18)		NRT gum (N=18)	
Variable	Statistic	Males (N=9)	Females (N=9)	Males (N=9)	Females (N=9)
C <sub>max</sub> (ng/mL)	n	9	9	9	9
	Geometric Mean (CV%)	8.55 (120.97%)	6.83 (103.15%)	6.16 (43.85%)	9.17 (59.28%)
	95% CI	4.12, 17.74	3.54, 13.14	4.46, 8.51	6.01, 13.99
	Median	9.07	7.50	6.47	8.01
	Q25, Q75	4.94, 11.00	3.71, 11.50	5.18, 8.92	7.19, 10.20
	Min, Max	2.4, 45.8	1.3, 20.3	2.7, 10.0	4.9, 34.1
	Mean (SD)	12.95 (13.866)	8.88 (5.975)	6.61 (2.441)	10.90 (8.896)

Note: NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_nic2.sas

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**Table 15.2.4.5.1 Descriptive Statistics of Pharmacokinetic Parameters of Nicotine by Sex - PK Population**

		Group-2 PK			
		THS 2.2 Menthol (N=18)		NRT gum (N=18)	
Variable	Statistic	Males (N=9)	Females (N=9)	Males (N=9)	Females (N=9)
AUC <sub>(0-last)</sub> (h*ng/mL)	n	9	9	9	9
	Geometric Mean (CV%)	20.50 (61.35%)	11.88 (81.89%)	26.01 (67.01%)	30.01 (81.72%)
	95% CI	13.27, 31.66	6.85, 20.61	16.28, 41.54	17.31, 52.00
	Median	18.59	10.72	29.41	20.91
	Q25, Q75	16.41, 28.31	8.09, 15.64	19.61, 39.13	16.77, 68.37
	Min, Max	9.9, 48.3	3.9, 42.0	9.6, 58.0	15.4, 86.4
	Mean (SD)	23.75 (14.285)	15.07 (12.002)	30.10 (15.683)	38.20 (29.073)

Note: NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_nic2.sas

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**Table 15.2.4.5.1 Descriptive Statistics of Pharmacokinetic Parameters of Nicotine by Sex - PK Population**

		Group-2 PK			
		THS 2.2 Menthol (N=18)		NRT gum (N=18)	
		Males (N=9)	Females (N=9)	Males (N=9)	Females (N=9)
Variable	Statistic				
AUC <sub>(0-∞)</sub> (h*ng/mL)	n	8	8	8	9
	Geometric Mean (CV%)	23.17 (61.65%)	12.15 (72.67%)	27.17 (69.94%)	32.16 (83.27%)
	95% CI	14.41, 37.25	7.04, 20.95	16.03, 46.06	18.41, 56.19
	Median	19.62	11.22	29.43	21.65
	Q25, Q75	15.44, 39.73	8.46, 15.50	16.26, 45.20	17.72, 75.81
	Min, Max	11.6, 53.3	4.9, 45.2	11.0, 62.2	16.3, 91.7
	Mean (SD)	26.81 (16.017)	15.06 (12.726)	31.88 (17.976)	41.14 (31.304)

Note: NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_nic2.sas

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**Table 15.2.4.5.1 Descriptive Statistics of Pharmacokinetic Parameters of Nicotine by Sex - PK Population**

		Group-2 PK			
		THS 2.2 Menthol (N=18)		NRT gum (N=18)	
Variable	Statistic	Males (N=9)	Females (N=9)	Males (N=9)	Females (N=9)
%AUC <sub>extrap</sub> (%)	n	8	8	8	9
	Geometric Mean (CV%)	7.56 (67.91%)	9.87 (60.45%)	7.68 (31.53%)	5.95 (49.02%)
	95% CI	4.51, 12.66	6.19, 15.74	5.93, 9.94	4.16, 8.50
	Median	7.26	9.95	7.68	5.39
	Q25, Q75	5.44, 11.96	6.85, 16.15	6.43, 9.29	4.52, 6.19
	Min, Max	2.7, 19.0	4.0, 20.1	4.6, 12.6	3.4, 15.4
	Mean (SD)	8.88 (5.393)	11.24 (5.991)	8.00 (2.456)	6.63 (3.765)

Note: NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_nic2.sas

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**Table 15.2.4.5.1 Descriptive Statistics of Pharmacokinetic Parameters of Nicotine by Sex - PK Population**

		Group-2 PK			
		THS 2.2 Menthol (N=18)		NRT gum (N=18)	
Variable	Statistic	Males (N=9)	Females (N=9)	Males (N=9)	Females (N=9)
AUC <sub>(0-t)</sub> (h*ng/mL)	n	9	9	9	9
	Geometric Mean (CV%)	4.04 (84.99%)	2.83 (63.95%)	2.66 (55.15%)	3.32 (46.08%)
	95% CI	2.29, 7.12	1.80, 4.44	1.79, 3.96	2.36, 4.65
	Median	4.24	2.68	3.38	2.81
	Q25, Q75	2.83, 6.44	2.11, 3.86	2.10, 3.69	2.51, 4.06
	Min, Max	1.0, 11.2	1.2, 7.7	1.1, 4.7	1.9, 7.6
	Mean (SD)	4.98 (3.144)	3.29 (2.023)	2.95 (1.222)	3.64 (1.843)

Note: NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Appendix 15.3.3.1

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**Table 15.2.4.5.1 Descriptive Statistics of Pharmacokinetic Parameters of Nicotine by Sex - PK Population**

Variable	Statistic	Group-2 PK			
		THS 2.2 Menthol (N=18)		NRT gum (N=18)	
		Males (N=9)	Females (N=9)	Males (N=9)	Females (N=9)
t <sub>max</sub> (min)	n	9	9	9	9
	Median	10.00	6.00	60.00	45.00
	Q25, Q75	6.00, 10.00	4.00, 15.00	40.00, 60.00	40.00, 60.00
	Min, Max	4.0, 15.0	4.0, 30.0	30.0, 60.0	25.0, 60.0

Note: NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_nic2.sas

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**Table 15.2.4.5.1 Descriptive Statistics of Pharmacokinetic Parameters of Nicotine by Sex - PK Population**

		Group-2 PK			
		THS 2.2 Menthol (N=18)		NRT gum (N=18)	
Variable	Statistic	Males (N=9)	Females (N=9)	Males (N=9)	Females (N=9)
t <sub>last</sub> (h)	n	9	9	9	9
	Median	12.00	9.00	12.00	12.00
	Q25, Q75	12.00, 24.00	6.00, 12.00	12.00, 24.00	12.00, 24.00
	Min, Max	9.0, 24.0	4.0, 24.0	6.0, 24.0	6.0, 24.0

Note: NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Appendix 15.3.3.1

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**Table 15.2.4.5.1 Descriptive Statistics of Pharmacokinetic Parameters of Nicotine by Sex - PK Population**

		Group-2 PK			
		THS 2.2 Menthol (N=18)		NRT gum (N=18)	
Variable	Statistic	Males (N=9)	Females (N=9)	Males (N=9)	Females (N=9)
t <sub>1/2</sub> (h)	n	8	8	8	9
	Geometric Mean (CV%)	4.17 (55.90%)	2.68 (64.46%)	3.88 (57.44%)	3.76 (78.91%)
	95% CI	2.69, 6.45	1.63, 4.40	2.48, 6.07	2.20, 6.43
	Median	4.60	2.50	3.70	2.80
	Q25, Q75	2.84, 6.46	1.65, 4.45	2.92, 6.17	2.49, 5.89
	Min, Max	1.7, 7.7	1.3, 6.6	1.5, 7.8	1.3, 11.6
	Mean (SD)	4.64 (2.132)	3.13 (1.891)	4.36 (2.133)	4.65 (3.280)

Note: NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Appendix 15.3.3.1

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**Table 15.2.4.5.1 Descriptive Statistics of Pharmacokinetic Parameters of Nicotine by Sex - PK Population**

Variable	Statistic	Group-2 PK			
		THS 2.2 Menthol (N=18)		NRT gum (N=18)	
		Males (N=9)	Females (N=9)	Males (N=9)	Females (N=9)
$\lambda_z$ (1/h)	n	8	8	8	9
	Geometric Mean (CV%)	0.17 (55.90%)	0.26 (64.46%)	0.18 (57.44%)	0.18 (78.91%)
	95% CI	0.10, 0.26	0.15, 0.43	0.11, 0.28	0.10, 0.32
	Median	0.15	0.28	0.19	0.25
	Q25, Q75	0.10, 0.25	0.15, 0.42	0.11, 0.24	0.11, 0.28
	Min, Max	0.1, 0.4	0.1, 0.6	0.1, 0.5	0.1, 0.6
	Mean (SD)	0.19 (0.108)	0.30 (0.158)	0.20 (0.118)	0.23 (0.152)

Note: NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_nic2.sas

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**Table 15.2.4.5.2 Descriptive Statistics of Pharmacokinetic Parameters of Nicotine by Nicotine Level - PK Population**

Variable	Statistic	Group-1 PK			
		THS 2.2 Menthol (N=43)		mCC (N=43)	
		<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)	<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)
C <sub>max</sub> (ng/mL)	n	25	18	25	18
	Geometric Mean (CV%)	14.20 (52.38%)	7.13 (186.47%)	11.96 (71.52%)	12.35 (113.31%)
	95% CI	11.59, 17.41	3.88, 13.12	9.17, 15.60	7.85, 19.41
	Median	14.10	7.27	12.10	12.20
	Q25, Q75	9.33, 19.70	4.91, 18.60	8.40, 17.90	6.93, 28.60
	Min, Max	5.9, 38.1	0.4, 43.8	2.6, 34.2	2.3, 58.1
	Mean (SD)	15.98 (8.339)	12.16 (11.818)	14.32 (8.569)	17.45 (14.439)

Note: mCC = menthol conventional cigarettes; THS = Tobacco Heating System.

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

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_nic3.sas

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**Table 15.2.4.5.2 Descriptive Statistics of Pharmacokinetic Parameters of Nicotine by Nicotine Level - PK Population**

Variable	Statistic	Group-1 PK			
		THS 2.2 Menthol (N=43)		mCC (N=43)	
		<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)	<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)
AUC <sub>(0-last)</sub> (h*ng/mL)	n	25	18	25	18
	Geometric Mean (CV%)	28.08 (90.91%)	19.00 (133.72%)	21.56 (74.48%)	29.25 (91.21%)
	95% CI	20.38, 38.69	11.48, 31.44	16.38, 28.36	19.86, 43.08
	Median	27.52	19.46	20.64	28.93
	Q25, Q75	18.03, 52.29	13.51, 29.01	14.49, 31.75	17.57, 61.08
	Min, Max	6.3, 117.7	1.2, 83.9	4.9, 60.8	7.2, 100.9
	Mean (SD)	36.97 (28.689)	27.77 (23.665)	26.05 (15.576)	38.37 (28.783)

Note: mCC = menthol conventional cigarettes; THS = Tobacco Heating System.

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

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_nic3.sas

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**Table 15.2.4.5.2 Descriptive Statistics of Pharmacokinetic Parameters of Nicotine by Nicotine Level - PK Population**

Variable	Statistic	Group-1 PK			
		THS 2.2 Menthol (N=43)		mCC (N=43)	
		<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)	<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)
AUC <sub>(0-∞)</sub> (h*ng/mL)	n	21	16	22	18
	Geometric Mean (CV%)	26.99 (85.35%)	24.81 (89.48%)	23.35 (80.03%)	32.16 (89.23%)
	95% CI	19.27, 37.81	16.48, 37.34	17.09, 31.90	21.98, 47.06
	Median	24.73	22.10	22.39	31.92
	Q25, Q75	17.41, 50.30	15.77, 42.76	14.68, 40.20	18.94, 66.36
	Min, Max	7.3, 111.0	5.7, 91.7	5.4, 69.0	8.5, 117.1
	Mean (SD)	34.61 (25.527)	32.76 (26.510)	28.87 (18.276)	42.09 (32.143)

Note: mCC = menthol conventional cigarettes; THS = Tobacco Heating System.

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

Appendix 15.3.3.1

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**Table 15.2.4.5.2 Descriptive Statistics of Pharmacokinetic Parameters of Nicotine by Nicotine Level - PK Population**

Variable	Statistic	Group-1 PK			
		THS 2.2 Menthol (N=43)		mCC (N=43)	
		<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)	<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)
%AUC <sub>extrap</sub> (%)	n	21	16	22	18
	Geometric Mean (CV%)	5.98 (55.67%)	9.51 (52.66%)	6.59 (43.93%)	7.93 (52.52%)
	95% CI	4.72, 7.58	7.30, 12.38	5.47, 7.95	6.20, 10.14
	Median	6.86	9.01	7.09	7.59
	Q25, Q75	4.16, 9.03	6.52, 13.82	5.17, 9.38	4.88, 11.80
	Min, Max	2.7, 15.7	4.6, 24.3	2.7, 11.8	4.2, 20.3
	Mean (SD)	6.79 (3.527)	10.69 (5.585)	7.12 (2.664)	8.92 (4.625)

Note: mCC = menthol conventional cigarettes; THS = Tobacco Heating System.

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

Appendix 15.3.3.1

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**Table 15.2.4.5.2 Descriptive Statistics of Pharmacokinetic Parameters of Nicotine by Nicotine Level - PK Population**

Variable	Statistic	Group-1 PK			
		THS 2.2 Menthol (N=43)		mCC (N=43)	
		<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)	<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)
AUC <sub>(0-t)</sub> (h*ng/mL)	n	25	18	25	18
	Geometric Mean (CV%)	0.79 (124.52%)	0.39 (185.47%)	0.82 (93.64%)	0.76 (107.46%)
	95% CI	0.53, 1.19	0.21, 0.72	0.59, 1.15	0.48, 1.18
	Median	0.75	0.48	0.86	0.68
	Q25, Q75	0.44, 1.70	0.17, 0.87	0.48, 1.36	0.45, 1.38
	Min, Max	0.1, 4.7	0.0, 2.1	0.1, 4.2	0.2, 3.8
	Mean (SD)	1.22 (1.223)	0.67 (0.638)	1.09 (0.885)	1.07 (0.938)

Note: mCC = menthol conventional cigarettes; THS = Tobacco Heating System.

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

Appendix 15.3.3.1

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**Table 15.2.4.5.2 Descriptive Statistics of Pharmacokinetic Parameters of Nicotine by Nicotine Level - PK Population**

Variable	Statistic	Group-1 PK			
		THS 2.2 Menthol (N=43)		mCC (N=43)	
		<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)	<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)
t <sub>max</sub> (min)	n	25	18	25	18
	Median	6.00	6.00	6.00	7.00
	Q25, Q75	6.00, 10.00	6.00, 15.00	4.00, 10.00	6.00, 8.00
	Min, Max	4.0, 720.0	4.0, 720.0	4.0, 30.0	4.0, 15.0

Note: mCC = menthol conventional cigarettes; THS = Tobacco Heating System.

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

Appendix 15.3.3.1

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**Table 15.2.4.5.2 Descriptive Statistics of Pharmacokinetic Parameters of Nicotine by Nicotine Level - PK Population**

Variable		Group-1 PK			
		THS 2.2 Menthol (N=43)		mCC (N=43)	
		<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)	<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)
t <sub>last</sub> (h)	n	25	18	25	18
	Median	24.00	12.00	12.00	24.00
	Q25, Q75	9.00, 24.00	12.00, 24.00	9.00, 24.00	9.00, 24.00
	Min, Max	6.0, 24.0	6.0, 24.0	6.0, 24.0	9.0, 24.0

Note: mCC = menthol conventional cigarettes; THS = Tobacco Heating System.

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

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**Table 15.2.4.5.2 Descriptive Statistics of Pharmacokinetic Parameters of Nicotine by Nicotine Level - PK Population**

Variable	Statistic	Group-1 PK			
		THS 2.2 Menthol (N=43)		mCC (N=43)	
		<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)	<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)
t <sub>1/2</sub> (h)	n	21	16	22	18
	Geometric Mean (CV%)	3.46 (68.06%)	4.77 (58.61%)	3.28 (67.16%)	4.76 (61.54%)
	95% CI	2.61, 4.58	3.57, 6.38	2.50, 4.30	3.58, 6.31
	Median	2.59	5.76	2.91	5.90
	Q25, Q75	2.16, 6.18	3.11, 7.51	1.88, 6.25	3.08, 7.06
	Min, Max	1.3, 12.0	1.4, 9.3	1.6, 8.4	1.9, 11.5
	Mean (SD)	4.16 (2.706)	5.39 (2.449)	3.92 (2.408)	5.46 (2.736)

Note: mCC = menthol conventional cigarettes; THS = Tobacco Heating System.

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

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_nic3.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.5.2 Descriptive Statistics of Pharmacokinetic Parameters of Nicotine by Nicotine Level - PK Population**

Variable	Statistic	Group-1 PK			
		THS 2.2 Menthol (N=43)		mCC (N=43)	
		<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)	<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)
$\lambda_z$ (1/h)	n	21	16	22	18
	Geometric Mean (CV%)	0.20 (68.06%)	0.15 (58.61%)	0.21 (67.16%)	0.15 (61.54%)
	95% CI	0.15, 0.27	0.10, 0.20	0.16, 0.28	0.10, 0.20
	Median	0.27	0.12	0.24	0.12
	Q25, Q75	0.11, 0.32	0.09, 0.23	0.11, 0.37	0.09, 0.23
	Min, Max	0.1, 0.5	0.1, 0.5	0.1, 0.4	0.1, 0.4
	Mean (SD)	0.24 (0.130)	0.17 (0.109)	0.25 (0.130)	0.17 (0.103)

Note: mCC = menthol conventional cigarettes; THS = Tobacco Heating System.

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

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_nic3.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.5.2 Descriptive Statistics of Pharmacokinetic Parameters of Nicotine by Nicotine Level - PK Population**

Variable	Statistic	Group-2 PK			
		THS 2.2 Menthol (N=18)		NRT gum (N=18)	
		<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)	<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)
C <sub>max</sub> (ng/mL)	n	10	8	10	8
	Geometric Mean (CV%)	6.79 (143.77%)	8.86 (70.64%)	8.87 (56.33%)	6.11 (47.62%)
	95% CI	3.18, 14.47	5.20, 15.09	6.09, 12.92	4.18, 8.92
	Median	6.06	9.99	8.08	6.49
	Q25, Q75	3.51, 11.50	7.23, 12.50	6.79, 9.37	4.61, 9.00
	Min, Max	1.3, 45.8	2.4, 20.3	5.2, 34.1	2.7, 10.2
	Mean (SD)	11.44 (13.720)	10.26 (5.321)	10.46 (8.489)	6.63 (2.668)

Note: NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_nic3.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.5.2 Descriptive Statistics of Pharmacokinetic Parameters of Nicotine by Nicotine Level - PK Population**

Variable	Statistic	Group-2 PK			
		THS 2.2 Menthol (N=18)		NRT gum (N=18)	
		<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)	<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)
AUC <sub>(0-last)</sub> (h*ng/mL)	n	10	8	10	8
	Geometric Mean (CV%)	14.56 (98.63%)	17.03 (53.75%)	33.90 (66.74%)	21.94 (73.06%)
	95% CI	8.07, 26.26	11.17, 25.95	21.95, 52.33	12.69, 37.91
	Median	14.26	17.50	32.60	19.99
	Q25, Q75	8.09, 28.31	10.68, 22.62	20.91, 58.00	13.93, 34.21
	Min, Max	3.9, 48.3	9.7, 42.0	15.4, 86.4	9.6, 68.4
	Mean (SD)	19.61 (16.016)	19.17 (10.789)	40.04 (24.769)	26.78 (19.718)

Note: NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_nic3.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.5.2 Descriptive Statistics of Pharmacokinetic Parameters of Nicotine by Nicotine Level - PK Population**

Variable	Statistic	Group-2 PK			
		THS 2.2 Menthol (N=18)		NRT gum (N=18)	
		<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)	<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)
AUC <sub>(0-∞)</sub> (h*ng/mL)	n	10	6	9	8
	Geometric Mean (CV%)	16.37 (91.24%)	17.49 (57.36%)	35.89 (72.56%)	24.02 (73.97%)
	95% CI	9.37, 28.56	9.99, 30.62	21.76, 59.17	13.82, 41.73
	Median	15.50	16.13	31.31	20.89
	Q25, Q75	8.87, 30.93	12.06, 20.14	21.64, 62.22	14.85, 38.78
	Min, Max	4.9, 53.3	10.4, 45.2	16.3, 91.7	11.0, 75.8
	Mean (SD)	21.49 (17.105)	20.01 (12.962)	43.27 (27.977)	29.49 (22.034)

Note: NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_nic3.sas

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**Table 15.2.4.5.2 Descriptive Statistics of Pharmacokinetic Parameters of Nicotine by Nicotine Level - PK Population**

Variable	Statistic	Group-2 PK			
		THS 2.2 Menthol (N=18)		NRT gum (N=18)	
		<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)	<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)
%AUC <sub>extrap</sub> (%)	n	10	6	9	8
	Geometric Mean (CV%)	9.55 (59.07%)	7.30 (73.39%)	5.89 (30.25%)	7.77 (51.27%)
	95% CI	6.46, 14.14	3.66, 14.55	4.68, 7.39	5.19, 11.65
	Median	9.11	6.86	6.08	8.40
	Q25, Q75	6.04, 14.46	5.63, 11.15	5.10, 6.79	4.98, 11.23
	Min, Max	4.0, 20.1	2.7, 19.0	3.4, 9.1	4.1, 15.4
	Mean (SD)	10.88 (5.728)	8.70 (5.728)	6.11 (1.735)	8.59 (4.026)

Note: NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_nic3.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.5.2 Descriptive Statistics of Pharmacokinetic Parameters of Nicotine by Nicotine Level - PK Population**

		Group-2 PK			
		THS 2.2 Menthol (N=18)		NRT gum (N=18)	
Variable	Statistic	<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)	<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)
AUC <sub>(0-t)</sub> (h*ng/mL)	n	10	8	10	8
	Geometric Mean (CV%)	3.00 (95.87%)	3.92 (48.59%)	3.23 (40.07%)	2.68 (63.71%)
	95% CI	1.68, 5.35	2.66, 5.76	2.45, 4.27	1.64, 4.36
	Median	2.87	3.91	3.27	2.74
	Q25, Q75	1.45, 5.56	2.75, 5.69	2.44, 3.69	1.90, 4.27
	Min, Max	1.0, 11.2	2.0, 7.7	1.9, 7.6	1.1, 5.6
	Mean (SD)	4.01 (3.285)	4.29 (1.967)	3.48 (1.600)	3.06 (1.581)

Note: NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_nic3.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.5.2 Descriptive Statistics of Pharmacokinetic Parameters of Nicotine by Nicotine Level - PK Population**

Variable	Statistic	Group-2 PK			
		THS 2.2 Menthol (N=18)		NRT gum (N=18)	
		<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)	<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)
t <sub>max</sub> (min)	n	10	8	10	8
	Median	8.00	8.00	45.00	52.50
	Q25, Q75	4.00, 10.00	5.00, 15.00	40.00, 60.00	37.50, 60.00
	Min, Max	4.0, 30.0	4.0, 15.0	25.0, 60.0	35.0, 60.0

Note: NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_nic3.sas

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**Table 15.2.4.5.2 Descriptive Statistics of Pharmacokinetic Parameters of Nicotine by Nicotine Level - PK Population**

Variable		Group-2 PK			
		THS 2.2 Menthol (N=18)		NRT gum (N=18)	
		<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)	<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)
t <sub>last</sub> (h)	n	10	8	10	8
	Median	12.00	12.00	24.00	12.00
	Q25, Q75	6.00, 24.00	9.00, 24.00	12.00, 24.00	12.00, 18.00
	Min, Max	4.0, 24.0	9.0, 24.0	6.0, 24.0	6.0, 24.0

Note: NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_nic3.sas

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**Table 15.2.4.5.2 Descriptive Statistics of Pharmacokinetic Parameters of Nicotine by Nicotine Level - PK Population**

Variable	Statistic	Group-2 PK			
		THS 2.2 Menthol (N=18)		NRT gum (N=18)	
		<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)	<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)
t <sub>1/2</sub> (h)	n	10	6	9	8
	Geometric Mean (CV%)	3.45 (71.86%)	3.17 (55.33%)	3.91 (67.77%)	3.72 (70.69%)
	95% CI	2.17, 5.49	1.84, 5.45	2.44, 6.28	2.18, 6.33
	Median	3.95	2.84	5.56	3.04
	Q25, Q75	1.69, 6.06	2.16, 5.18	2.49, 5.90	2.72, 5.72
	Min, Max	1.3, 7.7	1.7, 6.6	1.3, 7.8	1.5, 11.6
	Mean (SD)	4.09 (2.277)	3.55 (1.912)	4.52 (2.260)	4.51 (3.326)

Note: NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_nic3.sas

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**Table 15.2.4.5.2 Descriptive Statistics of Pharmacokinetic Parameters of Nicotine by Nicotine Level - PK Population**

Variable	Statistic	Group-2 PK			
		THS 2.2 Menthol (N=18)		NRT gum (N=18)	
		<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)	<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)
$\lambda_z$ (1/h)	n	10	6	9	8
	Geometric Mean (CV%)	0.20 (71.86%)	0.22 (55.33%)	0.18 (67.77%)	0.19 (70.69%)
	95% CI	0.12, 0.32	0.12, 0.38	0.11, 0.29	0.10, 0.32
	Median	0.18	0.24	0.12	0.23
	Q25, Q75	0.11, 0.42	0.13, 0.32	0.11, 0.28	0.13, 0.26
	Min, Max	0.1, 0.6	0.1, 0.4	0.1, 0.6	0.1, 0.5
	Mean (SD)	0.24 (0.163)	0.24 (0.114)	0.21 (0.151)	0.22 (0.121)

Note: NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Appendix 15.3.3.1

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_nic3.sas

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**Table 15.2.4.6 Descriptive Statistics of the Plasma Nicotine Concentrations (ng/mL) - PK Population**

Timepoint	Statistic	Group-1 PK		Group-2 PK	
		THS 2.2 Menthol (N=43)	mCC (N=43)	THS 2.2 Menthol (N=18)	NRT gum (N=18)
15 min < T <sub>0</sub>	n	43	43	18	18
	BLOQ - n (%)	23 (53.5%)	22 (51.2%)	12 (66.7%)	7 (38.9%)
	Geometric Mean (CV%)	NC	NC	NC	NC
	95% CI	NC	NC	NC	NC
	Median	0.00	0.00	0.00	0.22
	Q25, Q75	0.00, 0.33	0.00, 0.36	0.00, 0.27	0.00, 0.44
	Min, Max	0.0, 2.2	0.0, 2.0	0.0, 0.9	0.0, 0.9
	Mean (SD)	0.23 (0.396)	0.27 (0.426)	0.16 (0.279)	0.25 (0.264)

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Note: NC = Not calculated.

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: BLOQ values before T<sub>0</sub> are treated as zero. BLOQ values after the last quantifiable value are treated as missing. Any other BLOQ values are imputed by LLOQ/2.

Note: LLOQ = 0.2 ng/mL.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_pkconc.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.6 Descriptive Statistics of the Plasma Nicotine Concentrations (ng/mL) - PK Population**

Timepoint	Statistic	Group-1 PK		Group-2 PK	
		THS 2.2 Menthol (N=43)	mCC (N=43)	THS 2.2 Menthol (N=18)	NRT gum (N=18)
T <sub>0</sub> + 2 min	n	43	43	18	0
	BLOQ - n (%)	0	1 (2.3%)	0	
	Geometric Mean (CV%)	2.36 (186.95%)	3.32 (187.74%)	2.66 (144.75%)	
	95% CI	1.61, 3.45	2.27, 4.86	1.56, 4.52	
	Median	2.62	2.76	2.58	
	Q25, Q75	1.22, 6.42	1.69, 8.15	1.47, 4.81	
	Min, Max	0.2, 35.1	0.1, 36.6	0.3, 30.2	
	Mean (SD)	4.67 (6.336)	6.45 (8.097)	4.66 (6.775)	

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: BLOQ values before T<sub>0</sub> are treated as zero. BLOQ values after the last quantifiable value are treated as missing. Any other BLOQ values are imputed by LLOQ/2.

Note: LLOQ = 0.2 ng/mL.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_pkconc.sas

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**Table 15.2.4.6 Descriptive Statistics of the Plasma Nicotine Concentrations (ng/mL) - PK Population**

Timepoint	Statistic	Group-1 PK		Group-2 PK	
		THS 2.2 Menthol (N=43)	mCC (N=43)	THS 2.2 Menthol (N=18)	NRT gum (N=18)
T <sub>0</sub> + 4 min	n	43	43	18	0
	BLOQ - n (%)	0	0	0	
	Geometric Mean (CV%)	6.52 (153.26%)	8.11 (137.46%)	5.59 (138.48%)	
	95% CI	4.64, 9.14	5.90, 11.14	3.34, 9.36	
	Median	7.23	8.13	5.52	
	Q25, Q75	3.92, 14.80	4.71, 19.30	3.45, 11.50	
	Min, Max	0.3, 38.1	0.4, 47.6	0.6, 45.3	
	Mean (SD)	10.24 (8.943)	12.38 (10.874)	9.07 (10.652)	

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: BLOQ values before T<sub>0</sub> are treated as zero. BLOQ values after the last quantifiable value are treated as missing. Any other BLOQ values are imputed by LLOQ/2.

Note: LLOQ = 0.2 ng/mL.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_pkconc.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.6 Descriptive Statistics of the Plasma Nicotine Concentrations (ng/mL) - PK Population**

Timepoint	Statistic	Group-1 PK		Group-2 PK	
		THS 2.2 Menthol (N=43)	mCC (N=43)	THS 2.2 Menthol (N=18)	NRT gum (N=18)
T <sub>0</sub> + 6 min	n	43	43	18	0
	BLOQ - n (%)	0	0	0	
	Geometric Mean (CV%)	8.29 (129.69%)	9.21 (98.73%)	6.49 (112.24%)	
	95% CI	6.10, 11.26	7.14, 11.88	4.14, 10.18	
	Median	9.16	9.13	7.16	
	Q25, Q75	5.30, 17.60	5.47, 17.40	3.65, 9.84	
	Min, Max	0.4, 43.8	1.2, 58.1	1.2, 45.8	
	Mean (SD)	11.93 (9.404)	12.51 (10.610)	9.69 (10.609)	

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: BLOQ values before T<sub>0</sub> are treated as zero. BLOQ values after the last quantifiable value are treated as missing. Any other BLOQ values are imputed by LLOQ/2.

Note: LLOQ = 0.2 ng/mL.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_pkconc.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.6 Descriptive Statistics of the Plasma Nicotine Concentrations (ng/mL) - PK Population**

Timepoint	Statistic	Group-1 PK		Group-2 PK	
		THS 2.2 Menthol (N=43)	mCC (N=43)	THS 2.2 Menthol (N=18)	NRT gum (N=18)
T <sub>0</sub> + 8 min	n	43	43	18	0
	BLOQ - n (%)	0	0	0	
	Geometric Mean (CV%)	7.86 (113.03%)	8.57 (76.14%)	5.63 (92.24%)	
	95% CI	5.94, 10.40	6.96, 10.56	3.81, 8.32	
	Median	7.91	8.48	5.76	
	Q25, Q75	5.05, 15.90	6.06, 13.60	3.00, 8.63	
	Min, Max	0.4, 29.7	1.4, 36.0	1.2, 28.9	
	Mean (SD)	10.58 (7.231)	10.47 (6.707)	7.56 (6.701)	

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: BLOQ values before T<sub>0</sub> are treated as zero. BLOQ values after the last quantifiable value are treated as missing. Any other BLOQ values are imputed by LLOQ/2.

Note: LLOQ = 0.2 ng/mL.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_pkconc.sas

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**Table 15.2.4.6 Descriptive Statistics of the Plasma Nicotine Concentrations (ng/mL) - PK Population**

Timepoint	Statistic	Group-1 PK		Group-2 PK	
		THS 2.2 Menthol (N=43)	mCC (N=43)	THS 2.2 Menthol (N=18)	NRT gum (N=18)
T <sub>0</sub> + 10 min	n	43	43	18	18
	BLOQ - n (%)	0	0	0	0
	Geometric Mean (CV%)	7.30 (111.15%)	8.45 (67.95%)	5.67 (85.97%)	1.42 (57.33%)
	95% CI	5.53, 9.62	6.98, 10.22	3.91, 8.22	1.08, 1.85
	Median	8.17	8.54	6.67	1.55
	Q25, Q75	4.32, 14.00	6.02, 12.90	3.56, 8.17	0.98, 1.94
	Min, Max	0.3, 26.3	2.3, 31.5	1.1, 22.1	0.7, 4.2
	Mean (SD)	9.63 (6.246)	10.06 (6.126)	7.18 (5.090)	1.62 (0.893)

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: BLOQ values before T<sub>0</sub> are treated as zero. BLOQ values after the last quantifiable value are treated as missing. Any other BLOQ values are imputed by LLOQ/2.

Note: LLOQ = 0.2 ng/mL.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_pkconc.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.6 Descriptive Statistics of the Plasma Nicotine Concentrations (ng/mL) - PK Population**

Timepoint	Statistic	Group-1 PK		Group-2 PK	
		THS 2.2 Menthol (N=43)	mCC (N=43)	THS 2.2 Menthol (N=18)	NRT gum (N=18)
T <sub>0</sub> + 15 min	n	43	43	18	0
	BLOQ - n (%)	0	0	0	
	Geometric Mean (CV%)	6.12 (93.33%)	6.49 (57.01%)	4.71 (69.37%)	
	95% CI	4.80, 7.82	5.51, 7.65	3.44, 6.44	
	Median	6.79	6.96	4.43	
	Q25, Q75	4.62, 10.70	5.28, 8.76	3.33, 7.61	
	Min, Max	0.3, 20.4	1.3, 20.0	1.3, 13.4	
	Mean (SD)	7.65 (4.497)	7.33 (3.523)	5.60 (3.312)	

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: BLOQ values before T<sub>0</sub> are treated as zero. BLOQ values after the last quantifiable value are treated as missing. Any other BLOQ values are imputed by LLOQ/2.

Note: LLOQ = 0.2 ng/mL.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_pkconc.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.6 Descriptive Statistics of the Plasma Nicotine Concentrations (ng/mL) - PK Population**

Timepoint	Statistic	Group-1 PK		Group-2 PK	
		THS 2.2 Menthol (N=43)	mCC (N=43)	THS 2.2 Menthol (N=18)	NRT gum (N=18)
T <sub>0</sub> + 20 min	n	0	0	0	18
	BLOQ - n (%)				0
	Geometric Mean (CV%)				3.60 (51.02%)
	95% CI				2.83, 4.58
	Median				3.41
	Q25, Q75				2.45, 5.76
	Min, Max				1.8, 7.0
	Mean (SD)				4.00 (1.841)

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: BLOQ values before T<sub>0</sub> are treated as zero. BLOQ values after the last quantifiable value are treated as missing. Any other BLOQ values are imputed by LLOQ/2.

Note: LLOQ = 0.2 ng/mL.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_pkconc.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.6 Descriptive Statistics of the Plasma Nicotine Concentrations (ng/mL) - PK Population**

Timepoint	Statistic	Group-1 PK		Group-2 PK	
		THS 2.2 Menthol (N=43)	mCC (N=43)	THS 2.2 Menthol (N=18)	NRT gum (N=18)
T <sub>0</sub> + 25 min	n	0	0	0	18
	BLOQ - n (%)				0
	Geometric Mean (CV%)				4.15 (43.36%)
	95% CI				3.37, 5.11
	Median				3.78
	Q25, Q75				2.87, 5.78
	Min, Max				2.2, 8.6
	Mean (SD)				4.51 (1.943)

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: BLOQ values before T<sub>0</sub> are treated as zero. BLOQ values after the last quantifiable value are treated as missing. Any other BLOQ values are imputed by LLOQ/2.

Note: LLOQ = 0.2 ng/mL.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_pkconc.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.6 Descriptive Statistics of the Plasma Nicotine Concentrations (ng/mL) - PK Population**

Timepoint	Statistic	Group-1 PK		Group-2 PK	
		THS 2.2 Menthol (N=43)	mCC (N=43)	THS 2.2 Menthol (N=18)	NRT gum (N=18)
T <sub>0</sub> + 30 min	n	43	43	18	18
	BLOQ - n (%)	0	0	0	0
	Geometric Mean (CV%)	5.08 (85.00%)	5.46 (55.60%)	3.83 (63.00%)	4.94 (35.52%)
	95% CI	4.04, 6.38	4.65, 6.42	2.87, 5.11	4.16, 5.87
	Median	6.08	5.68	4.03	4.83
	Q25, Q75	3.88, 7.31	4.41, 7.87	2.49, 6.33	4.12, 6.13
	Min, Max	0.3, 13.3	1.8, 14.7	1.3, 9.3	2.5, 8.9
	Mean (SD)	6.16 (3.258)	6.16 (2.912)	4.43 (2.321)	5.23 (1.786)

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: BLOQ values before T<sub>0</sub> are treated as zero. BLOQ values after the last quantifiable value are treated as missing. Any other BLOQ values are imputed by LLOQ/2.

Note: LLOQ = 0.2 ng/mL.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_pkconc.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.6 Descriptive Statistics of the Plasma Nicotine Concentrations (ng/mL) - PK Population**

Timepoint	Statistic	Group-1 PK		Group-2 PK	
		THS 2.2 Menthol (N=43)	mCC (N=43)	THS 2.2 Menthol (N=18)	NRT gum (N=18)
T <sub>0</sub> + 35 min	n	0	0	0	18
	BLOQ - n (%)				0
	Geometric Mean (CV%)				5.11 (39.60%)
	95% CI				4.22, 6.18
	Median				5.20
	Q25, Q75				4.29, 6.31
	Min, Max				2.3, 9.8
	Mean (SD)				5.45 (1.969)

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: BLOQ values before T<sub>0</sub> are treated as zero. BLOQ values after the last quantifiable value are treated as missing. Any other BLOQ values are imputed by LLOQ/2.

Note: LLOQ = 0.2 ng/mL.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_pkconc.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.6 Descriptive Statistics of the Plasma Nicotine Concentrations (ng/mL) - PK Population**

Timepoint	Statistic	Group-1 PK		Group-2 PK	
		THS 2.2 Menthol (N=43)	mCC (N=43)	THS 2.2 Menthol (N=18)	NRT gum (N=18)
T <sub>0</sub> + 40 min	n	0	0	0	18
	BLOQ - n (%)				0
	Geometric Mean (CV%)				5.98 (35.25%)
	95% CI				5.04, 7.09
	Median				6.41
	Q25, Q75				4.72, 7.72
	Min, Max				2.7, 9.4
	Mean (SD)				6.29 (1.918)

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: BLOQ values before T<sub>0</sub> are treated as zero. BLOQ values after the last quantifiable value are treated as missing. Any other BLOQ values are imputed by LLOQ/2.

Note: LLOQ = 0.2 ng/mL.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_pkconc.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.6 Descriptive Statistics of the Plasma Nicotine Concentrations (ng/mL) - PK Population**

Timepoint	Statistic	Group-1 PK		Group-2 PK	
		THS 2.2 Menthol (N=43)	mCC (N=43)	THS 2.2 Menthol (N=18)	NRT gum (N=18)
T <sub>0</sub> + 45 min	n	43	43	18	18
	BLOQ - n (%)	1 (2.3%)	0	0	0
	Geometric Mean (CV%)	4.52 (98.65%)	5.01 (55.02%)	3.51 (63.70%)	6.05 (37.89%)
	95% CI	3.50, 5.83	4.28, 5.88	2.62, 4.69	5.04, 7.27
	Median	5.28	5.51	4.00	6.26
	Q25, Q75	3.64, 7.21	3.80, 7.05	2.15, 5.58	4.51, 7.54
	Min, Max	0.1, 13.6	1.5, 15.2	1.2, 8.3	2.5, 11.1
	Mean (SD)	5.56 (2.886)	5.65 (2.754)	4.07 (2.131)	6.42 (2.195)

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: BLOQ values before T<sub>0</sub> are treated as zero. BLOQ values after the last quantifiable value are treated as missing. Any other BLOQ values are imputed by LLOQ/2.

Note: LLOQ = 0.2 ng/mL.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_pkconc.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.6 Descriptive Statistics of the Plasma Nicotine Concentrations (ng/mL) - PK Population**

Timepoint	Statistic	Group-1 PK		Group-2 PK	
		THS 2.2 Menthol (N=43)	mCC (N=43)	THS 2.2 Menthol (N=18)	NRT gum (N=18)
T <sub>0</sub> + 60 min	n	43	43	18	18
	BLOQ - n (%)	0	0	0	0
	Geometric Mean (CV%)	4.32 (85.33%)	4.64 (57.68%)	3.12 (62.92%)	6.70 (56.77%)
	95% CI	3.44, 5.43	3.93, 5.47	2.34, 4.16	5.15, 8.72
	Median	4.77	5.14	3.20	6.10
	Q25, Q75	3.72, 6.71	3.46, 6.91	2.10, 5.16	5.18, 7.58
	Min, Max	0.2, 12.3	1.5, 13.4	1.1, 7.8	2.4, 34.1
	Mean (SD)	5.24 (2.774)	5.27 (2.619)	3.63 (2.008)	7.96 (6.830)

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: BLOQ values before T<sub>0</sub> are treated as zero. BLOQ values after the last quantifiable value are treated as missing. Any other BLOQ values are imputed by LLOQ/2.

Note: LLOQ = 0.2 ng/mL.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_pkconc.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.6 Descriptive Statistics of the Plasma Nicotine Concentrations (ng/mL) - PK Population**

Timepoint	Statistic	Group-1 PK		Group-2 PK	
		THS 2.2 Menthol (N=43)	mCC (N=43)	THS 2.2 Menthol (N=18)	NRT gum (N=18)
T <sub>0</sub> + 2 h	n	43	43	18	18
	BLOQ - n (%)	0	0	0	0
	Geometric Mean (CV%)	3.05 (85.28%)	3.22 (65.49%)	2.12 (64.76%)	4.36 (51.16%)
	95% CI	2.42, 3.83	2.67, 3.87	1.57, 2.85	3.43, 5.55
	Median	3.04	3.51	2.12	4.35
	Q25, Q75	2.31, 5.20	2.36, 4.75	1.41, 3.16	3.41, 6.58
	Min, Max	0.2, 10.6	0.9, 9.9	0.7, 5.3	1.6, 8.9
	Mean (SD)	3.80 (2.403)	3.78 (2.130)	2.48 (1.403)	4.85 (2.281)

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: BLOQ values before T<sub>0</sub> are treated as zero. BLOQ values after the last quantifiable value are treated as missing. Any other BLOQ values are imputed by LLOQ/2.

Note: LLOQ = 0.2 ng/mL.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_pkconc.sas

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**Table 15.2.4.6 Descriptive Statistics of the Plasma Nicotine Concentrations (ng/mL) - PK Population**

Timepoint	Statistic	Group-1 PK		Group-2 PK	
		THS 2.2 Menthol (N=43)	mCC (N=43)	THS 2.2 Menthol (N=18)	NRT gum (N=18)
T <sub>0</sub> + 3 h	n	0	0	0	18
	BLOQ - n (%)				0
	Geometric Mean (CV%)				3.11 (60.74%)
	95% CI				2.35, 4.11
	Median				2.94
	Q25, Q75				2.17, 5.39
	Min, Max				1.1, 7.3
	Mean (SD)				3.59 (1.976)

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: BLOQ values before T<sub>0</sub> are treated as zero. BLOQ values after the last quantifiable value are treated as missing. Any other BLOQ values are imputed by LLOQ/2.

Note: LLOQ = 0.2 ng/mL.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_pkconc.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.6 Descriptive Statistics of the Plasma Nicotine Concentrations (ng/mL) - PK Population**

Timepoint	Statistic	Group-1 PK		Group-2 PK	
		THS 2.2 Menthol (N=43)	mCC (N=43)	THS 2.2 Menthol (N=18)	NRT gum (N=18)
T <sub>0</sub> + 4 h	n	43	43	18	18
	BLOQ - n (%)	1 (2.3%)	0	0	0
	Geometric Mean (CV%)	1.65 (103.89%)	1.69 (82.57%)	1.20 (66.40%)	2.33 (71.30%)
	95% CI	1.26, 2.15	1.35, 2.12	0.88, 1.62	1.69, 3.21
	Median	1.61	1.82	1.08	2.14
	Q25, Q75	1.03, 3.57	0.91, 3.09	0.79, 1.80	1.55, 4.56
	Min, Max	0.1, 6.7	0.4, 6.6	0.4, 3.3	0.9, 6.4
	Mean (SD)	2.22 (1.658)	2.16 (1.526)	1.43 (0.913)	2.83 (1.837)

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: BLOQ values before T<sub>0</sub> are treated as zero. BLOQ values after the last quantifiable value are treated as missing. Any other BLOQ values are imputed by LLOQ/2.

Note: LLOQ = 0.2 ng/mL.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_pkconc.sas

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Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

**Table 15.2.4.6 Descriptive Statistics of the Plasma Nicotine Concentrations (ng/mL) - PK Population**

Timepoint	Statistic	Group-1 PK		Group-2 PK	
		THS 2.2 Menthol (N=43)	mCC (N=43)	THS 2.2 Menthol (N=18)	NRT gum (N=18)
T <sub>0</sub> + 6 h	n	43	43	17	18
	BLOQ - n (%)	0	0	1 (5.6%)	0
	Geometric Mean (CV%)	1.07 (99.41%)	1.08 (100.00%)	0.77 (70.09%)	1.33 (94.11%)
	95% CI	0.83, 1.39	0.83, 1.39	0.55, 1.06	0.89, 1.98
	Median	0.98	0.91	0.72	1.28
	Q25, Q75	0.58, 2.55	0.53, 2.00	0.46, 1.01	0.79, 2.86
	Min, Max	0.2, 4.9	0.2, 5.7	0.3, 2.6	0.4, 4.4
	Mean (SD)	1.49 (1.240)	1.50 (1.296)	0.94 (0.670)	1.77 (1.355)

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: BLOQ values before T<sub>0</sub> are treated as zero. BLOQ values after the last quantifiable value are treated as missing. Any other BLOQ values are imputed by LLOQ/2.

Note: LLOQ = 0.2 ng/mL.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_pkconc.sas

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Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

**Table 15.2.4.6 Descriptive Statistics of the Plasma Nicotine Concentrations (ng/mL) - PK Population**

Timepoint	Statistic	Group-1 PK		Group-2 PK	
		THS 2.2 Menthol (N=43)	mCC (N=43)	THS 2.2 Menthol (N=18)	NRT gum (N=18)
T <sub>0</sub> + 9 h	n	39	40	15	16
	BLOQ - n (%)	5 (11.6%)	3 (7.0%)	3 (16.7%)	2 (11.1%)
	Geometric Mean (CV%)	0.69 (111.52%)	0.70 (101.43%)	0.52 (67.51%)	0.86 (101.12%)
	95% CI	0.51, 0.93	0.53, 0.92	0.36, 0.73	0.54, 1.35
	Median	0.60	0.64	0.49	0.85
	Q25, Q75	0.33, 1.76	0.34, 1.13	0.33, 0.80	0.41, 2.03
	Min, Max	0.1, 3.9	0.2, 3.8	0.2, 1.9	0.3, 2.9
	Mean (SD)	1.02 (0.936)	1.00 (0.916)	0.62 (0.442)	1.19 (0.987)

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: BLOQ values before T<sub>0</sub> are treated as zero. BLOQ values after the last quantifiable value are treated as missing. Any other BLOQ values are imputed by LLOQ/2.

Note: LLOQ = 0.2 ng/mL.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_pkconc.sas

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Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

**Table 15.2.4.6 Descriptive Statistics of the Plasma Nicotine Concentrations (ng/mL) - PK Population**

Timepoint	Statistic	Group-1 PK		Group-2 PK	
		THS 2.2 Menthol (N=43)	mCC (N=43)	THS 2.2 Menthol (N=18)	NRT gum (N=18)
T <sub>0</sub> + 12 h	n	33	28	12	15
	BLOQ - n (%)	10 (23.3%)	15 (34.9%)	6 (33.3%)	3 (16.7%)
	Geometric Mean (CV%)	0.63 (136.32%)	0.69 (80.60%)	0.39 (57.10%)	0.57 (100.97%)
	95% CI	0.43, 0.91	0.52, 0.91	0.27, 0.55	0.36, 0.92
	Median	0.46	0.57	0.31	0.49
	Q25, Q75	0.26, 1.29	0.37, 1.32	0.27, 0.54	0.23, 1.51
	Min, Max	0.2, 10.6	0.3, 3.0	0.2, 1.3	0.2, 1.9
	Mean (SD)	1.20 (2.001)	0.89 (0.694)	0.45 (0.309)	0.79 (0.644)

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: BLOQ values before T<sub>0</sub> are treated as zero. BLOQ values after the last quantifiable value are treated as missing. Any other BLOQ values are imputed by LLOQ/2.

Note: LLOQ = 0.2 ng/mL.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_pkconc.sas

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Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

**Table 15.2.4.6 Descriptive Statistics of the Plasma Nicotine Concentrations (ng/mL) - PK Population**

Timepoint	Statistic	Group-1 PK		Group-2 PK	
		THS 2.2 Menthol (N=43)	mCC (N=43)	THS 2.2 Menthol (N=18)	NRT gum (N=18)
T <sub>0</sub> + 24 h	n	21	22	6	8
	BLOQ - n (%)	22 (51.2%)	21 (48.8%)	12 (66.7%)	10 (55.6%)
	Geometric Mean (CV%)	0.43 (69.61%)	0.38 (59.07%)	0.35 (30.82%)	0.43 (45.32%)
	95% CI	0.32, 0.58	0.29, 0.49	0.25, 0.49	0.30, 0.62
	Median	0.37	0.32	0.33	0.47
	Q25, Q75	0.30, 0.54	0.22, 0.61	0.29, 0.50	0.31, 0.62
	Min, Max	0.2, 2.9	0.2, 1.3	0.2, 0.5	0.2, 0.7
	Mean (SD)	0.56 (0.587)	0.44 (0.283)	0.37 (0.112)	0.46 (0.178)

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: BLOQ values before T<sub>0</sub> are treated as zero. BLOQ values after the last quantifiable value are treated as missing. Any other BLOQ values are imputed by LLOQ/2.

Note: LLOQ = 0.2 ng/mL.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_pkconc.sas

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Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

**Table 15.2.4.6.1 Descriptive Statistics of the Plasma Nicotine Concentrations (ng/mL) by Sex - PK Population**

Timepoint	Statistic	Group-1 PK			
		THS 2.2 Menthol (N=43)		mCC (N=43)	
		Males (N=23)	Females (N=20)	Males (N=23)	Females (N=20)
15 min < T <sub>0</sub>	n	23	20	23	20
	BLOQ - n (%)	10 (43.5%)	13 (65.0%)	9 (39.1%)	13 (65.0%)
	Geometric Mean (CV%)	NC	NC	NC	NC
	95% CI	NC	NC	NC	NC
	Median	0.22	0.00	0.21	0.00
	Q25, Q75	0.00, 0.47	0.00, 0.27	0.00, 0.34	0.00, 0.44
	Min, Max	0.0, 2.2	0.0, 0.3	0.0, 2.0	0.0, 1.4
	Mean (SD)	0.34 (0.504)	0.10 (0.146)	0.28 (0.424)	0.26 (0.440)

Note: mCC = menthol conventional cigarettes; THS = Tobacco Heating System.

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

Note: NC = Not calculated.

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: BLOQ values before T<sub>0</sub> are treated as zero. BLOQ values after the last quantifiable value are treated as missing. Any other BLOQ values are imputed by LLOQ/2.

Note: LLOQ = 0.2 ng/mL.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_pkconc2.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.6.1 Descriptive Statistics of the Plasma Nicotine Concentrations (ng/mL) by Sex - PK Population**

Timepoint	Statistic	Group-1 PK			
		THS 2.2 Menthol (N=43)		mCC (N=43)	
		Males (N=23)	Females (N=20)	Males (N=23)	Females (N=20)
T <sub>0</sub> + 2 min	n	23	20	23	20
	BLOQ - n (%)	0	0	0	1 (5.0%)
	Geometric Mean (CV%)	1.97 (200.12%)	2.91 (171.37%)	2.24 (110.07%)	5.22 (255.15%)
	95% CI	1.13, 3.41	1.68, 5.05	1.52, 3.30	2.68, 10.15
	Median	1.87	3.13	2.32	7.39
	Q25, Q75	1.22, 3.64	1.21, 7.00	1.32, 3.59	2.39, 13.95
	Min, Max	0.2, 19.2	0.3, 35.1	0.4, 20.9	0.1, 36.6
	Mean (SD)	3.88 (4.729)	5.58 (7.825)	3.39 (4.197)	9.97 (10.006)

Note: mCC = menthol conventional cigarettes; THS = Tobacco Heating System.

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

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: BLOQ values before T<sub>0</sub> are treated as zero. BLOQ values after the last quantifiable value are treated as missing. Any other BLOQ values are imputed by LLOQ/2.

Note: LLOQ = 0.2 ng/mL.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_pkconc2.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.6.1 Descriptive Statistics of the Plasma Nicotine Concentrations (ng/mL) by Sex - PK Population**

Timepoint	Statistic	Group-1 PK			
		THS 2.2 Menthol (N=43)		mCC (N=43)	
		Males (N=23)	Females (N=20)	Males (N=23)	Females (N=20)
T <sub>0</sub> + 4 min	n	23	20	23	20
	BLOQ - n (%)	0	0	0	0
	Geometric Mean (CV%)	5.81 (194.08%)	7.43 (113.66%)	6.33 (110.18%)	10.78 (159.55%)
	95% CI	3.38, 9.98	4.85, 11.39	4.30, 9.32	6.36, 18.26
	Median	8.86	6.34	7.33	14.55
	Q25, Q75	2.21, 14.80	4.09, 15.00	4.34, 11.30	4.84, 25.95
	Min, Max	0.3, 31.3	1.4, 38.1	0.4, 30.5	0.9, 47.6
	Mean (SD)	9.63 (7.848)	10.94 (10.222)	8.61 (6.852)	16.72 (13.038)

Note: mCC = menthol conventional cigarettes; THS = Tobacco Heating System.

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

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: BLOQ values before T<sub>0</sub> are treated as zero. BLOQ values after the last quantifiable value are treated as missing. Any other BLOQ values are imputed by LLOQ/2.

Note: LLOQ = 0.2 ng/mL.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_pkconc2.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.6.1 Descriptive Statistics of the Plasma Nicotine Concentrations (ng/mL) by Sex - PK Population**

Timepoint	Statistic	Group-1 PK			
		THS 2.2 Menthol (N=43)		mCC (N=43)	
		Males (N=23)	Females (N=20)	Males (N=23)	Females (N=20)
T <sub>0</sub> + 6 min	n	23	20	23	20
	BLOQ - n (%)	0	0	0	0
	Geometric Mean (CV%)	8.34 (163.00%)	8.23 (98.62%)	8.82 (84.91%)	9.68 (117.84%)
	95% CI	5.09, 13.65	5.59, 12.11	6.41, 12.13	6.25, 14.99
	Median	12.60	7.45	8.06	11.00
	Q25, Q75	3.99, 18.60	5.31, 12.79	6.13, 15.60	5.27, 18.20
	Min, Max	0.4, 26.2	1.3, 43.8	1.2, 31.5	1.4, 58.1
	Mean (SD)	12.27 (7.890)	11.54 (11.096)	11.12 (7.781)	14.11 (13.181)

Note: mCC = menthol conventional cigarettes; THS = Tobacco Heating System.

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

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: BLOQ values before T<sub>0</sub> are treated as zero. BLOQ values after the last quantifiable value are treated as missing. Any other BLOQ values are imputed by LLOQ/2.

Note: LLOQ = 0.2 ng/mL.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_pkconc2.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.6.1 Descriptive Statistics of the Plasma Nicotine Concentrations (ng/mL) by Sex - PK Population**

Timepoint	Statistic	Group-1 PK			
		THS 2.2 Menthol (N=43)		mCC (N=43)	
		Males (N=23)	Females (N=20)	Males (N=23)	Females (N=20)
T <sub>0</sub> + 8 min	n	23	20	23	20
	BLOQ - n (%)	0	0	0	0
	Geometric Mean (CV%)	8.04 (139.76%)	7.66 (87.08%)	8.65 (56.78%)	8.49 (99.39%)
	95% CI	5.12, 12.61	5.39, 10.89	6.88, 10.88	5.75, 12.51
	Median	11.10	6.97	8.49	8.03
	Q25, Q75	5.05, 15.90	4.76, 15.40	6.36, 13.20	5.30, 16.30
	Min, Max	0.4, 29.7	1.4, 27.5	1.9, 19.8	1.4, 36.0
	Mean (SD)	11.14 (7.029)	9.95 (7.588)	9.70 (4.385)	11.36 (8.696)

Note: mCC = menthol conventional cigarettes; THS = Tobacco Heating System.

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

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: BLOQ values before T<sub>0</sub> are treated as zero. BLOQ values after the last quantifiable value are treated as missing. Any other BLOQ values are imputed by LLOQ/2.

Note: LLOQ = 0.2 ng/mL.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_pkconc2.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.6.1 Descriptive Statistics of the Plasma Nicotine Concentrations (ng/mL) by Sex - PK Population**

Timepoint	Statistic	Group-1 PK			
		THS 2.2 Menthol (N=43)		mCC (N=43)	
		Males (N=23)	Females (N=20)	Males (N=23)	Females (N=20)
T <sub>0</sub> + 10 min	n	23	20	23	20
	BLOQ - n (%)	0	0	0	0
	Geometric Mean (CV%)	7.47 (141.77%)	7.11 (80.66%)	8.49 (52.12%)	8.40 (86.71%)
	95% CI	4.74, 11.76	5.10, 9.90	6.87, 10.50	5.91, 11.93
	Median	9.11	7.06	8.22	9.17
	Q25, Q75	5.79, 14.40	4.21, 11.70	6.55, 12.30	5.01, 13.50
	Min, Max	0.3, 26.3	1.3, 24.3	2.6, 19.2	2.3, 31.5
	Mean (SD)	10.25 (6.195)	8.93 (6.388)	9.45 (4.294)	10.77 (7.785)

Note: mCC = menthol conventional cigarettes; THS = Tobacco Heating System.

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

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: BLOQ values before T<sub>0</sub> are treated as zero. BLOQ values after the last quantifiable value are treated as missing. Any other BLOQ values are imputed by LLOQ/2.

Note: LLOQ = 0.2 ng/mL.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_pkconc2.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.6.1 Descriptive Statistics of the Plasma Nicotine Concentrations (ng/mL) by Sex - PK Population**

Timepoint	Statistic	Group-1 PK			
		THS 2.2 Menthol (N=43)		mCC (N=43)	
		Males (N=23)	Females (N=20)	Males (N=23)	Females (N=20)
T <sub>0</sub> + 15 min	n	23	20	23	20
	BLOQ - n (%)	0	0	0	0
	Geometric Mean (CV%)	5.82 (106.98%)	6.50 (80.00%)	6.65 (45.30%)	6.31 (70.60%)
	95% CI	3.98, 8.49	4.67, 9.03	5.52, 8.03	4.68, 8.51
	Median	6.74	6.94	6.48	7.04
	Q25, Q75	4.89, 9.75	4.25, 11.00	5.28, 8.82	4.72, 8.54
	Min, Max	0.3, 16.7	1.2, 20.4	2.6, 12.8	1.3, 20.0
	Mean (SD)	7.33 (3.916)	8.03 (5.165)	7.22 (2.840)	7.46 (4.251)

Note: mCC = menthol conventional cigarettes; THS = Tobacco Heating System.

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

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: BLOQ values before T<sub>0</sub> are treated as zero. BLOQ values after the last quantifiable value are treated as missing. Any other BLOQ values are imputed by LLOQ/2.

Note: LLOQ = 0.2 ng/mL.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_pkconc2.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.6.1 Descriptive Statistics of the Plasma Nicotine Concentrations (ng/mL) by Sex - PK Population**

Timepoint	Statistic	Group-1 PK			
		THS 2.2 Menthol (N=43)		mCC (N=43)	
		Males (N=23)	Females (N=20)	Males (N=23)	Females (N=20)
T <sub>0</sub> + 30 min	n	23	20	23	20
	BLOQ - n (%)	0	0	0	0
	Geometric Mean (CV%)	4.91 (96.18%)	5.27 (74.42%)	5.88 (49.61%)	5.02 (62.14%)
	95% CI	3.46, 6.98	3.86, 7.20	4.80, 7.21	3.84, 6.57
	Median	6.08	6.09	6.38	5.15
	Q25, Q75	4.19, 7.31	3.42, 7.70	4.81, 7.87	3.64, 7.94
	Min, Max	0.3, 12.1	1.3, 13.3	2.0, 12.1	1.8, 14.7
	Mean (SD)	6.02 (3.002)	6.33 (3.601)	6.45 (2.606)	5.82 (3.264)

Note: mCC = menthol conventional cigarettes; THS = Tobacco Heating System.

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

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: BLOQ values before T<sub>0</sub> are treated as zero. BLOQ values after the last quantifiable value are treated as missing. Any other BLOQ values are imputed by LLOQ/2.

Note: LLOQ = 0.2 ng/mL.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_pkconc2.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.6.1 Descriptive Statistics of the Plasma Nicotine Concentrations (ng/mL) by Sex - PK Population**

Timepoint	Statistic	Group-1 PK			
		THS 2.2 Menthol (N=43)		mCC (N=43)	
		Males (N=23)	Females (N=20)	Males (N=23)	Females (N=20)
T <sub>0</sub> + 45 min	n	23	20	23	20
	BLOQ - n (%)	1 (4.3%)	0	0	0
	Geometric Mean (CV%)	4.32 (122.61%)	4.76 (73.78%)	5.26 (42.24%)	4.75 (68.90%)
	95% CI	2.85, 6.54	3.49, 6.48	4.41, 6.27	3.54, 6.36
	Median	5.58	5.13	5.46	5.72
	Q25, Q75	3.78, 7.00	3.12, 7.59	4.52, 7.05	2.92, 6.71
	Min, Max	0.1, 9.3	1.3, 13.6	1.8, 11.5	1.5, 15.2
	Mean (SD)	5.41 (2.431)	5.73 (3.393)	5.66 (2.143)	5.64 (3.385)

Note: mCC = menthol conventional cigarettes; THS = Tobacco Heating System.

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

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: BLOQ values before T<sub>0</sub> are treated as zero. BLOQ values after the last quantifiable value are treated as missing. Any other BLOQ values are imputed by LLOQ/2.

Note: LLOQ = 0.2 ng/mL.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_pkconc2.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.6.1 Descriptive Statistics of the Plasma Nicotine Concentrations (ng/mL) by Sex - PK Population**

Timepoint	Statistic	Group-1 PK			
		THS 2.2 Menthol (N=43)		mCC (N=43)	
		Males (N=23)	Females (N=20)	Males (N=23)	Females (N=20)
T <sub>0</sub> + 60 min	n	23	20	23	20
	BLOQ - n (%)	0	0	0	0
	Geometric Mean (CV%)	4.39 (97.09%)	4.25 (74.32%)	4.97 (49.00%)	4.28 (67.24%)
	95% CI	3.08, 6.25	3.11, 5.80	4.06, 6.08	3.21, 5.70
	Median	5.46	4.53	5.18	4.99
	Q25, Q75	3.84, 7.00	3.12, 6.20	3.99, 7.00	2.56, 6.50
	Min, Max	0.2, 10.7	1.2, 12.3	1.6, 12.1	1.5, 13.4
	Mean (SD)	5.32 (2.433)	5.15 (3.184)	5.47 (2.375)	5.05 (2.921)

Note: mCC = menthol conventional cigarettes; THS = Tobacco Heating System.

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

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: BLOQ values before T<sub>0</sub> are treated as zero. BLOQ values after the last quantifiable value are treated as missing. Any other BLOQ values are imputed by LLOQ/2.

Note: LLOQ = 0.2 ng/mL.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_pkconc2.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.6.1 Descriptive Statistics of the Plasma Nicotine Concentrations (ng/mL) by Sex - PK Population**

Timepoint	Statistic	Group-1 PK			
		THS 2.2 Menthol (N=43)		mCC (N=43)	
		Males (N=23)	Females (N=20)	Males (N=23)	Females (N=20)
T <sub>0</sub> + 2 h	n	23	20	23	20
	BLOQ - n (%)	0	0	0	0
	Geometric Mean (CV%)	3.15 (90.55%)	2.94 (81.78%)	3.56 (54.16%)	2.87 (76.94%)
	95% CI	2.25, 4.40	2.10, 4.11	2.85, 4.44	2.08, 3.95
	Median	3.34	2.57	3.63	3.31
	Q25, Q75	2.47, 5.52	2.09, 4.49	2.68, 4.88	1.57, 4.11
	Min, Max	0.2, 8.8	0.9, 10.6	1.3, 9.9	0.9, 9.4
	Mean (SD)	3.82 (1.961)	3.77 (2.882)	4.00 (1.999)	3.53 (2.297)

Note: mCC = menthol conventional cigarettes; THS = Tobacco Heating System.

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

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: BLOQ values before T<sub>0</sub> are treated as zero. BLOQ values after the last quantifiable value are treated as missing. Any other BLOQ values are imputed by LLOQ/2.

Note: LLOQ = 0.2 ng/mL.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_pkconc2.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.6.1 Descriptive Statistics of the Plasma Nicotine Concentrations (ng/mL) by Sex - PK Population**

Timepoint	Statistic	Group-1 PK			
		THS 2.2 Menthol (N=43)		mCC (N=43)	
		Males (N=23)	Females (N=20)	Males (N=23)	Females (N=20)
T <sub>0</sub> + 4 h	n	23	20	23	20
	BLOQ - n (%)	1 (4.3%)	0	0	0
	Geometric Mean (CV%)	1.86 (108.30%)	1.43 (98.87%)	2.05 (64.05%)	1.36 (96.29%)
	95% CI	1.26, 2.72	0.97, 2.12	1.59, 2.65	0.93, 1.99
	Median	1.94	1.32	2.07	1.23
	Q25, Q75	1.34, 3.57	0.81, 2.79	1.30, 3.48	0.75, 2.28
	Min, Max	0.1, 6.2	0.4, 6.7	0.8, 6.6	0.4, 5.6
	Mean (SD)	2.41 (1.504)	2.01 (1.837)	2.41 (1.447)	1.87 (1.599)

Note: mCC = menthol conventional cigarettes; THS = Tobacco Heating System.

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

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: BLOQ values before T<sub>0</sub> are treated as zero. BLOQ values after the last quantifiable value are treated as missing. Any other BLOQ values are imputed by LLOQ/2.

Note: LLOQ = 0.2 ng/mL.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_pkconc2.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.6.1 Descriptive Statistics of the Plasma Nicotine Concentrations (ng/mL) by Sex - PK Population**

Timepoint	Statistic	Group-1 PK			
		THS 2.2 Menthol (N=43)		mCC (N=43)	
		Males (N=23)	Females (N=20)	Males (N=23)	Females (N=20)
T <sub>0</sub> + 6 h	n	23	20	23	20
	BLOQ - n (%)	0	0	0	0
	Geometric Mean (CV%)	1.29 (88.28%)	0.87 (106.93%)	1.34 (76.14%)	0.83 (118.27%)
	95% CI	0.92, 1.80	0.57, 1.31	1.00, 1.80	0.53, 1.30
	Median	1.13	0.74	1.34	0.63
	Q25, Q75	0.73, 2.62	0.45, 1.69	0.74, 2.08	0.45, 1.60
	Min, Max	0.3, 4.4	0.2, 4.9	0.5, 5.7	0.2, 4.5
	Mean (SD)	1.66 (1.173)	1.29 (1.315)	1.68 (1.256)	1.30 (1.342)

Note: mCC = menthol conventional cigarettes; THS = Tobacco Heating System.

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

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: BLOQ values before T<sub>0</sub> are treated as zero. BLOQ values after the last quantifiable value are treated as missing. Any other BLOQ values are imputed by LLOQ/2.

Note: LLOQ = 0.2 ng/mL.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_pkconc2.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.6.1 Descriptive Statistics of the Plasma Nicotine Concentrations (ng/mL) by Sex - PK Population**

Timepoint	Statistic	Group-1 PK			
		THS 2.2 Menthol (N=43)		mCC (N=43)	
		Males (N=23)	Females (N=20)	Males (N=23)	Females (N=20)
T <sub>0</sub> + 9 h	n	22	17	23	17
	BLOQ - n (%)	1 (4.3%)	4 (20.0%)	0	3 (15.0%)
	Geometric Mean (CV%)	0.81 (94.45%)	0.56 (131.45%)	0.81 (83.38%)	0.57 (123.00%)
	95% CI	0.56, 1.16	0.33, 0.95	0.59, 1.12	0.34, 0.94
	Median	0.72	0.41	0.87	0.37
	Q25, Q75	0.41, 1.88	0.30, 1.36	0.45, 1.15	0.24, 0.85
	Min, Max	0.2, 3.0	0.1, 3.9	0.3, 3.8	0.2, 2.9
	Mean (SD)	1.09 (0.859)	0.93 (1.046)	1.06 (0.884)	0.91 (0.978)

Note: mCC = menthol conventional cigarettes; THS = Tobacco Heating System.

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

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: BLOQ values before T<sub>0</sub> are treated as zero. BLOQ values after the last quantifiable value are treated as missing. Any other BLOQ values are imputed by LLOQ/2.

Note: LLOQ = 0.2 ng/mL.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_pkconc2.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.6.1 Descriptive Statistics of the Plasma Nicotine Concentrations (ng/mL) by Sex - PK Population**

Timepoint	Statistic	Group-1 PK			
		THS 2.2 Menthol (N=43)		mCC (N=43)	
		Males (N=23)	Females (N=20)	Males (N=23)	Females (N=20)
T <sub>0</sub> + 12 h	n	19	14	20	8
	BLOQ - n (%)	4 (17.4%)	6 (30.0%)	3 (13.0%)	12 (60.0%)
	Geometric Mean (CV%)	0.70 (112.90%)	0.55 (176.09%)	0.61 (78.34%)	0.91 (81.01%)
	95% CI	0.45, 1.08	0.27, 1.09	0.44, 0.85	0.50, 1.66
	Median	0.63	0.32	0.53	0.99
	Q25, Q75	0.30, 1.34	0.22, 1.25	0.32, 0.99	0.49, 1.77
	Min, Max	0.2, 5.8	0.2, 10.6	0.3, 3.0	0.4, 2.1
	Mean (SD)	1.07 (1.276)	1.36 (2.747)	0.79 (0.683)	1.12 (0.709)

Note: mCC = menthol conventional cigarettes; THS = Tobacco Heating System.

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

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: BLOQ values before T<sub>0</sub> are treated as zero. BLOQ values after the last quantifiable value are treated as missing. Any other BLOQ values are imputed by LLOQ/2.

Note: LLOQ = 0.2 ng/mL.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_pkconc2.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.6.1 Descriptive Statistics of the Plasma Nicotine Concentrations (ng/mL) by Sex - PK Population**

Timepoint	Statistic	Group-1 PK			
		THS 2.2 Menthol (N=43)		mCC (N=43)	
		Males (N=23)	Females (N=20)	Males (N=23)	Females (N=20)
T <sub>0</sub> + 24 h	n	13	8	15	7
	BLOQ - n (%)	10 (43.5%)	12 (60.0%)	8 (34.8%)	13 (65.0%)
	Geometric Mean (CV%)	0.48 (77.44%)	0.37 (56.00%)	0.35 (55.94%)	0.45 (66.70%)
	95% CI	0.31, 0.73	0.23, 0.57	0.26, 0.48	0.25, 0.79
	Median	0.45	0.33	0.30	0.59
	Q25, Q75	0.30, 0.55	0.24, 0.48	0.22, 0.50	0.21, 0.76
	Min, Max	0.2, 2.9	0.2, 1.1	0.2, 1.3	0.2, 0.9
	Mean (SD)	0.64 (0.714)	0.42 (0.276)	0.41 (0.288)	0.52 (0.279)

Note: mCC = menthol conventional cigarettes; THS = Tobacco Heating System.

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

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: BLOQ values before T<sub>0</sub> are treated as zero. BLOQ values after the last quantifiable value are treated as missing. Any other BLOQ values are imputed by LLOQ/2.

Note: LLOQ = 0.2 ng/mL.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_pkconc2.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.6.1 Descriptive Statistics of the Plasma Nicotine Concentrations (ng/mL) by Sex - PK Population**

Timepoint	Statistic	Group-2 PK			
		THS 2.2 Menthol (N=18)		NRT gum (N=18)	
		Males (N=9)	Females (N=9)	Males (N=9)	Females (N=9)
15 min < T <sub>0</sub>	n	9	9	9	9
	BLOQ - n (%)	5 (55.6%)	7 (77.8%)	2 (22.2%)	5 (55.6%)
	Geometric Mean (CV%)	NC	NC	NC	NC
	95% CI	NC	NC	NC	NC
	Median	0.00	0.00	0.22	0.00
	Q25, Q75	0.00, 0.33	0.00, 0.00	0.21, 0.33	0.00, 0.49
	Min, Max	0.0, 0.9	0.0, 0.3	0.0, 0.5	0.0, 0.9
	Mean (SD)	0.25 (0.361)	0.06 (0.114)	0.24 (0.174)	0.25 (0.342)

Note: NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Note: NC = Not calculated.

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: BLOQ values before T<sub>0</sub> are treated as zero. BLOQ values after the last quantifiable value are treated as missing. Any other BLOQ values are imputed by LLOQ/2.

Note: LLOQ = 0.2 ng/mL.

Appendix 15.3.3.2

Path: /cvm/projects/prj/development/000000106326/dev/tables/t\_pkconc2.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.6.1 Descriptive Statistics of the Plasma Nicotine Concentrations (ng/mL) by Sex - PK Population**

Timepoint	Statistic	Group-2 PK			
		THS 2.2 Menthol (N=18)		NRT gum (N=18)	
		Males (N=9)	Females (N=9)	Males (N=9)	Females (N=9)
T <sub>0</sub> + 2 min	n	9	9	0	0
	BLOQ - n (%)	0	0		
	Geometric Mean (CV%)	2.68 (221.59%)	2.64 (93.10%)		
	95% CI	0.96, 7.46	1.43, 4.85		
	Median	1.97	3.18		
	Q25, Q75	1.48, 4.17	1.47, 4.81		
	Min, Max	0.3, 30.2	0.8, 7.2		
	Mean (SD)	5.94 (9.416)	3.39 (2.280)		

Note: NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: BLOQ values before T<sub>0</sub> are treated as zero. BLOQ values after the last quantifiable value are treated as missing. Any other BLOQ values are imputed by LLOQ/2.

Note: LLOQ = 0.2 ng/mL.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_pkconc2.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.6.1 Descriptive Statistics of the Plasma Nicotine Concentrations (ng/mL) by Sex - PK Population**

Timepoint	Statistic	Group-2 PK			
		THS 2.2 Menthol (N=18)		NRT gum (N=18)	
		Males (N=9)	Females (N=9)	Males (N=9)	Females (N=9)
T <sub>0</sub> + 4 min	n	9	9	0	0
	BLOQ - n (%)	0	0		
	Geometric Mean (CV%)	5.64 (197.17%)	5.54 (99.54%)		
	95% CI	2.14, 14.86	2.92, 10.49		
	Median	5.37	5.73		
	Q25, Q75	3.45, 8.17	3.51, 11.50		
	Min, Max	0.6, 45.3	1.1, 14.0		
	Mean (SD)	11.02 (14.526)	7.13 (4.645)		

Note: NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: BLOQ values before T<sub>0</sub> are treated as zero. BLOQ values after the last quantifiable value are treated as missing. Any other BLOQ values are imputed by LLOQ/2.

Note: LLOQ = 0.2 ng/mL.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_pkconc2.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.6.1 Descriptive Statistics of the Plasma Nicotine Concentrations (ng/mL) by Sex - PK Population**

Timepoint	Statistic	Group-2 PK			
		THS 2.2 Menthol (N=18)		NRT gum (N=18)	
		Males (N=9)	Females (N=9)	Males (N=9)	Females (N=9)
T <sub>0</sub> + 6 min	n	9	9	0	0
	BLOQ - n (%)	0	0		
	Geometric Mean (CV%)	7.32 (135.88%)	5.76 (96.06%)		
	95% CI	3.33, 16.08	3.09, 10.72		
	Median	7.13	7.18		
	Q25, Q75	4.04, 11.00	3.65, 8.90		
	Min, Max	2.0, 45.8	1.2, 20.3		
	Mean (SD)	11.94 (14.018)	7.44 (5.594)		

Note: NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: BLOQ values before T<sub>0</sub> are treated as zero. BLOQ values after the last quantifiable value are treated as missing. Any other BLOQ values are imputed by LLOQ/2.

Note: LLOQ = 0.2 ng/mL.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_pkconc2.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.6.1 Descriptive Statistics of the Plasma Nicotine Concentrations (ng/mL) by Sex - PK Population**

Timepoint	Statistic	Group-2 PK			
		THS 2.2 Menthol (N=18)		NRT gum (N=18)	
		Males (N=9)	Females (N=9)	Males (N=9)	Females (N=9)
T <sub>0</sub> + 8 min	n	9	9	0	0
	BLOQ - n (%)	0	0		
	Geometric Mean (CV%)	6.59 (99.39%)	4.81 (87.24%)		
	95% CI	3.48, 12.46	2.69, 8.59		
	Median	5.77	5.15		
	Q25, Q75	4.78, 9.06	3.00, 7.13		
	Min, Max	2.1, 28.9	1.2, 15.8		
	Mean (SD)	9.05 (8.452)	6.07 (4.358)		

Note: NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: BLOQ values before T<sub>0</sub> are treated as zero. BLOQ values after the last quantifiable value are treated as missing. Any other BLOQ values are imputed by LLOQ/2.

Note: LLOQ = 0.2 ng/mL.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_pkconc2.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.6.1 Descriptive Statistics of the Plasma Nicotine Concentrations (ng/mL) by Sex - PK Population**

Timepoint	Statistic	Group-2 PK			
		THS 2.2 Menthol (N=18)		NRT gum (N=18)	
		Males (N=9)	Females (N=9)	Males (N=9)	Females (N=9)
T <sub>0</sub> + 10 min	n	9	9	9	9
	BLOQ - n (%)	0	0	0	0
	Geometric Mean (CV%)	6.73 (83.53%)	4.78 (88.76%)	1.24 (56.64%)	1.62 (57.46%)
	95% CI	3.84, 11.78	2.66, 8.60	0.82, 1.87	1.07, 2.45
	Median	6.97	6.24	1.15	1.70
	Q25, Q75	4.94, 9.97	3.56, 7.22	0.69, 1.87	1.34, 1.94
	Min, Max	2.1, 22.1	1.1, 13.8	0.7, 2.5	0.7, 4.2
	Mean (SD)	8.42 (6.104)	5.95 (3.788)	1.40 (0.704)	1.84 (1.043)

Note: NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: BLOQ values before T<sub>0</sub> are treated as zero. BLOQ values after the last quantifiable value are treated as missing. Any other BLOQ values are imputed by LLOQ/2.

Note: LLOQ = 0.2 ng/mL.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_pkconc2.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.6.1 Descriptive Statistics of the Plasma Nicotine Concentrations (ng/mL) by Sex - PK Population**

Timepoint	Statistic	Group-2 PK			
		THS 2.2 Menthol (N=18)		NRT gum (N=18)	
		Males (N=9)	Females (N=9)	Males (N=9)	Females (N=9)
T <sub>0</sub> + 15 min	n	9	9	0	0
	BLOQ - n (%)	0	0		
	Geometric Mean (CV%)	5.46 (63.07%)	4.06 (75.47%)		
	95% CI	3.49, 8.52	2.42, 6.81		
	Median	5.77	3.71		
	Q25, Q75	4.29, 8.60	3.33, 7.37		
	Min, Max	2.4, 13.4	1.3, 10.9		
	Mean (SD)	6.30 (3.549)	4.90 (3.099)		

Note: NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: BLOQ values before T<sub>0</sub> are treated as zero. BLOQ values after the last quantifiable value are treated as missing. Any other BLOQ values are imputed by LLOQ/2.

Note: LLOQ = 0.2 ng/mL.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_pkconc2.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.6.1 Descriptive Statistics of the Plasma Nicotine Concentrations (ng/mL) by Sex - PK Population**

Timepoint	Statistic	Group-2 PK			
		THS 2.2 Menthol (N=18)		NRT gum (N=18)	
		Males (N=9)	Females (N=9)	Males (N=9)	Females (N=9)
T <sub>0</sub> + 20 min	n	0	0	9	9
	BLOQ - n (%)			0	0
	Geometric Mean (CV%)			3.11 (51.73%)	4.16 (48.04%)
	95% CI			2.14, 4.53	2.92, 5.90
	Median			2.96	5.17
	Q25, Q75			1.93, 3.95	2.84, 6.17
	Min, Max			1.8, 6.6	2.2, 7.0
	Mean (SD)			3.47 (1.738)	4.53 (1.882)

Note: NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: BLOQ values before T<sub>0</sub> are treated as zero. BLOQ values after the last quantifiable value are treated as missing. Any other BLOQ values are imputed by LLOQ/2.

Note: LLOQ = 0.2 ng/mL.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_pkconc2.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.6.1 Descriptive Statistics of the Plasma Nicotine Concentrations (ng/mL) by Sex - PK Population**

Timepoint	Statistic	Group-2 PK			
		THS 2.2 Menthol (N=18)		NRT gum (N=18)	
		Males (N=9)	Females (N=9)	Males (N=9)	Females (N=9)
T <sub>0</sub> + 25 min	n	0	0	9	9
	BLOQ - n (%)			0	0
	Geometric Mean (CV%)			3.73 (44.79%)	4.62 (41.24%)
	95% CI			2.68, 5.18	3.40, 6.28
	Median			3.63	4.85
	Q25, Q75			2.73, 3.84	3.67, 5.78
	Min, Max			2.4, 7.8	2.2, 8.6
	Mean (SD)			4.07 (1.995)	4.95 (1.901)

Note: NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: BLOQ values before T<sub>0</sub> are treated as zero. BLOQ values after the last quantifiable value are treated as missing. Any other BLOQ values are imputed by LLOQ/2.

Note: LLOQ = 0.2 ng/mL.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_pkconc2.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.6.1 Descriptive Statistics of the Plasma Nicotine Concentrations (ng/mL) by Sex - PK Population**

Timepoint	Statistic	Group-2 PK			
		THS 2.2 Menthol (N=18)		NRT gum (N=18)	
		Males (N=9)	Females (N=9)	Males (N=9)	Females (N=9)
T <sub>0</sub> + 30 min	n	9	9	9	9
	BLOQ - n (%)	0	0	0	0
	Geometric Mean (CV%)	4.44 (58.72%)	3.30 (66.27%)	4.54 (43.18%)	5.38 (25.95%)
	95% CI	2.92, 6.75	2.07, 5.25	3.30, 6.25	4.42, 6.56
	Median	5.37	3.01	4.27	5.81
	Q25, Q75	3.67, 6.33	2.49, 5.66	3.38, 5.02	4.63, 6.13
	Min, Max	1.9, 9.3	1.3, 7.5	2.5, 8.9	3.2, 7.6
	Mean (SD)	4.99 (2.343)	3.86 (2.289)	4.92 (2.212)	5.53 (1.294)

Note: NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: BLOQ values before T<sub>0</sub> are treated as zero. BLOQ values after the last quantifiable value are treated as missing. Any other BLOQ values are imputed by LLOQ/2.

Note: LLOQ = 0.2 ng/mL.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_pkconc2.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.6.1 Descriptive Statistics of the Plasma Nicotine Concentrations (ng/mL) by Sex - PK Population**

Timepoint	Statistic	Group-2 PK			
		THS 2.2 Menthol (N=18)		NRT gum (N=18)	
		Males (N=9)	Females (N=9)	Males (N=9)	Females (N=9)
T <sub>0</sub> + 35 min	n	0	0	9	9
	BLOQ - n (%)			0	0
	Geometric Mean (CV%)			4.49 (38.72%)	5.81 (37.64%)
	95% CI			3.37, 5.99	4.39, 7.69
	Median			4.34	6.22
	Q25, Q75			3.60, 5.21	5.19, 6.87
	Min, Max			2.3, 8.3	2.7, 9.8
	Mean (SD)			4.77 (1.766)	6.14 (2.020)

Note: NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: BLOQ values before T<sub>0</sub> are treated as zero. BLOQ values after the last quantifiable value are treated as missing. Any other BLOQ values are imputed by LLOQ/2.

Note: LLOQ = 0.2 ng/mL.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_pkconc2.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.6.1 Descriptive Statistics of the Plasma Nicotine Concentrations (ng/mL) by Sex - PK Population**

Timepoint	Statistic	Group-2 PK			
		THS 2.2 Menthol (N=18)		NRT gum (N=18)	
		Males (N=9)	Females (N=9)	Males (N=9)	Females (N=9)
T <sub>0</sub> + 40 min	n	0	0	9	9
	BLOQ - n (%)			0	0
	Geometric Mean (CV%)			5.37 (39.75%)	6.66 (28.18%)
	95% CI			4.00, 7.22	5.38, 8.24
	Median			5.24	7.19
	Q25, Q75			4.43, 6.79	6.02, 7.72
	Min, Max			2.7, 9.4	3.5, 8.7
	Mean (SD)			5.73 (2.150)	6.86 (1.572)

Note: NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: BLOQ values before T<sub>0</sub> are treated as zero. BLOQ values after the last quantifiable value are treated as missing. Any other BLOQ values are imputed by LLOQ/2.

Note: LLOQ = 0.2 ng/mL.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_pkconc2.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.6.1 Descriptive Statistics of the Plasma Nicotine Concentrations (ng/mL) by Sex - PK Population**

Timepoint	Statistic	Group-2 PK			
		THS 2.2 Menthol (N=18)		NRT gum (N=18)	
		Males (N=9)	Females (N=9)	Males (N=9)	Females (N=9)
T <sub>0</sub> + 45 min	n	9	9	9	9
	BLOQ - n (%)	0	0	0	0
	Geometric Mean (CV%)	4.34 (53.24%)	2.83 (67.14%)	5.23 (41.03%)	7.01 (29.08%)
	95% CI	2.95, 6.37	1.77, 4.53	3.86, 7.08	5.63, 8.73
	Median	5.25	2.57	5.20	6.89
	Q25, Q75	3.40, 5.86	2.14, 4.59	4.34, 6.01	6.50, 7.54
	Min, Max	1.9, 8.3	1.2, 7.4	2.5, 9.1	4.1, 11.1
	Mean (SD)	4.79 (2.043)	3.34 (2.074)	5.59 (2.145)	7.26 (2.018)

Note: NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: BLOQ values before T<sub>0</sub> are treated as zero. BLOQ values after the last quantifiable value are treated as missing. Any other BLOQ values are imputed by LLOQ/2.

Note: LLOQ = 0.2 ng/mL.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_pkconc2.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.6.1 Descriptive Statistics of the Plasma Nicotine Concentrations (ng/mL) by Sex - PK Population**

Timepoint	Statistic	Group-2 PK			
		THS 2.2 Menthol (N=18)		NRT gum (N=18)	
		Males (N=9)	Females (N=9)	Males (N=9)	Females (N=9)
T <sub>0</sub> + 60 min	n	9	9	9	9
	BLOQ - n (%)	0	0	0	0
	Geometric Mean (CV%)	3.72 (51.78%)	2.62 (70.32%)	5.77 (42.05%)	7.79 (67.98%)
	95% CI	2.55, 5.42	1.60, 4.27	4.22, 7.87	4.85, 12.52
	Median	4.31	2.27	6.47	5.95
	Q25, Q75	2.58, 5.16	2.10, 4.03	5.18, 6.79	5.86, 10.00
	Min, Max	1.8, 7.5	1.1, 7.8	2.4, 10.0	4.9, 34.1
	Mean (SD)	4.11 (1.832)	3.15 (2.169)	6.15 (2.122)	9.78 (9.339)

Note: NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: BLOQ values before T<sub>0</sub> are treated as zero. BLOQ values after the last quantifiable value are treated as missing. Any other BLOQ values are imputed by LLOQ/2.

Note: LLOQ = 0.2 ng/mL.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_pkconc2.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.6.1 Descriptive Statistics of the Plasma Nicotine Concentrations (ng/mL) by Sex - PK Population**

Timepoint	Statistic	Group-2 PK			
		THS 2.2 Menthol (N=18)		NRT gum (N=18)	
		Males (N=9)	Females (N=9)	Males (N=9)	Females (N=9)
T <sub>0</sub> + 2 h	n	9	9	9	9
	BLOQ - n (%)	0	0	0	0
	Geometric Mean (CV%)	2.62 (52.73%)	1.71 (69.89%)	3.99 (52.26%)	4.77 (51.44%)
	95% CI	1.79, 3.84	1.05, 2.79	2.73, 5.83	3.28, 6.92
	Median	2.61	1.61	4.40	4.29
	Q25, Q75	1.66, 3.58	1.37, 2.87	3.82, 4.88	3.41, 8.50
	Min, Max	1.3, 5.3	0.7, 5.0	1.6, 7.6	2.7, 8.9
	Mean (SD)	2.91 (1.400)	2.05 (1.341)	4.39 (1.886)	5.31 (2.652)

Note: NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: BLOQ values before T<sub>0</sub> are treated as zero. BLOQ values after the last quantifiable value are treated as missing. Any other BLOQ values are imputed by LLOQ/2.

Note: LLOQ = 0.2 ng/mL.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_pkconc2.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.6.1 Descriptive Statistics of the Plasma Nicotine Concentrations (ng/mL) by Sex - PK Population**

Timepoint	Statistic	Group-2 PK			
		THS 2.2 Menthol (N=18)		NRT gum (N=18)	
		Males (N=9)	Females (N=9)	Males (N=9)	Females (N=9)
T <sub>0</sub> + 3 h	n	0	0	9	9
	BLOQ - n (%)			0	0
	Geometric Mean (CV%)			2.98 (59.90%)	3.23 (65.51%)
	95% CI			1.95, 4.57	2.04, 5.12
	Median			3.43	2.70
	Q25, Q75			2.40, 3.84	2.17, 6.72
	Min, Max			1.1, 5.8	1.6, 7.3
	Mean (SD)			3.36 (1.573)	3.82 (2.388)

Note: NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: BLOQ values before T<sub>0</sub> are treated as zero. BLOQ values after the last quantifiable value are treated as missing. Any other BLOQ values are imputed by LLOQ/2.

Note: LLOQ = 0.2 ng/mL.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_pkconc2.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.6.1 Descriptive Statistics of the Plasma Nicotine Concentrations (ng/mL) by Sex - PK Population**

Timepoint	Statistic	Group-2 PK			
		THS 2.2 Menthol (N=18)		NRT gum (N=18)	
		Males (N=9)	Females (N=9)	Males (N=9)	Females (N=9)
T <sub>0</sub> + 4 h	n	9	9	9	9
	BLOQ - n (%)	0	0	0	0
	Geometric Mean (CV%)	1.53 (55.76%)	0.94 (66.89%)	2.37 (63.43%)	2.29 (84.07%)
	95% CI	1.02, 2.28	0.58, 1.50	1.51, 3.71	1.30, 4.01
	Median	1.50	0.80	2.80	1.61
	Q25, Q75	0.96, 2.00	0.64, 1.19	1.66, 2.93	1.55, 5.11
	Min, Max	0.8, 3.3	0.4, 3.0	0.9, 4.9	1.1, 6.4
	Mean (SD)	1.73 (0.943)	1.12 (0.823)	2.71 (1.380)	2.94 (2.287)

Note: NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: BLOQ values before T<sub>0</sub> are treated as zero. BLOQ values after the last quantifiable value are treated as missing. Any other BLOQ values are imputed by LLOQ/2.

Note: LLOQ = 0.2 ng/mL.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_pkconc2.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.6.1 Descriptive Statistics of the Plasma Nicotine Concentrations (ng/mL) by Sex - PK Population**

Timepoint	Statistic	Group-2 PK			
		THS 2.2 Menthol (N=18)		NRT gum (N=18)	
		Males (N=9)	Females (N=9)	Males (N=9)	Females (N=9)
T <sub>0</sub> + 6 h	n	9	8	9	9
	BLOQ - n (%)	0	1 (11.1%)	0	0
	Geometric Mean (CV%)	0.94 (64.33%)	0.61 (70.32%)	1.36 (87.02%)	1.30 (108.99%)
	95% CI	0.59, 1.48	0.35, 1.04	0.76, 2.42	0.65, 2.57
	Median	0.83	0.49	1.69	0.92
	Q25, Q75	0.59, 1.35	0.37, 0.94	0.94, 1.88	0.79, 3.50
	Min, Max	0.5, 2.6	0.3, 2.0	0.4, 3.7	0.4, 4.4
	Mean (SD)	1.11 (0.734)	0.74 (0.573)	1.69 (1.062)	1.86 (1.660)

Note: NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: BLOQ values before T<sub>0</sub> are treated as zero. BLOQ values after the last quantifiable value are treated as missing. Any other BLOQ values are imputed by LLOQ/2.

Note: LLOQ = 0.2 ng/mL.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_pkconc2.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.6.1 Descriptive Statistics of the Plasma Nicotine Concentrations (ng/mL) by Sex - PK Population**

Timepoint	Statistic	Group-2 PK			
		THS 2.2 Menthol (N=18)		NRT gum (N=18)	
		Males (N=9)	Females (N=9)	Males (N=9)	Females (N=9)
T <sub>0</sub> + 9 h	n	9	6	8	8
	BLOQ - n (%)	0	3 (33.3%)	1 (11.1%)	1 (11.1%)
	Geometric Mean (CV%)	0.58 (69.75%)	0.44 (66.35%)	0.90 (77.71%)	0.82 (134.47%)
	95% CI	0.35, 0.94	0.23, 0.83	0.50, 1.60	0.35, 1.93
	Median	0.52	0.41	0.89	0.49
	Q25, Q75	0.40, 0.80	0.29, 0.58	0.61, 1.33	0.41, 2.68
	Min, Max	0.2, 1.9	0.2, 1.2	0.3, 2.8	0.3, 2.9
	Mean (SD)	0.69 (0.495)	0.52 (0.363)	1.10 (0.780)	1.28 (1.208)

Note: NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: BLOQ values before T<sub>0</sub> are treated as zero. BLOQ values after the last quantifiable value are treated as missing. Any other BLOQ values are imputed by LLOQ/2.

Note: LLOQ = 0.2 ng/mL.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_pkconc2.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.6.1 Descriptive Statistics of the Plasma Nicotine Concentrations (ng/mL) by Sex - PK Population**

Timepoint	Statistic	Group-2 PK			
		THS 2.2 Menthol (N=18)		NRT gum (N=18)	
		Males (N=9)	Females (N=9)	Males (N=9)	Females (N=9)
T <sub>0</sub> + 12 h	n	8	4	8	7
	BLOQ - n (%)	1 (11.1%)	5 (55.6%)	1 (11.1%)	2 (22.2%)
	Geometric Mean (CV%)	0.40 (58.85%)	0.36 (62.71%)	0.55 (71.50%)	0.60 (147.29%)
	95% CI	0.25, 0.64	0.14, 0.92	0.32, 0.95	0.22, 1.63
	Median	0.31	0.31	0.51	0.44
	Q25, Q75	0.27, 0.54	0.25, 0.59	0.36, 0.89	0.21, 1.82
	Min, Max	0.3, 1.3	0.2, 0.8	0.2, 1.5	0.2, 1.9
	Mean (SD)	0.47 (0.339)	0.42 (0.282)	0.66 (0.425)	0.94 (0.841)

Note: NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: BLOQ values before T<sub>0</sub> are treated as zero. BLOQ values after the last quantifiable value are treated as missing. Any other BLOQ values are imputed by LLOQ/2.

Note: LLOQ = 0.2 ng/mL.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_pkconc2.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.6.1 Descriptive Statistics of the Plasma Nicotine Concentrations (ng/mL) by Sex - PK Population**

Timepoint	Statistic	Group-2 PK			
		THS 2.2 Menthol (N=18)		NRT gum (N=18)	
		Males (N=9)	Females (N=9)	Males (N=9)	Females (N=9)
T <sub>0</sub> + 24 h	n	4	2	4	4
	BLOQ - n (%)	5 (55.6%)	7 (77.8%)	5 (55.6%)	5 (55.6%)
	Geometric Mean (CV%)	0.33 (33.23%)	NC	0.36 (41.13%)	0.52 (44.38%)
	95% CI	0.19, 0.56	NC	0.18, 0.67	0.26, 1.03
	Median	0.31	NC	0.40	0.62
	Q25, Q75	0.26, 0.43	NC	0.27, 0.48	0.44, 0.67
	Min, Max	0.2, 0.5	NC	0.2, 0.5	0.3, 0.7
	Mean (SD)	0.34 (0.119)	NC	0.37 (0.129)	0.55 (0.189)

Note: NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Note: NC = Not calculated.

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: BLOQ values before T<sub>0</sub> are treated as zero. BLOQ values after the last quantifiable value are treated as missing. Any other BLOQ values are imputed by LLOQ/2.

Note: LLOQ = 0.2 ng/mL.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_pkconc2.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.6.2 Descriptive Statistics of the Plasma Nicotine Concentrations (ng/mL) by Nicotine Level - PK Population**

Timepoint	Statistic	Group-1 PK			
		THS 2.2 Menthol (N=43)		mCC (N=43)	
		<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)	<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)
15 min < T <sub>0</sub>	n	25	18	25	18
	BLOQ - n (%)	13 (52.0%)	10 (55.6%)	15 (60.0%)	7 (38.9%)
	Geometric Mean (CV%)	NC	NC	NC	NC
	95% CI	NC	NC	NC	NC
	Median	0.00	0.00	0.00	0.21
	Q25, Q75	0.00, 0.32	0.00, 0.38	0.00, 0.33	0.00, 0.45
	Min, Max	0.0, 0.7	0.0, 2.2	0.0, 1.4	0.0, 2.0
	Mean (SD)	0.18 (0.211)	0.30 (0.561)	0.20 (0.335)	0.37 (0.521)

Note: mCC = menthol conventional cigarettes; THS = Tobacco Heating System.

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

Note: NC = Not calculated.

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: BLOQ values before T<sub>0</sub> are treated as zero. BLOQ values after the last quantifiable value are treated as missing. Any other BLOQ values are imputed by LLOQ/2.

Note: LLOQ = 0.2 ng/mL.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_pkconc3.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.6.2 Descriptive Statistics of the Plasma Nicotine Concentrations (ng/mL) by Nicotine Level - PK Population**

Timepoint	Statistic	Group-1 PK			
		THS 2.2 Menthol (N=43)		mCC (N=43)	
		<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)	<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)
T <sub>0</sub> + 2 min	n	25	18	25	18
	BLOQ - n (%)	0	0	0	1 (5.6%)
	Geometric Mean (CV%)	3.16 (162.93%)	1.57 (196.59%)	3.65 (125.87%)	2.92 (309.33%)
	95% CI	1.97, 5.07	0.84, 2.94	2.44, 5.46	1.35, 6.26
	Median	3.56	1.50	2.63	3.06
	Q25, Q75	1.70, 7.57	0.74, 2.95	1.77, 8.15	0.97, 6.69
	Min, Max	0.2, 35.1	0.2, 19.2	0.7, 29.0	0.1, 36.6
	Mean (SD)	5.59 (7.113)	3.38 (4.975)	5.98 (6.998)	7.10 (9.594)

Note: mCC = menthol conventional cigarettes; THS = Tobacco Heating System.

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

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: BLOQ values before T<sub>0</sub> are treated as zero. BLOQ values after the last quantifiable value are treated as missing. Any other BLOQ values are imputed by LLOQ/2.

Note: LLOQ = 0.2 ng/mL.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_pkconc3.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.6.2 Descriptive Statistics of the Plasma Nicotine Concentrations (ng/mL) by Nicotine Level - PK Population**

Timepoint	Statistic	Group-1 PK			
		THS 2.2 Menthol (N=43)		mCC (N=43)	
		<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)	<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)
T <sub>0</sub> + 4 min	n	25	18	25	18
	BLOQ - n (%)	0	0	0	0
	Geometric Mean (CV%)	8.97 (89.93%)	4.18 (223.47%)	8.31 (99.60%)	7.84 (204.66%)
	95% CI	6.52, 12.33	2.14, 8.14	5.89, 11.71	4.14, 14.85
	Median	11.40	5.02	8.40	7.94
	Q25, Q75	5.07, 15.70	1.54, 11.70	5.03, 13.00	3.59, 23.20
	Min, Max	1.9, 38.1	0.3, 31.3	1.2, 34.2	0.4, 47.6
	Mean (SD)	11.64 (8.486)	8.29 (9.435)	11.33 (9.292)	13.84 (12.899)

Note: mCC = menthol conventional cigarettes; THS = Tobacco Heating System.

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

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: BLOQ values before T<sub>0</sub> are treated as zero. BLOQ values after the last quantifiable value are treated as missing. Any other BLOQ values are imputed by LLOQ/2.

Note: LLOQ = 0.2 ng/mL.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_pkconc3.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.6.2 Descriptive Statistics of the Plasma Nicotine Concentrations (ng/mL) by Nicotine Level - PK Population**

Timepoint	Statistic	Group-1 PK			
		THS 2.2 Menthol (N=43)		mCC (N=43)	
		<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)	<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)
T <sub>0</sub> + 6 min	n	25	18	25	18
	BLOQ - n (%)	0	0	0	0
	Geometric Mean (CV%)	10.80 (61.82%)	5.74 (215.92%)	9.02 (73.27%)	9.48 (138.70%)
	95% CI	8.54, 13.67	2.98, 11.05	6.88, 11.83	5.66, 15.87
	Median	9.27	5.77	8.97	9.92
	Q25, Q75	7.59, 16.80	2.93, 18.60	6.05, 17.40	5.39, 15.60
	Min, Max	3.7, 34.8	0.4, 43.8	2.4, 24.4	1.2, 58.1
	Mean (SD)	12.60 (7.343)	11.01 (11.867)	10.85 (6.253)	14.81 (14.605)

Note: mCC = menthol conventional cigarettes; THS = Tobacco Heating System.

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

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: BLOQ values before T<sub>0</sub> are treated as zero. BLOQ values after the last quantifiable value are treated as missing. Any other BLOQ values are imputed by LLOQ/2.

Note: LLOQ = 0.2 ng/mL.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_pkconc3.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.6.2 Descriptive Statistics of the Plasma Nicotine Concentrations (ng/mL) by Nicotine Level - PK Population**

Timepoint	Statistic	Group-1 PK			
		THS 2.2 Menthol (N=43)		mCC (N=43)	
		<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)	<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)
T <sub>0</sub> + 8 min	n	25	18	25	18
	BLOQ - n (%)	0	0	0	0
	Geometric Mean (CV%)	10.47 (59.13%)	5.28 (165.94%)	8.21 (57.15%)	9.11 (103.09%)
	95% CI	8.34, 13.12	2.98, 9.37	6.58, 10.22	5.96, 13.91
	Median	11.10	5.70	7.74	9.57
	Q25, Q75	6.96, 15.90	3.44, 13.20	6.06, 12.90	6.36, 17.80
	Min, Max	4.1, 29.7	0.4, 27.5	2.6, 20.4	1.4, 36.0
	Mean (SD)	12.07 (6.666)	8.52 (7.662)	9.30 (4.575)	12.10 (8.761)

Note: mCC = menthol conventional cigarettes; THS = Tobacco Heating System.

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

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: BLOQ values before T<sub>0</sub> are treated as zero. BLOQ values after the last quantifiable value are treated as missing. Any other BLOQ values are imputed by LLOQ/2.

Note: LLOQ = 0.2 ng/mL.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_pkconc3.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.6.2 Descriptive Statistics of the Plasma Nicotine Concentrations (ng/mL) by Nicotine Level - PK Population**

Timepoint	Statistic	Group-1 PK			
		THS 2.2 Menthol (N=43)		mCC (N=43)	
		<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)	<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)
T <sub>0</sub> + 10 min	n	25	18	25	18
	BLOQ - n (%)	0	0	0	0
	Geometric Mean (CV%)	9.75 (55.70%)	4.88 (164.73%)	8.29 (57.67%)	8.68 (83.72%)
	95% CI	7.86, 12.08	2.76, 8.64	6.64, 10.34	6.03, 12.47
	Median	9.11	5.52	8.22	9.06
	Q25, Q75	6.88, 14.00	3.63, 9.38	6.55, 12.10	5.03, 14.00
	Min, Max	3.7, 26.3	0.3, 24.3	2.5, 20.1	2.3, 31.5
	Mean (SD)	11.06 (5.711)	7.66 (6.575)	9.40 (4.631)	10.99 (7.802)

Note: mCC = menthol conventional cigarettes; THS = Tobacco Heating System.

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

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: BLOQ values before T<sub>0</sub> are treated as zero. BLOQ values after the last quantifiable value are treated as missing. Any other BLOQ values are imputed by LLOQ/2.

Note: LLOQ = 0.2 ng/mL.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_pkconc3.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.6.2 Descriptive Statistics of the Plasma Nicotine Concentrations (ng/mL) by Nicotine Level - PK Population**

Timepoint	Statistic	Group-1 PK			
		THS 2.2 Menthol (N=43)		mCC (N=43)	
		<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)	<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)
T <sub>0</sub> + 15 min	n	25	18	25	18
	BLOQ - n (%)	0	0	0	0
	Geometric Mean (CV%)	7.96 (54.73%)	4.26 (124.68%)	6.25 (50.59%)	6.85 (66.75%)
	95% CI	6.44, 9.83	2.63, 6.90	5.13, 7.62	5.06, 9.26
	Median	8.13	5.19	6.48	7.61
	Q25, Q75	6.69, 11.90	3.84, 6.86	5.47, 8.53	4.96, 10.70
	Min, Max	2.3, 17.2	0.3, 20.4	1.3, 12.7	2.2, 20.0
	Mean (SD)	8.91 (4.073)	5.91 (4.583)	6.84 (2.619)	8.02 (4.485)

Note: mCC = menthol conventional cigarettes; THS = Tobacco Heating System.

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

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: BLOQ values before T<sub>0</sub> are treated as zero. BLOQ values after the last quantifiable value are treated as missing. Any other BLOQ values are imputed by LLOQ/2.

Note: LLOQ = 0.2 ng/mL.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_pkconc3.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.6.2 Descriptive Statistics of the Plasma Nicotine Concentrations (ng/mL) by Nicotine Level - PK Population**

Timepoint	Statistic	Group-1 PK			
		THS 2.2 Menthol (N=43)		mCC (N=43)	
		<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)	<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)
T <sub>0</sub> + 30 min	n	25	18	25	18
	BLOQ - n (%)	0	0	0	0
	Geometric Mean (CV%)	6.32 (54.28%)	3.75 (111.88%)	5.20 (49.16%)	5.85 (64.88%)
	95% CI	5.12, 7.79	2.39, 5.88	4.29, 6.31	4.35, 7.86
	Median	6.79	4.40	5.42	6.67
	Q25, Q75	5.46, 8.69	3.02, 6.33	4.63, 7.14	4.07, 8.43
	Min, Max	1.9, 13.1	0.3, 13.3	1.8, 11.4	2.0, 14.7
	Mean (SD)	7.01 (2.927)	4.98 (3.403)	5.71 (2.320)	6.79 (3.554)

Note: mCC = menthol conventional cigarettes; THS = Tobacco Heating System.

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

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: BLOQ values before T<sub>0</sub> are treated as zero. BLOQ values after the last quantifiable value are treated as missing. Any other BLOQ values are imputed by LLOQ/2.

Note: LLOQ = 0.2 ng/mL.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_pkconc3.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.6.2 Descriptive Statistics of the Plasma Nicotine Concentrations (ng/mL) by Nicotine Level - PK Population**

Timepoint	Statistic	Group-1 PK			
		THS 2.2 Menthol (N=43)		mCC (N=43)	
		<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)	<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)
T <sub>0</sub> + 45 min	n	25	18	25	18
	BLOQ - n (%)	0	1 (5.6%)	0	0
	Geometric Mean (CV%)	5.83 (56.36%)	3.17 (137.51%)	4.66 (49.53%)	5.55 (61.97%)
	95% CI	4.69, 7.25	1.90, 5.30	3.84, 5.66	4.17, 7.37
	Median	5.94	4.10	5.39	5.78
	Q25, Q75	5.18, 7.97	3.05, 5.18	3.80, 6.06	4.74, 7.28
	Min, Max	1.3, 13.6	0.1, 9.6	1.5, 9.3	1.8, 15.2
	Mean (SD)	6.52 (2.834)	4.22 (2.450)	5.10 (1.971)	6.40 (3.496)

Note: mCC = menthol conventional cigarettes; THS = Tobacco Heating System.

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

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: BLOQ values before T<sub>0</sub> are treated as zero. BLOQ values after the last quantifiable value are treated as missing. Any other BLOQ values are imputed by LLOQ/2.

Note: LLOQ = 0.2 ng/mL.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_pkconc3.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.6.2 Descriptive Statistics of the Plasma Nicotine Concentrations (ng/mL) by Nicotine Level - PK Population**

Timepoint	Statistic	Group-1 PK			
		THS 2.2 Menthol (N=43)		mCC (N=43)	
		<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)	<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)
T <sub>0</sub> + 60 min	n	25	18	25	18
	BLOQ - n (%)	0	0	0	0
	Geometric Mean (CV%)	5.25 (56.18%)	3.30 (113.96%)	4.33 (51.69%)	5.09 (65.73%)
	95% CI	4.22, 6.52	2.09, 5.20	3.54, 5.30	3.78, 6.87
	Median	5.56	3.92	5.02	5.64
	Q25, Q75	4.52, 7.31	2.52, 4.91	3.27, 6.10	4.02, 7.13
	Min, Max	1.4, 12.3	0.2, 10.9	1.5, 9.1	1.6, 13.4
	Mean (SD)	5.86 (2.552)	4.37 (2.905)	4.79 (1.988)	5.94 (3.249)

Note: mCC = menthol conventional cigarettes; THS = Tobacco Heating System.

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

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: BLOQ values before T<sub>0</sub> are treated as zero. BLOQ values after the last quantifiable value are treated as missing. Any other BLOQ values are imputed by LLOQ/2.

Note: LLOQ = 0.2 ng/mL.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_pkconc3.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.6.2 Descriptive Statistics of the Plasma Nicotine Concentrations (ng/mL) by Nicotine Level - PK Population**

Timepoint	Statistic	Group-1 PK			
		THS 2.2 Menthol (N=43)		mCC (N=43)	
		<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)	<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)
T <sub>0</sub> + 2 h	n	25	18	25	18
	BLOQ - n (%)	0	0	0	0
	Geometric Mean (CV%)	3.59 (67.08%)	2.43 (103.75%)	2.92 (61.22%)	3.69 (69.94%)
	95% CI	2.79, 4.63	1.58, 3.72	2.31, 3.69	2.69, 5.06
	Median	3.78	2.54	3.34	3.91
	Q25, Q75	2.45, 5.52	2.05, 3.59	2.14, 4.28	2.68, 6.50
	Min, Max	0.9, 10.6	0.2, 8.8	0.9, 6.6	1.2, 9.9
	Mean (SD)	4.23 (2.404)	3.19 (2.333)	3.34 (1.597)	4.41 (2.629)

Note: mCC = menthol conventional cigarettes; THS = Tobacco Heating System.

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

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: BLOQ values before T<sub>0</sub> are treated as zero. BLOQ values after the last quantifiable value are treated as missing. Any other BLOQ values are imputed by LLOQ/2.

Note: LLOQ = 0.2 ng/mL.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_pkconc3.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.6.2 Descriptive Statistics of the Plasma Nicotine Concentrations (ng/mL) by Nicotine Level - PK Population**

Timepoint	Statistic	Group-1 PK			
		THS 2.2 Menthol (N=43)		mCC (N=43)	
		<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)	<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)
T <sub>0</sub> + 4 h	n	25	18	25	18
	BLOQ - n (%)	0	1 (5.6%)	0	0
	Geometric Mean (CV%)	1.84 (90.14%)	1.41 (123.29%)	1.46 (78.88%)	2.09 (82.80%)
	95% CI	1.34, 2.54	0.87, 2.28	1.09, 1.95	1.45, 2.99
	Median	1.69	1.38	1.40	1.87
	Q25, Q75	1.24, 3.57	0.88, 2.10	0.84, 2.32	1.29, 4.31
	Min, Max	0.4, 6.7	0.1, 6.2	0.4, 3.8	0.6, 6.6
	Mean (SD)	2.38 (1.651)	2.01 (1.691)	1.80 (1.144)	2.65 (1.861)

Note: mCC = menthol conventional cigarettes; THS = Tobacco Heating System.

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

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: BLOQ values before T<sub>0</sub> are treated as zero. BLOQ values after the last quantifiable value are treated as missing. Any other BLOQ values are imputed by LLOQ/2.

Note: LLOQ = 0.2 ng/mL.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_pkconc3.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.6.2 Descriptive Statistics of the Plasma Nicotine Concentrations (ng/mL) by Nicotine Level - PK Population**

Timepoint	Statistic	Group-1 PK			
		THS 2.2 Menthol (N=43)		mCC (N=43)	
		<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)	<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)
T <sub>0</sub> + 6 h	n	25	18	25	18
	BLOQ - n (%)	0	0	0	0
	Geometric Mean (CV%)	1.11 (104.58%)	1.02 (95.64%)	0.90 (93.28%)	1.37 (102.60%)
	95% CI	0.78, 1.59	0.68, 1.52	0.65, 1.25	0.90, 2.10
	Median	0.95	0.99	0.74	1.24
	Q25, Q75	0.63, 2.62	0.53, 1.55	0.51, 1.54	0.70, 3.23
	Min, Max	0.2, 4.9	0.3, 4.4	0.2, 3.2	0.4, 5.7
	Mean (SD)	1.56 (1.278)	1.39 (1.214)	1.20 (0.932)	1.92 (1.615)

Note: mCC = menthol conventional cigarettes; THS = Tobacco Heating System.

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

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: BLOQ values before T<sub>0</sub> are treated as zero. BLOQ values after the last quantifiable value are treated as missing. Any other BLOQ values are imputed by LLOQ/2.

Note: LLOQ = 0.2 ng/mL.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_pkconc3.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.6.2 Descriptive Statistics of the Plasma Nicotine Concentrations (ng/mL) by Nicotine Level - PK Population**

Timepoint	Statistic	Group-1 PK			
		THS 2.2 Menthol (N=43)		mCC (N=43)	
		<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)	<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)
T <sub>0</sub> + 9 h	n	23	16	22	18
	BLOQ - n (%)	3 (12.0%)	2 (11.1%)	3 (12.0%)	0
	Geometric Mean (CV%)	0.68 (124.33%)	0.71 (98.21%)	0.59 (91.15%)	0.85 (110.35%)
	95% CI	0.44, 1.04	0.45, 1.10	0.41, 0.84	0.54, 1.34
	Median	0.51	0.62	0.49	0.77
	Q25, Q75	0.30, 1.88	0.39, 1.36	0.31, 1.01	0.38, 2.25
	Min, Max	0.1, 3.9	0.2, 3.0	0.2, 2.3	0.2, 3.8
	Mean (SD)	1.03 (0.968)	1.00 (0.918)	0.80 (0.664)	1.25 (1.124)

Note: mCC = menthol conventional cigarettes; THS = Tobacco Heating System.

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

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: BLOQ values before T<sub>0</sub> are treated as zero. BLOQ values after the last quantifiable value are treated as missing. Any other BLOQ values are imputed by LLOQ/2.

Note: LLOQ = 0.2 ng/mL.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_pkconc3.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.6.2 Descriptive Statistics of the Plasma Nicotine Concentrations (ng/mL) by Nicotine Level - PK Population**

Timepoint	Statistic	Group-1 PK			
		THS 2.2 Menthol (N=43)		mCC (N=43)	
		<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)	<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)
T <sub>0</sub> + 12 h	n	18	15	15	13
	BLOQ - n (%)	7 (28.0%)	3 (16.7%)	10 (40.0%)	5 (27.8%)
	Geometric Mean (CV%)	0.66 (143.01%)	0.59 (135.44%)	0.60 (68.29%)	0.80 (94.22%)
	95% CI	0.39, 1.12	0.33, 1.05	0.42, 0.86	0.49, 1.30
	Median	0.54	0.40	0.54	0.66
	Q25, Q75	0.26, 1.25	0.25, 1.38	0.34, 1.20	0.46, 1.76
	Min, Max	0.2, 10.6	0.2, 5.8	0.3, 1.7	0.3, 3.0
	Mean (SD)	1.32 (2.405)	1.05 (1.444)	0.73 (0.489)	1.07 (0.858)

Note: mCC = menthol conventional cigarettes; THS = Tobacco Heating System.

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

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: BLOQ values before T<sub>0</sub> are treated as zero. BLOQ values after the last quantifiable value are treated as missing. Any other BLOQ values are imputed by LLOQ/2.

Note: LLOQ = 0.2 ng/mL.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_pkconc3.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.6.2 Descriptive Statistics of the Plasma Nicotine Concentrations (ng/mL) by Nicotine Level - PK Population**

Timepoint	Statistic	Group-1 PK			
		THS 2.2 Menthol (N=43)		mCC (N=43)	
		<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)	<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)
T <sub>0</sub> + 24 h	n	13	8	11	11
	BLOQ - n (%)	12 (48.0%)	10 (55.6%)	14 (56.0%)	7 (38.9%)
	Geometric Mean (CV%)	0.43 (80.63%)	0.43 (55.73%)	0.34 (45.65%)	0.42 (72.01%)
	95% CI	0.28, 0.67	0.27, 0.67	0.25, 0.47	0.27, 0.65
	Median	0.37	0.42	0.31	0.33
	Q25, Q75	0.31, 0.46	0.28, 0.69	0.21, 0.50	0.22, 0.68
	Min, Max	0.2, 2.9	0.2, 0.9	0.2, 0.8	0.2, 1.3
	Mean (SD)	0.60 (0.730)	0.48 (0.251)	0.38 (0.178)	0.51 (0.356)

Note: mCC = menthol conventional cigarettes; THS = Tobacco Heating System.

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

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: BLOQ values before T<sub>0</sub> are treated as zero. BLOQ values after the last quantifiable value are treated as missing. Any other BLOQ values are imputed by LLOQ/2.

Note: LLOQ = 0.2 ng/mL.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_pkconc3.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.6.2 Descriptive Statistics of the Plasma Nicotine Concentrations (ng/mL) by Nicotine Level - PK Population**

Timepoint	Statistic	Group-2 PK			
		THS 2.2 Menthol (N=18)		NRT gum (N=18)	
		<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)	<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)
15 min < T <sub>0</sub>	n	10	8	10	8
	BLOQ - n (%)	7 (70.0%)	5 (62.5%)	4 (40.0%)	3 (37.5%)
	Geometric Mean (CV%)	NC	NC	NC	NC
	95% CI	NC	NC	NC	NC
	Median	0.00	0.00	0.23	0.22
	Q25, Q75	0.00, 0.26	0.00, 0.30	0.00, 0.44	0.00, 0.41
	Min, Max	0.0, 0.8	0.0, 0.9	0.0, 0.6	0.0, 0.9
	Mean (SD)	0.13 (0.265)	0.18 (0.311)	0.23 (0.225)	0.27 (0.320)

Note: NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Note: NC = Not calculated.

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: BLOQ values before T<sub>0</sub> are treated as zero. BLOQ values after the last quantifiable value are treated as missing. Any other BLOQ values are imputed by LLOQ/2.

Note: LLOQ = 0.2 ng/mL.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_pkconc3.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.6.2 Descriptive Statistics of the Plasma Nicotine Concentrations (ng/mL) by Nicotine Level - PK Population**

Timepoint	Statistic	Group-2 PK			
		THS 2.2 Menthol (N=18)		NRT gum (N=18)	
		<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)	<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)
T <sub>0</sub> + 2 min	n	10	8	0	0
	BLOQ - n (%)	0	0		
	Geometric Mean (CV%)	2.47 (238.61%)	2.92 (57.05%)		
	95% CI	0.92, 6.62	1.87, 4.55		
	Median	1.72	3.53		
	Q25, Q75	1.21, 7.23	1.64, 4.41		
	Min, Max	0.3, 30.2	1.5, 5.6		
	Mean (SD)	5.78 (9.041)	3.27 (1.551)		

Note: NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: BLOQ values before T<sub>0</sub> are treated as zero. BLOQ values after the last quantifiable value are treated as missing. Any other BLOQ values are imputed by LLOQ/2.

Note: LLOQ = 0.2 ng/mL.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_pkconc3.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.6.2 Descriptive Statistics of the Plasma Nicotine Concentrations (ng/mL) by Nicotine Level - PK Population**

Timepoint	Statistic	Group-2 PK			
		THS 2.2 Menthol (N=18)		NRT gum (N=18)	
		<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)	<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)
T <sub>0</sub> + 4 min	n	10	8	0	0
	BLOQ - n (%)	0	0		
	Geometric Mean (CV%)	4.90 (213.28%)	6.59 (63.85%)		
	95% CI	1.92, 12.51	4.04, 10.75		
	Median	4.36	6.62		
	Q25, Q75	2.80, 11.50	5.00, 10.49		
	Min, Max	0.6, 45.3	2.3, 14.0		
	Mean (SD)	10.28 (14.070)	7.56 (4.036)		

Note: NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: BLOQ values before T<sub>0</sub> are treated as zero. BLOQ values after the last quantifiable value are treated as missing. Any other BLOQ values are imputed by LLOQ/2.

Note: LLOQ = 0.2 ng/mL.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_pkconc3.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.6.2 Descriptive Statistics of the Plasma Nicotine Concentrations (ng/mL) by Nicotine Level - PK Population**

Timepoint	Statistic	Group-2 PK			
		THS 2.2 Menthol (N=18)		NRT gum (N=18)	
		<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)	<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)
T <sub>0</sub> + 6 min	n	10	8	0	0
	BLOQ - n (%)	0	0		
	Geometric Mean (CV%)	5.96 (150.61%)	7.23 (73.69%)		
	95% CI	2.73, 12.99	4.16, 12.54		
	Median	5.59	8.11		
	Q25, Q75	3.28, 9.84	5.18, 9.95		
	Min, Max	1.2, 45.8	2.1, 20.3		
	Mean (SD)	10.55 (13.694)	8.61 (5.463)		

Note: NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: BLOQ values before T<sub>0</sub> are treated as zero. BLOQ values after the last quantifiable value are treated as missing. Any other BLOQ values are imputed by LLOQ/2.

Note: LLOQ = 0.2 ng/mL.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_pkconc3.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.6.2 Descriptive Statistics of the Plasma Nicotine Concentrations (ng/mL) by Nicotine Level - PK Population**

Timepoint	Statistic	Group-2 PK			
		THS 2.2 Menthol (N=18)		NRT gum (N=18)	
		<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)	<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)
T <sub>0</sub> + 8 min	n	10	8	0	0
	BLOQ - n (%)	0	0		
	Geometric Mean (CV%)	5.04 (118.40%)	6.46 (62.49%)		
	95% CI	2.58, 9.86	3.99, 10.44		
	Median	4.57	6.95		
	Q25, Q75	2.55, 8.63	5.46, 8.12		
	Min, Max	1.2, 28.9	2.1, 15.8		
	Mean (SD)	7.71 (8.517)	7.36 (3.966)		

Note: NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: BLOQ values before T<sub>0</sub> are treated as zero. BLOQ values after the last quantifiable value are treated as missing. Any other BLOQ values are imputed by LLOQ/2.

Note: LLOQ = 0.2 ng/mL.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_pkconc3.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.6.2 Descriptive Statistics of the Plasma Nicotine Concentrations (ng/mL) by Nicotine Level - PK Population**

Timepoint	Statistic	Group-2 PK			
		THS 2.2 Menthol (N=18)		NRT gum (N=18)	
		<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)	<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)
T <sub>0</sub> + 10 min	n	10	8	10	8
	BLOQ - n (%)	0	0	0	0
	Geometric Mean (CV%)	5.22 (110.11%)	6.29 (59.26%)	1.49 (68.88%)	1.33 (44.80%)
	95% CI	2.76, 9.89	3.97, 9.95	0.95, 2.33	0.93, 1.91
	Median	5.59	6.92	1.69	1.29
	Q25, Q75	2.79, 9.97	5.39, 7.98	0.69, 2.32	1.03, 1.79
	Min, Max	1.1, 22.1	2.1, 13.8	0.7, 4.2	0.7, 2.7
	Mean (SD)	7.29 (6.328)	7.06 (3.378)	1.76 (1.073)	1.44 (0.627)

Note: NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: BLOQ values before T<sub>0</sub> are treated as zero. BLOQ values after the last quantifiable value are treated as missing. Any other BLOQ values are imputed by LLOQ/2.

Note: LLOQ = 0.2 ng/mL.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_pkconc3.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.6.2 Descriptive Statistics of the Plasma Nicotine Concentrations (ng/mL) by Nicotine Level - PK Population**

Timepoint	Statistic	Group-2 PK			
		THS 2.2 Menthol (N=18)		NRT gum (N=18)	
		<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)	<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)
T <sub>0</sub> + 15 min	n	10	8	0	0
	BLOQ - n (%)	0	0		
	Geometric Mean (CV%)	4.11 (77.06%)	5.59 (58.55%)		
	95% CI	2.52, 6.70	3.54, 8.80		
	Median	4.09	5.88		
	Q25, Q75	2.55, 6.22	3.85, 8.84		
	Min, Max	1.3, 13.4	2.4, 10.9		
	Mean (SD)	5.04 (3.536)	6.30 (3.091)		

Note: NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: BLOQ values before T<sub>0</sub> are treated as zero. BLOQ values after the last quantifiable value are treated as missing. Any other BLOQ values are imputed by LLOQ/2.

Note: LLOQ = 0.2 ng/mL.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_pkconc3.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.6.2 Descriptive Statistics of the Plasma Nicotine Concentrations (ng/mL) by Nicotine Level - PK Population**

Timepoint	Statistic	Group-2 PK			
		THS 2.2 Menthol (N=18)		NRT gum (N=18)	
		<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)	<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)
T <sub>0</sub> + 20 min	n	0	0	10	8
	BLOQ - n (%)			0	0
	Geometric Mean (CV%)			4.18 (56.22%)	2.98 (38.17%)
	95% CI			2.87, 6.09	2.19, 4.06
	Median			5.44	2.90
	Q25, Q75			2.45, 6.19	2.43, 3.41
	Min, Max			1.8, 7.0	1.8, 6.2
	Mean (SD)			4.66 (1.980)	3.18 (1.344)

Note: NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: BLOQ values before T<sub>0</sub> are treated as zero. BLOQ values after the last quantifiable value are treated as missing. Any other BLOQ values are imputed by LLOQ/2.

Note: LLOQ = 0.2 ng/mL.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_pkconc3.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.6.2 Descriptive Statistics of the Plasma Nicotine Concentrations (ng/mL) by Nicotine Level - PK Population**

Timepoint	Statistic	Group-2 PK			
		THS 2.2 Menthol (N=18)		NRT gum (N=18)	
		<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)	<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)
T <sub>0</sub> + 25 min	n	0	0	10	8
	BLOQ - n (%)			0	0
	Geometric Mean (CV%)			4.64 (45.48%)	3.61 (38.35%)
	95% CI			3.40, 6.33	2.65, 4.93
	Median			4.35	3.67
	Q25, Q75			3.45, 7.10	2.63, 4.62
	Min, Max			2.6, 8.6	2.2, 6.7
	Mean (SD)			5.04 (2.177)	3.84 (1.469)

Note: NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: BLOQ values before T<sub>0</sub> are treated as zero. BLOQ values after the last quantifiable value are treated as missing. Any other BLOQ values are imputed by LLOQ/2.

Note: LLOQ = 0.2 ng/mL.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_pkconc3.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.6.2 Descriptive Statistics of the Plasma Nicotine Concentrations (ng/mL) by Nicotine Level - PK Population**

Timepoint	Statistic	Group-2 PK			
		THS 2.2 Menthol (N=18)		NRT gum (N=18)	
		<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)	<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)
T <sub>0</sub> + 30 min	n	10	8	10	8
	BLOQ - n (%)	0	0	0	0
	Geometric Mean (CV%)	3.53 (73.09%)	4.23 (52.18%)	5.55 (35.37%)	4.28 (31.40%)
	95% CI	2.21, 5.64	2.80, 6.38	4.34, 7.10	3.31, 5.53
	Median	3.75	4.67	5.69	4.36
	Q25, Q75	1.91, 5.67	2.92, 6.39	4.44, 7.59	3.66, 5.22
	Min, Max	1.3, 9.3	1.9, 7.5	3.3, 8.9	2.5, 6.6
	Mean (SD)	4.24 (2.609)	4.66 (2.053)	5.84 (1.932)	4.45 (1.311)

Note: NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: BLOQ values before T<sub>0</sub> are treated as zero. BLOQ values after the last quantifiable value are treated as missing. Any other BLOQ values are imputed by LLOQ/2.

Note: LLOQ = 0.2 ng/mL.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_pkconc3.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.6.2 Descriptive Statistics of the Plasma Nicotine Concentrations (ng/mL) by Nicotine Level - PK Population**

Timepoint	Statistic	Group-2 PK			
		THS 2.2 Menthol (N=18)		NRT gum (N=18)	
		<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)	<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)
T <sub>0</sub> + 35 min	n	0	0	10	8
	BLOQ - n (%)			0	0
	Geometric Mean (CV%)			5.66 (33.34%)	4.50 (45.10%)
	95% CI			4.48, 7.14	3.14, 6.45
	Median			5.72	4.77
	Q25, Q75			4.94, 6.31	3.49, 6.04
	Min, Max			3.5, 9.8	2.3, 8.0
	Mean (SD)			5.93 (1.959)	4.86 (1.937)

Note: NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: BLOQ values before T<sub>0</sub> are treated as zero. BLOQ values after the last quantifiable value are treated as missing. Any other BLOQ values are imputed by LLOQ/2.

Note: LLOQ = 0.2 ng/mL.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_pkconc3.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.6.2 Descriptive Statistics of the Plasma Nicotine Concentrations (ng/mL) by Nicotine Level - PK Population**

Timepoint	Statistic	Group-2 PK			
		THS 2.2 Menthol (N=18)		NRT gum (N=18)	
		<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)	<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)
T <sub>0</sub> + 40 min	n	0	0	10	8
	BLOQ - n (%)			0	0
	Geometric Mean (CV%)			6.68 (27.05%)	5.21 (40.84%)
	95% CI			5.52, 8.08	3.74, 7.24
	Median			7.04	5.67
	Q25, Q75			5.24, 8.59	3.88, 7.17
	Min, Max			4.4, 9.4	2.7, 8.2
	Mean (SD)			6.89 (1.746)	5.54 (1.966)

Note: NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: BLOQ values before T<sub>0</sub> are treated as zero. BLOQ values after the last quantifiable value are treated as missing. Any other BLOQ values are imputed by LLOQ/2.

Note: LLOQ = 0.2 ng/mL.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_pkconc3.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.6.2 Descriptive Statistics of the Plasma Nicotine Concentrations (ng/mL) by Nicotine Level - PK Population**

Timepoint	Statistic	Group-2 PK			
		THS 2.2 Menthol (N=18)		NRT gum (N=18)	
		<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)	<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)
T <sub>0</sub> + 45 min	n	10	8	10	8
	BLOQ - n (%)	0	0	0	0
	Geometric Mean (CV%)	3.31 (75.04%)	3.77 (52.35%)	6.75 (32.54%)	5.28 (41.42%)
	95% CI	2.05, 5.35	2.49, 5.69	5.37, 8.47	3.78, 7.37
	Median	3.64	4.00	7.21	5.84
	Q25, Q75	2.15, 5.58	2.51, 5.56	5.20, 8.71	4.19, 6.62
	Min, Max	1.2, 8.3	1.9, 7.4	4.2, 11.1	2.5, 9.3
	Mean (SD)	3.98 (2.378)	4.17 (1.933)	7.06 (2.202)	5.63 (2.045)

Note: NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: BLOQ values before T<sub>0</sub> are treated as zero. BLOQ values after the last quantifiable value are treated as missing. Any other BLOQ values are imputed by LLOQ/2.

Note: LLOQ = 0.2 ng/mL.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_pkconc3.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.6.2 Descriptive Statistics of the Plasma Nicotine Concentrations (ng/mL) by Nicotine Level - PK Population**

Timepoint	Statistic	Group-2 PK			
		THS 2.2 Menthol (N=18)		NRT gum (N=18)	
		<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)	<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)
T <sub>0</sub> + 60 min	n	10	8	10	8
	BLOQ - n (%)	0	0	0	0
	Geometric Mean (CV%)	2.91 (72.73%)	3.41 (53.04%)	7.60 (61.35%)	5.73 (49.19%)
	95% CI	1.82, 4.64	2.24, 5.18	5.07, 11.39	3.88, 8.45
	Median	3.01	3.31	6.51	5.88
	Q25, Q75	1.86, 5.16	2.30, 4.84	5.80, 7.58	4.61, 8.23
	Min, Max	1.1, 7.5	1.8, 7.8	4.9, 34.1	2.4, 10.2
	Mean (SD)	3.48 (2.099)	3.82 (2.015)	9.33 (8.824)	6.25 (2.678)

Note: NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: BLOQ values before T<sub>0</sub> are treated as zero. BLOQ values after the last quantifiable value are treated as missing. Any other BLOQ values are imputed by LLOQ/2.

Note: LLOQ = 0.2 ng/mL.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_pkconc3.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.6.2 Descriptive Statistics of the Plasma Nicotine Concentrations (ng/mL) by Nicotine Level - PK Population**

Timepoint	Statistic	Group-2 PK			
		THS 2.2 Menthol (N=18)		NRT gum (N=18)	
		<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)	<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)
T <sub>0</sub> + 2 h	n	10	8	10	8
	BLOQ - n (%)	0	0	0	0
	Geometric Mean (CV%)	2.04 (79.27%)	2.22 (49.49%)	4.93 (38.33%)	3.74 (63.57%)
	95% CI	1.23, 3.37	1.50, 3.29	3.78, 6.44	2.29, 6.09
	Median	2.12	2.14	4.56	3.74
	Q25, Q75	1.37, 3.58	1.51, 3.04	3.96, 6.58	2.44, 5.94
	Min, Max	0.7, 5.3	1.3, 5.0	2.9, 8.9	1.6, 8.9
	Mean (SD)	2.50 (1.590)	2.46 (1.236)	5.26 (2.060)	4.34 (2.578)

Note: NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: BLOQ values before T<sub>0</sub> are treated as zero. BLOQ values after the last quantifiable value are treated as missing. Any other BLOQ values are imputed by LLOQ/2.

Note: LLOQ = 0.2 ng/mL.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_pkconc3.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.6.2 Descriptive Statistics of the Plasma Nicotine Concentrations (ng/mL) by Nicotine Level - PK Population**

Timepoint	Statistic	Group-2 PK			
		THS 2.2 Menthol (N=18)		NRT gum (N=18)	
		<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)	<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)
T <sub>0</sub> + 3 h	n	0	0	10	8
	BLOQ - n (%)			0	0
	Geometric Mean (CV%)			3.57 (53.19%)	2.61 (68.02%)
	95% CI			2.49, 5.11	1.55, 4.38
	Median			3.58	2.35
	Q25, Q75			2.70, 5.39	1.88, 4.29
	Min, Max			1.6, 7.3	1.1, 6.7
	Mean (SD)			3.98 (1.945)	3.10 (2.029)

Note: NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: BLOQ values before T<sub>0</sub> are treated as zero. BLOQ values after the last quantifiable value are treated as missing. Any other BLOQ values are imputed by LLOQ/2.

Note: LLOQ = 0.2 ng/mL.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_pkconc3.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.6.2 Descriptive Statistics of the Plasma Nicotine Concentrations (ng/mL) by Nicotine Level - PK Population**

Timepoint	Statistic	Group-2 PK			
		THS 2.2 Menthol (N=18)		NRT gum (N=18)	
		<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)	<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)
T <sub>0</sub> + 4 h	n	10	8	10	8
	BLOQ - n (%)	0	0	0	0
	Geometric Mean (CV%)	1.15 (81.98%)	1.25 (49.86%)	2.75 (71.10%)	1.89 (68.35%)
	95% CI	0.69, 1.93	0.84, 1.86	1.73, 4.35	1.12, 3.18
	Median	1.01	1.15	2.87	1.64
	Q25, Q75	0.64, 2.00	0.86, 1.65	1.59, 4.90	1.33, 3.12
	Min, Max	0.4, 3.3	0.7, 3.0	1.1, 6.4	0.9, 5.1
	Mean (SD)	1.45 (1.062)	1.39 (0.757)	3.27 (1.964)	2.27 (1.612)

Note: NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: BLOQ values before T<sub>0</sub> are treated as zero. BLOQ values after the last quantifiable value are treated as missing. Any other BLOQ values are imputed by LLOQ/2.

Note: LLOQ = 0.2 ng/mL.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_pkconc3.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.6.2 Descriptive Statistics of the Plasma Nicotine Concentrations (ng/mL) by Nicotine Level - PK Population**

Timepoint	Statistic	Group-2 PK			
		THS 2.2 Menthol (N=18)		NRT gum (N=18)	
		<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)	<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)
T <sub>0</sub> + 6 h	n	9	8	10	8
	BLOQ - n (%)	1 (10.0%)	0	0	0
	Geometric Mean (CV%)	0.81 (85.84%)	0.71 (55.93%)	1.61 (95.58%)	1.04 (89.16%)
	95% CI	0.45, 1.44	0.46, 1.11	0.90, 2.88	0.55, 1.98
	Median	0.83	0.66	1.70	0.93
	Q25, Q75	0.45, 1.35	0.48, 0.92	0.79, 3.66	0.67, 1.93
	Min, Max	0.3, 2.6	0.4, 2.0	0.4, 4.4	0.4, 3.5
	Mean (SD)	1.04 (0.793)	0.82 (0.527)	2.10 (1.474)	1.37 (1.153)

Note: NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: BLOQ values before T<sub>0</sub> are treated as zero. BLOQ values after the last quantifiable value are treated as missing. Any other BLOQ values are imputed by LLOQ/2.

Note: LLOQ = 0.2 ng/mL.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_pkconc3.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.6.2 Descriptive Statistics of the Plasma Nicotine Concentrations (ng/mL) by Nicotine Level - PK Population**

Timepoint	Statistic	Group-2 PK			
		THS 2.2 Menthol (N=18)		NRT gum (N=18)	
		<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)	<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)
T <sub>0</sub> + 9 h	n	7	8	9	7
	BLOQ - n (%)	3 (30.0%)	0	1 (10.0%)	1 (12.5%)
	Geometric Mean (CV%)	0.61 (72.52%)	0.45 (62.95%)	1.06 (106.21%)	0.65 (90.68%)
	95% CI	0.33, 1.12	0.27, 0.73	0.54, 2.08	0.31, 1.35
	Median	0.49	0.48	0.93	0.45
	Q25, Q75	0.37, 1.01	0.27, 0.60	0.84, 2.77	0.38, 1.46
	Min, Max	0.3, 1.9	0.2, 1.2	0.3, 2.9	0.3, 2.6
	Mean (SD)	0.75 (0.550)	0.52 (0.320)	1.43 (1.068)	0.88 (0.847)

Note: NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: BLOQ values before T<sub>0</sub> are treated as zero. BLOQ values after the last quantifiable value are treated as missing. Any other BLOQ values are imputed by LLOQ/2.

Note: LLOQ = 0.2 ng/mL.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_pkconc3.sas

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Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

**Table 15.2.4.6.2 Descriptive Statistics of the Plasma Nicotine Concentrations (ng/mL) by Nicotine Level - PK Population**

Timepoint	Statistic	Group-2 PK			
		THS 2.2 Menthol (N=18)		NRT gum (N=18)	
		<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)	<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)
T <sub>0</sub> + 12 h	n	7	5	8	7
	BLOQ - n (%)	3 (30.0%)	3 (37.5%)	2 (20.0%)	1 (12.5%)
	Geometric Mean (CV%)	0.41 (65.61%)	0.36 (50.42%)	0.76 (91.92%)	0.42 (100.99%)
	95% CI	0.23, 0.72	0.20, 0.66	0.39, 1.46	0.19, 0.91
	Median	0.33	0.29	0.67	0.24
	Q25, Q75	0.26, 0.56	0.27, 0.34	0.48, 1.64	0.23, 0.95
	Min, Max	0.2, 1.3	0.3, 0.8	0.2, 1.9	0.2, 1.8
	Mean (SD)	0.49 (0.363)	0.40 (0.245)	0.97 (0.666)	0.59 (0.602)

Note: NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: BLOQ values before T<sub>0</sub> are treated as zero. BLOQ values after the last quantifiable value are treated as missing. Any other BLOQ values are imputed by LLOQ/2.

Note: LLOQ = 0.2 ng/mL.

Appendix 15.3.3.2

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_pkconc3.sas

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Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

**Table 15.2.4.6.2 Descriptive Statistics of the Plasma Nicotine Concentrations (ng/mL) by Nicotine Level - PK Population**

Timepoint	Statistic	Group-2 PK			
		THS 2.2 Menthol (N=18)		NRT gum (N=18)	
		<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)	<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)
T <sub>0</sub> + 24 h	n	3	3	6	2
	BLOQ - n (%)	7 (70.0%)	5 (62.5%)	4 (40.0%)	6 (75.0%)
	Geometric Mean (CV%)	NC	NC	0.43 (43.71%)	NC
	95% CI	NC	NC	0.27, 0.67	NC
	Median	NC	NC	0.47	NC
	Q25, Q75	NC	NC	0.34, 0.61	NC
	Min, Max	NC	NC	0.2, 0.6	NC
	Mean (SD)	NC	NC	0.46 (0.160)	NC

Note: NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Note: NC = Not calculated.

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: BLOQ values before T<sub>0</sub> are treated as zero. BLOQ values after the last quantifiable value are treated as missing. Any other BLOQ values are imputed by LLOQ/2.

Note: LLOQ = 0.2 ng/mL.

Appendix 15.3.3.2

Path: /cvm/projects/prj/development/000000106326/dev/tables/t\_pkconc3.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.7 Analysis of Blood COHb (%) - PK Population**

Variable	Time point	Statistic	Group-1 PK		
			THS 2.2 Menthol (N=43)	mCC (N=43)	THS 2.2 Menthol: mCC Ratio (%)
Blood COHb (%)	Overall	n	43	43	
		Geometric LS Mean (CV%)	2.54	3.12	81.47 (12.68)
		95% CI	2.47, 2.61	3.03, 3.21	78.95, 84.07
	T <sub>0</sub> + 15 min	n	43	43	
		Geometric LS Mean (CV%)	2.55	3.53	72.20 (14.54)
		95% CI	2.46, 2.64	3.41, 3.65	69.29, 75.24
	T <sub>0</sub> + 60 min	n	43	43	
		Geometric LS Mean (CV%)	2.61	3.32	78.51 (14.38)
		95% CI	2.52, 2.70	3.21, 3.43	75.41, 81.74

Note: mCC = menthol conventional cigarettes; THS = Tobacco Heating System.

Note: Geometric LS Mean and 95% CI are the adjusted geometric least squares means based on an ANOVA model. Geometrical CV% of the ratio is estimated only for the ratio.

Note: T<sub>0</sub> = Time of first product-use at single use day

Note: Comparison overall time points is the main comparison.

Appendix 15.4.4.7 and 15.3.3.4

Path: /cvn/projects/prj/development/000000106326/dev/tables/tlf\_anlcohb.sas

Program Run: 28OCT14 cvn\_ahedge Program Status: FINAL

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**Table 15.2.4.7 Analysis of Blood COHb (%) - PK Population**

Variable	Time point	Statistic	Group-1 PK		
			THS 2.2 Menthol (N=43)	mCC (N=43)	THS 2.2 Menthol: mCC Ratio (%)
Blood COHb (%)	T <sub>0</sub> + 4 h	n	43	43	
		Geometric LS Mean (CV%)	2.59	3.10	83.43 (14.13)
		95% CI	2.51, 2.68	3.00, 3.21	80.24, 86.74
	T <sub>0</sub> + 12 h	n	43	43	
		Geometric LS Mean (CV%)	2.42	2.60	93.16 (14.71)
		95% CI	2.34, 2.51	2.51, 2.69	89.32, 97.17

Note: mCC = menthol conventional cigarettes; THS = Tobacco Heating System.

Note: Geometric LS Mean and 95% CI are the adjusted geometric least squares means based on an ANOVA model. Geometrical CV% of the ratio is estimated only for the ratio.

Note: T<sub>0</sub> = Time of first product-use at single use day

Note: Comparison overall time points is the main comparison.

Appendix 15.4.4.7 and 15.3.3.4

Path: /cvn/projects/prj/development/000000106326/dev/tables/tlf\_anlcohb.sas

Program Run: 28OCT14 cvn\_ahedge Program Status: FINAL

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**Table 15.2.4.7 Analysis of Blood COHb (%) - PK Population**

Variable	Time point	Statistic	Group-2 PK		
			THS 2.2 Menthol (N=18)	NRT gum (N=18)	THS 2.2 Menthol: NRT gum Ratio (%)
Blood COHb (%)	Overall	n	18	18	
		Geometric LS Mean (CV%)	2.45	2.46	99.63 (13.61)
		95% CI	2.35, 2.55	2.36, 2.56	96.07, 103.32
	T <sub>0</sub> + 15 min	n	18	18	
		Geometric LS Mean (CV%)	2.42	2.48	97.32 (14.59)
		95% CI	2.31, 2.52	2.38, 2.59	93.34, 101.46
	T <sub>0</sub> + 60 min	n	18	18	
		Geometric LS Mean (CV%)	2.44	2.48	98.56 (19.63)
		95% CI	2.29, 2.60	2.33, 2.64	91.45, 106.23

Note: NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Note: Geometric LS Mean and 95% CI are the adjusted geometric least squares means based on an ANOVA model. Geometrical CV% of the ratio is estimated only for the ratio.

Note: T<sub>0</sub> = Time of first product-use at single use day

Note: Comparison overall time points is the main comparison.

Appendix 15.4.4.7 and 15.3.3.4

Path: /cvn/projects/prj/development/000000106326/dev/tables/tlf\_anlcohb.sas

Program Run: 28OCT14 cvn\_ahedge Program Status: FINAL

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**Table 15.2.4.7 Analysis of Blood COHb (%) - PK Population**

Variable	Time point	Statistic	Group-2 PK		
			THS 2.2 Menthol (N=18)	NRT gum (N=18)	THS 2.2 Menthol: NRT gum Ratio (%)
Blood COHb (%)	T <sub>0</sub> + 4 h	n	18	18	
		Geometric LS Mean (CV%)	2.49	2.50	99.68 (15.80)
		95% CI	2.38, 2.62	2.39, 2.62	94.92, 104.67
	T <sub>0</sub> + 12 h	n	18	18	
		Geometric LS Mean (CV%)	2.43	2.36	103.04 (17.64)
		95% CI	2.30, 2.57	2.24, 2.49	96.97, 109.48

Note: NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Note: Geometric LS Mean and 95% CI are the adjusted geometric least squares means based on an ANOVA model. Geometrical CV% of the ratio is estimated only for the ratio.

Note: T<sub>0</sub> = Time of first product-use at single use day

Note: Comparison overall time points is the main comparison.

Appendix 15.4.4.7 and 15.3.3.4

Path: /cvn/projects/prj/development/000000106326/dev/tables/tlf\_anlcohb.sas

Program Run: 28OCT14 cvn\_ahedge Program Status: FINAL

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**Table 15.2.4.8.1 Descriptive Statistics of Blood COHb (%) Continuous Measurements - PK Population**

Timepoint	Statistic	Group-1 PK		Group-2 PK	
		THS 2.2 Menthol (N=43)	mCC (N=43)	THS 2.2 Menthol (N=18)	NRT gum (N=18)
15 min < T <sub>0</sub>	n	43	43	18	18
	Geometric Mean (CV%)	2.50 (9.83%)	2.46 (8.50%)	2.38 (11.16%)	2.41 (9.53%)
	95% CI	2.42, 2.58	2.39, 2.53	2.24, 2.52	2.29, 2.53
	Median	2.50	2.40	2.45	2.40
	Q25, Q75	2.30, 2.70	2.30, 2.60	2.20, 2.60	2.30, 2.50
	Min, Max	2.1, 3.1	2.2, 3.1	2.0, 2.8	2.0, 3.0
	Mean (SD)	2.51 (0.248)	2.47 (0.217)	2.39 (0.261)	2.42 (0.236)

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Note: T<sub>0</sub> = Time of first product use at single use day.

Appendix 15.3.3.4

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_cohb.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.8.1 Descriptive Statistics of Blood COHb (%) Continuous Measurements - PK Population**

Timepoint	Statistic	Group-1 PK		Group-2 PK	
		THS 2.2 Menthol (N=43)	mCC (N=43)	THS 2.2 Menthol (N=18)	NRT gum (N=18)
T <sub>0</sub> + 15 min	n	43	43	18	18
	Geometric Mean (CV%)	2.55 (9.67%)	3.53 (12.56%)	2.42 (7.59%)	2.48 (10.13%)
	95% CI	2.47, 2.63	3.39, 3.67	2.32, 2.51	2.36, 2.62
	Median	2.50	3.50	2.40	2.55
	Q25, Q75	2.40, 2.70	3.20, 3.90	2.40, 2.50	2.30, 2.60
	Min, Max	2.1, 3.3	2.8, 4.8	2.0, 2.8	2.1, 3.0
	Mean (SD)	2.56 (0.255)	3.56 (0.458)	2.42 (0.181)	2.49 (0.256)

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Note: T<sub>0</sub> = Time of first product use at single use day.

Appendix 15.3.3.4

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_cohb.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.8.1 Descriptive Statistics of Blood COHb (%) Continuous Measurements - PK Population**

Timepoint	Statistic	Group-1 PK		Group-2 PK	
		THS 2.2 Menthol (N=43)	mCC (N=43)	THS 2.2 Menthol (N=18)	NRT gum (N=18)
T <sub>0</sub> + 60 min	n	43	43	18	18
	Geometric Mean (CV%)	2.61 (8.95%)	3.32 (11.45%)	2.44 (7.72%)	2.48 (12.97%)
	95% CI	2.53, 2.69	3.20, 3.44	2.35, 2.54	2.32, 2.65
	Median	2.60	3.30	2.40	2.50
	Q25, Q75	2.40, 2.70	3.10, 3.50	2.30, 2.60	2.30, 2.50
	Min, Max	2.3, 3.5	2.6, 4.7	2.1, 2.8	2.1, 3.8
	Mean (SD)	2.62 (0.247)	3.34 (0.392)	2.45 (0.189)	2.50 (0.368)

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Note: T<sub>0</sub> = Time of first product use at single use day.

Appendix 15.3.3.4

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_cohb.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.8.1 Descriptive Statistics of Blood COHb (%) Continuous Measurements - PK Population**

Timepoint	Statistic	Group-1 PK		Group-2 PK	
		THS 2.2 Menthol (N=43)	mCC (N=43)	THS 2.2 Menthol (N=18)	NRT gum (N=18)
T <sub>0</sub> + 4 h	n	43	43	18	18
	Geometric Mean (CV%)	2.59 (10.74%)	3.10 (10.02%)	2.49 (9.60%)	2.50 (10.87%)
	95% CI	2.50, 2.68	3.00, 3.21	2.37, 2.62	2.37, 2.65
	Median	2.60	3.10	2.50	2.50
	Q25, Q75	2.30, 2.80	2.90, 3.30	2.40, 2.60	2.40, 2.70
	Min, Max	2.2, 3.2	2.6, 3.9	2.0, 3.0	2.0, 3.0
	Mean (SD)	2.60 (0.281)	3.12 (0.319)	2.51 (0.237)	2.52 (0.269)

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Note: T<sub>0</sub> = Time of first product use at single use day.

Appendix 15.3.3.4

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_cohb.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.8.1 Descriptive Statistics of Blood COHb (%) Continuous Measurements - PK Population**

Timepoint	Statistic	Group-1 PK		Group-2 PK	
		THS 2.2 Menthol (N=43)	mCC (N=43)	THS 2.2 Menthol (N=18)	NRT gum (N=18)
T <sub>0</sub> + 12 h	n	43	43	18	18
	Geometric Mean (CV%)	2.42 (11.23%)	2.60 (10.69%)	2.43 (13.97%)	2.36 (12.35%)
	95% CI	2.34, 2.51	2.51, 2.69	2.26, 2.61	2.22, 2.51
	Median	2.40	2.60	2.45	2.30
	Q25, Q75	2.30, 2.60	2.40, 2.70	2.20, 2.60	2.20, 2.50
	Min, Max	2.0, 3.0	2.1, 3.4	1.9, 3.6	2.0, 3.1
	Mean (SD)	2.44 (0.275)	2.62 (0.289)	2.46 (0.365)	2.38 (0.301)

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Note: T<sub>0</sub> = Time of first product use at single use day.

Appendix 15.3.3.4

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_cohb.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.8.1.1 Descriptive Statistics of Blood COHb (%) by Sex - PK Population**

Timepoint	Statistic	Group-1 PK			
		THS 2.2 Menthol (N=43)		mCC (N=43)	
		Males (N=23)	Females (N=20)	Males (N=23)	Females (N=20)
15 min < T <sub>0</sub>	n	23	20	23	20
	Geometric Mean (CV%)	2.47 (9.20%)	2.52 (10.66%)	2.50 (7.86%)	2.41 (9.02%)
	95% CI	2.37, 2.58	2.39, 2.65	2.41, 2.59	2.31, 2.52
	Median	2.40	2.55	2.50	2.35
	Q25, Q75	2.30, 2.70	2.30, 2.75	2.40, 2.60	2.30, 2.45
	Min, Max	2.1, 2.9	2.1, 3.1	2.2, 3.1	2.2, 3.0
	Mean (SD)	2.48 (0.229)	2.54 (0.271)	2.50 (0.204)	2.42 (0.229)

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Note: T<sub>0</sub> = Time of first product use at single use day.

Appendix 15.3.3.4

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_cohb2.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.8.1.1 Descriptive Statistics of Blood COHb (%) by Sex - PK Population**

Timepoint	Statistic	Group-1 PK			
		THS 2.2 Menthol (N=43)		mCC (N=43)	
		Males (N=23)	Females (N=20)	Males (N=23)	Females (N=20)
T <sub>0</sub> + 15 min	n	23	20	23	20
	Geometric Mean (CV%)	2.54 (7.57%)	2.56 (11.84%)	3.47 (13.46%)	3.60 (11.47%)
	95% CI	2.45, 2.63	2.42, 2.71	3.27, 3.68	3.41, 3.80
	Median	2.50	2.50	3.40	3.60
	Q25, Q75	2.40, 2.70	2.35, 2.70	3.20, 3.70	3.30, 3.90
	Min, Max	2.2, 2.9	2.1, 3.3	2.9, 4.8	2.8, 4.4
	Mean (SD)	2.55 (0.193)	2.58 (0.317)	3.50 (0.499)	3.62 (0.409)

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Note: T<sub>0</sub> = Time of first product use at single use day.

Appendix 15.3.3.4

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_cohb2.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.8.1.1 Descriptive Statistics of Blood COHb (%) by Sex - PK Population**

Timepoint	Statistic	Group-1 PK			
		THS 2.2 Menthol (N=43)		mCC (N=43)	
		Males (N=23)	Females (N=20)	Males (N=23)	Females (N=20)
T <sub>0</sub> + 60 min	n	23	20	23	20
	Geometric Mean (CV%)	2.62 (10.72%)	2.60 (6.61%)	3.28 (12.35%)	3.37 (10.42%)
	95% CI	2.49, 2.74	2.52, 2.69	3.10, 3.46	3.21, 3.55
	Median	2.50	2.60	3.20	3.40
	Q25, Q75	2.40, 2.80	2.45, 2.70	3.10, 3.50	3.05, 3.70
	Min, Max	2.3, 3.5	2.4, 3.1	2.6, 4.7	2.8, 4.0
	Mean (SD)	2.63 (0.298)	2.61 (0.177)	3.30 (0.428)	3.39 (0.350)

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Note: T<sub>0</sub> = Time of first product use at single use day.

Appendix 15.3.3.4

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_cohb2.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.8.1.1 Descriptive Statistics of Blood COHb (%) by Sex - PK Population**

Timepoint	Statistic	Group-1 PK			
		THS 2.2 Menthol (N=43)		mCC (N=43)	
		Males (N=23)	Females (N=20)	Males (N=23)	Females (N=20)
T <sub>0</sub> + 4 h	n	23	20	23	20
	Geometric Mean (CV%)	2.54 (10.03%)	2.64 (11.41%)	3.09 (11.86%)	3.12 (7.66%)
	95% CI	2.43, 2.66	2.50, 2.79	2.93, 3.25	3.01, 3.24
	Median	2.60	2.75	3.00	3.20
	Q25, Q75	2.30, 2.70	2.35, 2.85	2.80, 3.30	2.90, 3.35
	Min, Max	2.2, 3.2	2.2, 3.2	2.6, 3.9	2.7, 3.5
	Mean (SD)	2.56 (0.260)	2.66 (0.300)	3.11 (0.382)	3.13 (0.237)

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Note: T<sub>0</sub> = Time of first product use at single use day.

Appendix 15.3.3.4

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_cohb2.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.8.1.1 Descriptive Statistics of Blood COHb (%) by Sex - PK Population**

Timepoint	Statistic	Group-1 PK			
		THS 2.2 Menthol (N=43)		mCC (N=43)	
		Males (N=23)	Females (N=20)	Males (N=23)	Females (N=20)
T <sub>0</sub> + 12 h	n	23	20	23	20
	Geometric Mean (CV%)	2.42 (10.75%)	2.43 (12.03%)	2.59 (11.64%)	2.62 (9.75%)
	95% CI	2.30, 2.54	2.30, 2.58	2.46, 2.73	2.50, 2.74
	Median	2.40	2.40	2.60	2.60
	Q25, Q75	2.30, 2.60	2.25, 2.65	2.40, 2.80	2.45, 2.70
	Min, Max	2.0, 3.0	2.0, 3.0	2.1, 3.4	2.3, 3.4
	Mean (SD)	2.43 (0.264)	2.45 (0.293)	2.60 (0.312)	2.63 (0.268)

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Note: T<sub>0</sub> = Time of first product use at single use day.

Appendix 15.3.3.4

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_cohb2.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.8.1.1 Descriptive Statistics of Blood COHb (%) by Sex - PK Population**

Timepoint	Statistic	Group-2 PK			
		THS 2.2 Menthol (N=18)		NRT gum (N=18)	
		Males (N=9)	Females (N=9)	Males (N=9)	Females (N=9)
15 min < T <sub>0</sub>	n	9	9	9	9
	Geometric Mean (CV%)	2.36 (12.37%)	2.39 (10.54%)	2.42 (12.64%)	2.40 (5.79%)
	95% CI	2.14, 2.60	2.20, 2.59	2.19, 2.67	2.29, 2.51
	Median	2.50	2.40	2.40	2.40
	Q25, Q75	2.10, 2.60	2.20, 2.60	2.20, 2.50	2.30, 2.40
	Min, Max	2.0, 2.7	2.0, 2.8	2.0, 3.0	2.2, 2.7
	Mean (SD)	2.38 (0.287)	2.40 (0.250)	2.43 (0.313)	2.40 (0.142)

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Note: T<sub>0</sub> = Time of first product use at single use day.

Appendix 15.3.3.4

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_cohb2.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.8.1.1 Descriptive Statistics of Blood COHb (%) by Sex - PK Population**

Timepoint	Statistic	Group-2 PK			
		THS 2.2 Menthol (N=18)		NRT gum (N=18)	
		Males (N=9)	Females (N=9)	Males (N=9)	Females (N=9)
T <sub>0</sub> + 15 min	n	9	9	9	9
	Geometric Mean (CV%)	2.42 (5.03%)	2.41 (9.85%)	2.53 (12.80%)	2.44 (6.90%)
	95% CI	2.32, 2.52	2.23, 2.61	2.29, 2.79	2.31, 2.58
	Median	2.40	2.40	2.60	2.50
	Q25, Q75	2.40, 2.50	2.40, 2.60	2.30, 2.70	2.30, 2.60
	Min, Max	2.2, 2.6	2.0, 2.8	2.1, 3.0	2.2, 2.6
	Mean (SD)	2.42 (0.121)	2.42 (0.234)	2.54 (0.325)	2.44 (0.167)

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Note: T<sub>0</sub> = Time of first product use at single use day.

Appendix 15.3.3.4

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_cohb2.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.8.1.1 Descriptive Statistics of Blood COHb (%) by Sex - PK Population**

Timepoint	Statistic	Group-2 PK			
		THS 2.2 Menthol (N=18)		NRT gum (N=18)	
		Males (N=9)	Females (N=9)	Males (N=9)	Females (N=9)
T <sub>0</sub> + 60 min	n	9	9	9	9
	Geometric Mean (CV%)	2.45 (8.95%)	2.44 (6.83%)	2.40 (7.99%)	2.56 (16.55%)
	95% CI	2.28, 2.63	2.31, 2.58	2.26, 2.56	2.25, 2.90
	Median	2.40	2.40	2.40	2.50
	Q25, Q75	2.30, 2.60	2.30, 2.60	2.30, 2.50	2.40, 2.50
	Min, Max	2.1, 2.8	2.2, 2.7	2.1, 2.7	2.1, 3.8
	Mean (SD)	2.46 (0.219)	2.44 (0.167)	2.41 (0.191)	2.59 (0.484)

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Note: T<sub>0</sub> = Time of first product use at single use day.

Appendix 15.3.3.4

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_cohb2.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.8.1.1 Descriptive Statistics of Blood COHb (%) by Sex - PK Population**

Timepoint	Statistic	Group-2 PK			
		THS 2.2 Menthol (N=18)		NRT gum (N=18)	
		Males (N=9)	Females (N=9)	Males (N=9)	Females (N=9)
T <sub>0</sub> + 4 h	n	9	9	9	9
	Geometric Mean (CV%)	2.44 (10.18%)	2.55 (9.09%)	2.50 (11.18%)	2.51 (11.24%)
	95% CI	2.26, 2.65	2.37, 2.74	2.29, 2.73	2.30, 2.74
	Median	2.50	2.50	2.50	2.50
	Q25, Q75	2.40, 2.60	2.40, 2.60	2.40, 2.50	2.30, 2.70
	Min, Max	2.0, 2.8	2.2, 3.0	2.0, 3.0	2.1, 2.9
	Mean (SD)	2.46 (0.241)	2.56 (0.236)	2.51 (0.276)	2.52 (0.278)

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Note: T<sub>0</sub> = Time of first product use at single use day.

Appendix 15.3.3.4

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_cohb2.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.8.1.1 Descriptive Statistics of Blood COHb (%) by Sex - PK Population**

Timepoint	Statistic	Group-2 PK			
		THS 2.2 Menthol (N=18)		NRT gum (N=18)	
		Males (N=9)	Females (N=9)	Males (N=9)	Females (N=9)
T <sub>0</sub> + 12 h	n	9	9	9	9
	Geometric Mean (CV%)	2.41 (10.60%)	2.45 (17.38%)	2.40 (13.62%)	2.32 (11.47%)
	95% CI	2.22, 2.62	2.14, 2.81	2.16, 2.67	2.12, 2.54
	Median	2.50	2.40	2.50	2.20
	Q25, Q75	2.40, 2.60	2.20, 2.60	2.30, 2.50	2.20, 2.50
	Min, Max	1.9, 2.6	2.0, 3.6	2.0, 3.1	2.0, 2.8
	Mean (SD)	2.42 (0.239)	2.49 (0.473)	2.42 (0.335)	2.33 (0.274)

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Note: T<sub>0</sub> = Time of first product use at single use day.

Appendix 15.3.3.4

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_cohb2.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.8.1.2 Descriptive Statistics of Blood COHb (%) by Nicotine Level - PK Population**

Timepoint	Statistic	Group-1 PK			
		THS 2.2 Menthol (N=43)		mCC (N=43)	
		<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)	<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)
15 min < T <sub>0</sub>	n	25	18	25	18
	Geometric Mean (CV%)	2.48 (10.81%)	2.51 (8.53%)	2.43 (8.93%)	2.50 (7.80%)
	95% CI	2.37, 2.60	2.40, 2.63	2.33, 2.52	2.40, 2.60
	Median	2.50	2.45	2.40	2.50
	Q25, Q75	2.30, 2.60	2.30, 2.70	2.30, 2.50	2.40, 2.60
	Min, Max	2.1, 3.1	2.2, 2.9	2.2, 3.1	2.2, 2.9
	Mean (SD)	2.50 (0.272)	2.52 (0.216)	2.44 (0.231)	2.51 (0.196)

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Note: T<sub>0</sub> = Time of first product use at single use day.

Appendix 15.3.3.4

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_cohb3.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.8.1.2 Descriptive Statistics of Blood COHb (%) by Nicotine Level - PK Population**

Timepoint	Statistic	Group-1 PK			
		THS 2.2 Menthol (N=43)		mCC (N=43)	
		<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)	<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)
T <sub>0</sub> + 15 min	n	25	18	25	18
	Geometric Mean (CV%)	2.54 (10.63%)	2.56 (8.42%)	3.72 (11.25%)	3.28 (10.50%)
	95% CI	2.42, 2.66	2.45, 2.68	3.55, 3.90	3.11, 3.46
	Median	2.50	2.55	3.70	3.25
	Q25, Q75	2.40, 2.70	2.40, 2.70	3.40, 4.00	3.10, 3.50
	Min, Max	2.1, 3.3	2.3, 3.2	3.1, 4.8	2.8, 4.4
	Mean (SD)	2.55 (0.279)	2.57 (0.225)	3.74 (0.430)	3.29 (0.364)

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Note: T<sub>0</sub> = Time of first product use at single use day.

Appendix 15.3.3.4

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_cohb3.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.8.1.2 Descriptive Statistics of Blood COHb (%) by Nicotine Level - PK Population**

Timepoint	Statistic	Group-1 PK			
		THS 2.2 Menthol (N=43)		mCC (N=43)	
		<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)	<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)
T <sub>0</sub> + 60 min	n	25	18	25	18
	Geometric Mean (CV%)	2.57 (8.45%)	2.65 (9.54%)	3.46 (10.75%)	3.13 (9.84%)
	95% CI	2.48, 2.67	2.53, 2.79	3.31, 3.63	2.98, 3.29
	Median	2.50	2.60	3.40	3.10
	Q25, Q75	2.40, 2.70	2.50, 2.80	3.30, 3.70	3.00, 3.40
	Min, Max	2.3, 3.2	2.4, 3.5	2.8, 4.7	2.6, 3.8
	Mean (SD)	2.58 (0.227)	2.67 (0.271)	3.48 (0.387)	3.14 (0.310)

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Note: T<sub>0</sub> = Time of first product use at single use day.

Appendix 15.3.3.4

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_cohb3.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.8.1.2 Descriptive Statistics of Blood COHb (%) by Nicotine Level - PK Population**

Timepoint	Statistic	Group-1 PK			
		THS 2.2 Menthol (N=43)		mCC (N=43)	
		<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)	<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)
T <sub>0</sub> + 4 h	n	25	18	25	18
	Geometric Mean (CV%)	2.59 (12.04%)	2.58 (8.96%)	3.20 (10.47%)	2.97 (7.61%)
	95% CI	2.46, 2.73	2.47, 2.71	3.06, 3.35	2.85, 3.09
	Median	2.70	2.60	3.20	2.95
	Q25, Q75	2.30, 2.80	2.40, 2.70	3.00, 3.40	2.80, 3.10
	Min, Max	2.2, 3.2	2.3, 3.2	2.7, 3.9	2.6, 3.4
	Mean (SD)	2.61 (0.312)	2.59 (0.239)	3.22 (0.341)	2.98 (0.227)

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Note: T<sub>0</sub> = Time of first product use at single use day.

Appendix 15.3.3.4

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_cohb3.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.8.1.2 Descriptive Statistics of Blood COHb (%) by Nicotine Level - PK Population**

Timepoint	Statistic	Group-1 PK			
		THS 2.2 Menthol (N=43)		mCC (N=43)	
		<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)	<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)
T <sub>0</sub> + 12 h	n	25	18	25	18
	Geometric Mean (CV%)	2.40 (11.44%)	2.46 (11.10%)	2.67 (12.22%)	2.51 (6.98%)
	95% CI	2.29, 2.52	2.32, 2.60	2.53, 2.81	2.42, 2.60
	Median	2.40	2.45	2.70	2.55
	Q25, Q75	2.30, 2.60	2.30, 2.70	2.50, 2.80	2.40, 2.60
	Min, Max	2.0, 3.0	2.1, 3.0	2.1, 3.4	2.3, 3.0
	Mean (SD)	2.42 (0.277)	2.47 (0.277)	2.69 (0.333)	2.52 (0.180)

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Note: T<sub>0</sub> = Time of first product use at single use day.

Appendix 15.3.3.4

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_cohb3.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.8.1.2 Descriptive Statistics of Blood COHb (%) by Nicotine Level - PK Population**

Timepoint	Statistic	Group-2 PK			
		THS 2.2 Menthol (N=18)		NRT gum (N=18)	
		<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)	<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)
15 min < T <sub>0</sub>	n	10	8	10	8
	Geometric Mean (CV%)	2.42 (11.02%)	2.32 (11.67%)	2.44 (7.94%)	2.36 (11.51%)
	95% CI	2.23, 2.62	2.10, 2.57	2.30, 2.59	2.14, 2.60
	Median	2.55	2.35	2.45	2.35
	Q25, Q75	2.20, 2.60	2.10, 2.50	2.30, 2.50	2.25, 2.40
	Min, Max	2.0, 2.7	2.0, 2.8	2.2, 2.8	2.0, 3.0
	Mean (SD)	2.43 (0.259)	2.34 (0.273)	2.45 (0.196)	2.38 (0.287)

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Note: T<sub>0</sub> = Time of first product use at single use day.

Appendix 15.3.3.4

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_cohb3.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.8.1.2 Descriptive Statistics of Blood COHb (%) by Nicotine Level - PK Population**

Timepoint	Statistic	Group-2 PK			
		THS 2.2 Menthol (N=18)		NRT gum (N=18)	
		<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)	<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)
T <sub>0</sub> + 15 min	n	10	8	10	8
	Geometric Mean (CV%)	2.40 (5.62%)	2.44 (9.86%)	2.48 (9.48%)	2.49 (11.56%)
	95% CI	2.30, 2.50	2.24, 2.65	2.31, 2.66	2.25, 2.74
	Median	2.40	2.40	2.45	2.60
	Q25, Q75	2.30, 2.50	2.40, 2.60	2.30, 2.60	2.25, 2.60
	Min, Max	2.2, 2.6	2.0, 2.8	2.2, 3.0	2.1, 3.0
	Mean (SD)	2.40 (0.134)	2.45 (0.233)	2.49 (0.243)	2.50 (0.288)

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Note: T<sub>0</sub> = Time of first product use at single use day.

Appendix 15.3.3.4

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_cohb3.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.8.1.2 Descriptive Statistics of Blood COHb (%) by Nicotine Level - PK Population**

Timepoint	Statistic	Group-2 PK			
		THS 2.2 Menthol (N=18)		NRT gum (N=18)	
		<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)	<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)
T <sub>0</sub> + 60 min	n	10	8	10	8
	Geometric Mean (CV%)	2.48 (6.15%)	2.40 (9.53%)	2.52 (16.31%)	2.43 (7.76%)
	95% CI	2.36, 2.59	2.21, 2.61	2.24, 2.83	2.27, 2.60
	Median	2.45	2.35	2.50	2.45
	Q25, Q75	2.40, 2.60	2.25, 2.65	2.30, 2.50	2.35, 2.55
	Min, Max	2.3, 2.8	2.1, 2.7	2.1, 3.8	2.1, 2.7
	Mean (SD)	2.48 (0.155)	2.41 (0.230)	2.55 (0.472)	2.44 (0.185)

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Note: T<sub>0</sub> = Time of first product use at single use day.

Appendix 15.3.3.4

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_cohb3.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.8.1.2 Descriptive Statistics of Blood COHb (%) by Nicotine Level - PK Population**

Timepoint	Statistic	Group-2 PK			
		THS 2.2 Menthol (N=18)		NRT gum (N=18)	
		<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)	<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)
T <sub>0</sub> + 4 h	n	10	8	10	8
	Geometric Mean (CV%)	2.51 (10.16%)	2.48 (9.48%)	2.50 (9.54%)	2.51 (13.05%)
	95% CI	2.33, 2.70	2.28, 2.69	2.33, 2.68	2.24, 2.80
	Median	2.55	2.45	2.50	2.50
	Q25, Q75	2.40, 2.60	2.30, 2.70	2.40, 2.70	2.35, 2.75
	Min, Max	2.0, 3.0	2.2, 2.8	2.1, 2.9	2.0, 3.0
	Mean (SD)	2.52 (0.249)	2.49 (0.236)	2.51 (0.238)	2.53 (0.320)

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Note: T<sub>0</sub> = Time of first product use at single use day.

Appendix 15.3.3.4

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_cohb3.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.8.1.2 Descriptive Statistics of Blood COHb (%) by Nicotine Level - PK Population**

Timepoint	Statistic	Group-2 PK			
		THS 2.2 Menthol (N=18)		NRT gum (N=18)	
		<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)	<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)
T <sub>0</sub> + 12 h	n	10	8	10	8
	Geometric Mean (CV%)	2.41 (10.89%)	2.46 (17.88%)	2.35 (9.96%)	2.37 (15.58%)
	95% CI	2.22, 2.61	2.12, 2.86	2.18, 2.53	2.08, 2.71
	Median	2.50	2.40	2.40	2.30
	Q25, Q75	2.20, 2.60	2.20, 2.60	2.20, 2.50	2.10, 2.65
	Min, Max	1.9, 2.7	2.0, 3.6	2.0, 2.7	2.0, 3.1
	Mean (SD)	2.42 (0.249)	2.50 (0.490)	2.36 (0.232)	2.40 (0.386)

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Note: T<sub>0</sub> = Time of first product use at single use day.

Appendix 15.3.3.4

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_cohb3.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.8.2 Descriptive Statistics of Blood COHb (%) Categorical Measurements - PK Population**

Timepoint	Statistic	Group-1 PK		Group-2 PK	
		THS 2.2 Menthol (N=43)	mCC (N=43)	THS 2.2 Menthol (N=18)	NRT gum (N=18)
15 min < T <sub>0</sub>	COHb <= 2% - n (%)	0	0	3 ( 16.7%)	1 ( 5.6%)
	COHb > 2% - n (%)	43 (100 %)	43 (100 %)	15 ( 83.3%)	17 ( 94.4%)
T <sub>0</sub> + 15 min	COHb <= 2% - n (%)	0	0	1 ( 5.6%)	0
	COHb > 2% - n (%)	43 (100 %)	43 (100 %)	17 ( 94.4%)	18 (100 %)
T <sub>0</sub> + 60 min	COHb <= 2% - n (%)	0	0	0	0
	COHb > 2% - n (%)	43 (100 %)	43 (100 %)	18 (100 %)	18 (100 %)
T <sub>0</sub> + 4 h	COHb <= 2% - n (%)	0	0	1 ( 5.6%)	1 ( 5.6%)
	COHb > 2% - n (%)	43 (100 %)	43 (100 %)	17 ( 94.4%)	17 ( 94.4%)
T <sub>0</sub> + 12 h	COHb <= 2% - n (%)	3 ( 7.0%)	0	2 ( 11.1%)	3 ( 16.7%)
	COHb > 2% - n (%)	40 ( 93.0%)	43 (100 %)	16 ( 88.9%)	15 ( 83.3%)

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Note: T<sub>0</sub> = Time of first product use at single use day.

Appendix 15.3.3.4

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_cohbcat.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.9 Analysis of Exhaled CO (ppm) During Single Use Day - PK Population**

Variable	Time point	Statistic	Group-1 PK		
			THS 2.2 Menthol (N=43)	mCC (N=43)	THS 2.2 Menthol - mCC
Exhaled CO (ppm)	Overall	n	43	43	
		LS Mean (SE)	3.43	5.05	-1.62 (0.153)
		95% CI	3.08, 3.78	4.70, 5.40	-1.92, -1.31
	12:00-01:30 PM	n	43	43	
		LS Mean (SE)	3.19	5.78	-2.60 (0.240)
		95% CI	2.75, 3.63	5.35, 6.22	-3.07, -2.12
	04:00-05:30 PM	n	43	43	
		LS Mean (SE)	3.68	5.06	-1.39 (0.255)
		95% CI	3.22, 4.13	4.61, 5.52	-1.89, -0.88
	08:00-09:30 PM	n	43	43	
		LS Mean (SE)	3.42	4.30	-0.87 (0.235)
		95% CI	2.99, 3.85	3.86, 4.73	-1.34, -0.41

Note: mCC = menthol conventional cigarettes; THS = Tobacco Heating System.

Note: Means and 95% CI are the adjusted least squares means and confidence intervals from an ANOVA model.

Note: Comparison overall time points is the main comparison.

Appendix 15.4.4.9 and 15.3.3.5

Path: /cvn/projects/prj/development/000000106326/dev/tables/tlf\_anlco.sas

Program Run: 28OCT14 cvn\_ahedge Program Status: FINAL

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**Table 15.2.4.9 Analysis of Exhaled CO (ppm) During Single Use Day - PK Population**

Variable	Time point	Statistic	Group-2 PK		
			THS 2.2 Menthol (N=18)	NRT gum (N=18)	THS 2.2 Menthol - NRT gum
Exhaled CO (ppm)	Overall	n	18	18	
		LS Mean (SE)	3.20	2.94	0.26 (0.176)
		95% CI	2.84, 3.57	2.58, 3.31	-0.09, 0.61
	12:00-01:30 PM	n	18	18	
		LS Mean (SE)	3.00	2.72	0.28 (0.219)
		95% CI	2.59, 3.41	2.31, 3.13	-0.16, 0.72
	04:00-05:30 PM	n	18	18	
		LS Mean (SE)	2.94	2.83	0.11 (0.241)
		95% CI	2.51, 3.38	2.40, 3.26	-0.37, 0.59
	08:00-09:30 PM	n	18	18	
		LS Mean (SE)	3.67	3.28	0.39 (0.406)
		95% CI	3.04, 4.30	2.65, 3.91	-0.42, 1.20

Note: NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Note: Means and 95% CI are the adjusted least squares means and confidence intervals from an ANOVA model.

Note: Comparison overall time points is the main comparison.

Appendix 15.4.4.9 and 15.3.3.5

Path: /cvn/projects/prj/development/000000106326/dev/tables/tlf\_anlco.sas

Program Run: 28OCT14 cvn\_ahedge Program Status: FINAL

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**Table 15.2.4.10.1 Descriptive Statistics of Exhaled CO (ppm) During Single Use Continuous Measurements - PK Population**

Timepoint	Statistic	Group-1 PK		Group-2 PK	
		THS 2.2 Menthol (N=43)	mCC (N=43)	THS 2.2 Menthol (N=18)	NRT gum (N=18)
15 min < T <sub>0</sub>	n	43	43	18	18
	Mean (SD)	3.5 (1.23)	3.7 (1.08)	3.7 (1.02)	3.7 (1.08)
	95% CI	3.1, 3.9	3.4, 4.1	3.2, 4.3	3.1, 4.3
	Median	3.0	4.0	4.0	4.0
	Q25, Q75	3.0, 4.0	3.0, 4.0	3.0, 5.0	3.0, 4.0
	Min, Max	2, 9	2, 7	2, 5	2, 7
12:00-01:30 PM	n	43	43	18	18
	Mean (SD)	3.2 (0.70)	5.8 (1.46)	3.0 (0.85)	2.7 (0.90)
	95% CI	2.9, 3.5	5.3, 6.3	2.5, 3.5	2.2, 3.2
	Median	3.0	6.0	3.0	2.5
	Q25, Q75	3.0, 4.0	5.0, 7.0	2.0, 4.0	2.0, 3.0
	Min, Max	2, 5	3, 9	2, 4	2, 5

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Note: T<sub>0</sub> = Time of first product use at single use day.

Appendix 15.3.3.5

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_cosing.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.10.1 Descriptive Statistics of Exhaled CO (ppm) During Single Use Continuous Measurements - PK Population**

Timepoint	Statistic	Group-1 PK		Group-2 PK	
		THS 2.2 Menthol (N=43)	mCC (N=43)	THS 2.2 Menthol (N=18)	NRT gum (N=18)
04:00-05:30 PM	n	43	43	18	18
	Mean (SD)	3.7 (1.66)	5.1 (1.73)	2.9 (0.94)	2.8 (0.93)
	95% CI	3.1, 4.2	4.5, 5.7	2.4, 3.5	2.3, 3.3
	Median	3.0	5.0	3.0	3.0
	Q25, Q75	2.0, 4.0	4.0, 6.0	2.0, 4.0	2.0, 3.0
	Min, Max	2, 10	2, 11	2, 5	1, 5
08:00-09:30 PM	n	43	43	18	18
	Mean (SD)	3.4 (1.57)	4.3 (1.44)	3.7 (1.42)	3.3 (1.13)
	95% CI	2.9, 3.9	3.8, 4.8	2.9, 4.4	2.7, 3.9
	Median	3.0	4.0	4.0	3.0
	Q25, Q75	2.0, 4.0	3.0, 5.0	2.0, 5.0	2.0, 4.0
	Min, Max	2, 8	2, 8	2, 7	2, 5

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Note: T<sub>0</sub> = Time of first product use at single use day.

Appendix 15.3.3.5

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_cosing.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.10.1.1 Descriptive Statistics of Exhaled CO (ppm) During Single Use by Sex - PK Population**

Timepoint	Statistic	Group-1 PK			
		THS 2.2 Menthol		mCC	
		Males (N=23)	Females (N=20)	Males (N=23)	Females (N=20)
15 min < T <sub>0</sub>	n	23	20	23	20
	Mean (SD)	3.6 (1.34)	3.4 (1.10)	3.8 (0.86)	3.7 (1.31)
	95% CI	3.0, 4.2	2.8, 4.0	3.4, 4.2	3.0, 4.4
	Median	3.0	3.0	4.0	3.5
	Q25, Q75	3.0, 4.0	2.5, 4.0	3.0, 4.0	3.0, 4.0
	Min, Max	2, 9	2, 5	2, 5	2, 7
12:00-01:30 PM	n	23	20	23	20
	Mean (SD)	3.4 (0.79)	3.0 (0.52)	6.0 (1.23)	5.5 (1.68)
	95% CI	3.0, 3.8	2.7, 3.2	5.5, 6.6	4.7, 6.3
	Median	3.0	3.0	6.0	5.0
	Q25, Q75	3.0, 4.0	3.0, 3.0	5.0, 7.0	4.0, 6.5
	Min, Max	2, 5	2, 4	4, 9	3, 9

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Note: T<sub>0</sub> = Time of first product use at single use day.

Appendix 15.3.3.5

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_cosing2.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.10.1.1 Descriptive Statistics of Exhaled CO (ppm) During Single Use by Sex - PK Population**

Timepoint	Statistic	Group-1 PK			
		THS 2.2 Menthol		mCC	
		Males (N=23)	Females (N=20)	Males (N=23)	Females (N=20)
04:00-05:30 PM	n	23	20	23	20
	Mean (SD)	4.2 (1.93)	3.1 (1.08)	5.4 (1.78)	4.7 (1.63)
	95% CI	3.3, 5.1	2.5, 3.7	4.6, 6.2	3.9, 5.5
	Median	4.0	3.0	5.0	5.0
	Q25, Q75	3.0, 5.0	2.0, 4.0	4.0, 7.0	4.0, 5.0
	Min, Max	2, 10	2, 5	3, 11	2, 9
08:00-09:30 PM	n	23	20	23	20
	Mean (SD)	3.6 (1.78)	3.2 (1.29)	4.6 (1.48)	4.0 (1.38)
	95% CI	2.8, 4.4	2.6, 3.8	3.9, 5.3	3.3, 4.7
	Median	3.0	3.0	4.0	4.0
	Q25, Q75	2.0, 4.0	2.0, 4.0	3.0, 6.0	3.0, 5.0
	Min, Max	2, 8	2, 6	2, 8	2, 6

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Note: T<sub>0</sub> = Time of first product use at single use day.

Appendix 15.3.3.5

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_cosing2.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.10.1.1 Descriptive Statistics of Exhaled CO (ppm) During Single Use by Sex - PK Population**

Timepoint	Statistic	Group-2 PK			
		THS 2.2 Menthol		NRT gum	
		Males (N=9)	Females (N=9)	Males (N=9)	Females (N=9)
15 min < T <sub>0</sub>	n	9	9	9	9
	Mean (SD)	3.6 (1.02)	3.9 (1.06)	3.9 (1.27)	3.6 (0.89)
	95% CI	2.7, 4.4	3.0, 4.7	2.9, 4.9	2.8, 4.3
	Median	3.0	4.0	4.0	4.0
	Q25, Q75	3.0, 4.0	3.0, 5.0	3.0, 4.0	3.0, 4.0
	Min, Max	2, 5	2, 5	3, 7	2, 5
12:00-01:30 PM	n	9	9	9	9
	Mean (SD)	3.3 (0.71)	2.7 (0.87)	3.2 (0.98)	2.2 (0.45)
	95% CI	2.7, 3.9	2.0, 3.4	2.4, 4.0	1.8, 2.6
	Median	3.0	2.0	3.0	2.0
	Q25, Q75	3.0, 4.0	2.0, 3.0	3.0, 4.0	2.0, 2.0
	Min, Max	2, 4	2, 4	2, 5	2, 3

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Note: T<sub>0</sub> = Time of first product use at single use day.

Appendix 15.3.3.5

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_cosing2.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.10.1.1 Descriptive Statistics of Exhaled CO (ppm) During Single Use by Sex - PK Population**

Timepoint	Statistic	Group-2 PK			
		THS 2.2 Menthol		NRT gum	
		Males (N=9)	Females (N=9)	Males (N=9)	Females (N=9)
04:00-05:30 PM	n	9	9	9	9
	Mean (SD)	3.1 (0.93)	2.8 (0.98)	2.8 (0.84)	2.9 (1.06)
	95% CI	2.3, 3.9	2.0, 3.6	2.1, 3.5	2.0, 3.7
	Median	3.0	2.0	3.0	3.0
	Q25, Q75	3.0, 3.0	2.0, 4.0	3.0, 3.0	2.0, 3.0
	Min, Max	2, 5	2, 4	1, 4	2, 5
08:00-09:30 PM	n	9	9	9	9
	Mean (SD)	3.7 (1.66)	3.7 (1.23)	3.2 (0.98)	3.3 (1.33)
	95% CI	2.3, 5.0	2.7, 4.7	2.4, 4.0	2.3, 4.4
	Median	4.0	4.0	3.0	3.0
	Q25, Q75	2.0, 4.0	3.0, 5.0	3.0, 4.0	2.0, 5.0
	Min, Max	2, 7	2, 5	2, 5	2, 5

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Note: T<sub>0</sub> = Time of first product use at single use day.

Appendix 15.3.3.5

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_cosing2.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.10.1.2 Descriptive Statistics of Exhaled CO (ppm) During Single Use by Nicotine Level - PK Population**

Timepoint	Statistic	Group-1 PK			
		THS 2.2 Menthol		mCC	
		<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)	<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)
15 min < T <sub>0</sub>	n	25	18	25	18
	Mean (SD)	3.2 (0.93)	3.9 (1.50)	3.5 (1.05)	4.1 (1.03)
	95% CI	2.8, 3.7	3.1, 4.7	3.0, 4.0	3.6, 4.7
	Median	3.0	4.0	3.0	4.0
	Q25, Q75	3.0, 4.0	3.0, 4.0	3.0, 4.0	3.0, 5.0
	Min, Max	2, 5	2, 9	2, 6	3, 7
12:00-01:30 PM	n	25	18	25	18
	Mean (SD)	3.1 (0.67)	3.3 (0.76)	6.0 (1.40)	5.6 (1.55)
	95% CI	2.8, 3.4	2.9, 3.7	5.3, 6.6	4.7, 6.4
	Median	3.0	3.0	6.0	5.5
	Q25, Q75	3.0, 3.0	3.0, 4.0	5.0, 7.0	5.0, 6.0
	Min, Max	2, 5	2, 5	3, 8	3, 9

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Note: T<sub>0</sub> = Time of first product use at single use day.

Appendix 15.3.3.5

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_cosing3.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.10.1.2 Descriptive Statistics of Exhaled CO (ppm) During Single Use by Nicotine Level - PK Population**

Timepoint	Statistic	Group-1 PK			
		THS 2.2 Menthol		mCC	
		<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)	<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)
04:00-05:30 PM	n	25	18	25	18
	Mean (SD)	3.4 (1.29)	4.1 (2.03)	5.0 (1.42)	5.2 (2.13)
	95% CI	2.8, 3.9	3.1, 5.2	4.4, 5.6	4.1, 6.3
	Median	3.0	3.5	5.0	5.0
	Q25, Q75	2.0, 4.0	3.0, 5.0	4.0, 6.0	4.0, 5.0
	Min, Max	2, 7	2, 10	2, 8	3, 11
08:00-09:30 PM	n	25	18	25	18
	Mean (SD)	3.2 (1.45)	3.7 (1.71)	4.0 (1.40)	4.7 (1.46)
	95% CI	2.6, 3.8	2.8, 4.6	3.4, 4.7	3.9, 5.4
	Median	3.0	3.0	4.0	4.5
	Q25, Q75	2.0, 4.0	2.0, 5.0	3.0, 5.0	3.0, 6.0
	Min, Max	2, 8	2, 8	2, 7	3, 8

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Note: T<sub>0</sub> = Time of first product use at single use day.

Appendix 15.3.3.5

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_cosing3.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.10.1.2 Descriptive Statistics of Exhaled CO (ppm) During Single Use by Nicotine Level - PK Population**

Timepoint	Statistic	Group-2 PK			
		THS 2.2 Menthol		NRT gum	
		<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)	<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)
15 min < T <sub>0</sub>	n	10	8	10	8
	Mean (SD)	3.8 (0.92)	3.6 (1.19)	3.8 (0.64)	3.6 (1.51)
	95% CI	3.1, 4.5	2.6, 4.7	3.3, 4.3	2.3, 4.9
	Median	3.5	4.0	4.0	3.0
	Q25, Q75	3.0, 5.0	2.5, 4.5	3.0, 4.0	3.0, 4.0
	Min, Max	3, 5	2, 5	3, 5	2, 7
12:00-01:30 PM	n	10	8	10	8
	Mean (SD)	3.0 (0.82)	3.0 (0.93)	2.7 (0.68)	2.8 (1.17)
	95% CI	2.4, 3.6	2.2, 3.8	2.2, 3.2	1.7, 3.8
	Median	3.0	3.0	3.0	2.0
	Q25, Q75	2.0, 4.0	2.0, 4.0	2.0, 3.0	2.0, 3.5
	Min, Max	2, 4	2, 4	2, 4	2, 5

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Note: T<sub>0</sub> = Time of first product use at single use day.

Appendix 15.3.3.5

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_cosing3.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.10.1.2 Descriptive Statistics of Exhaled CO (ppm) During Single Use by Nicotine Level - PK Population**

Timepoint	Statistic	Group-2 PK			
		THS 2.2 Menthol		NRT gum	
		<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)	<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)
04:00-05:30 PM	n	10	8	10	8
	Mean (SD)	3.0 (0.67)	2.9 (1.25)	3.1 (1.00)	2.5 (0.76)
	95% CI	2.5, 3.5	1.8, 4.0	2.3, 3.9	1.8, 3.2
	Median	3.0	2.0	3.0	3.0
	Q25, Q75	3.0, 3.0	2.0, 4.0	2.0, 4.0	2.0, 3.0
	Min, Max	2, 4	2, 5	2, 5	1, 3
08:00-09:30 PM	n	10	8	10	8
	Mean (SD)	3.4 (1.18)	4.0 (1.70)	3.6 (1.08)	2.9 (1.13)
	95% CI	2.5, 4.3	2.5, 5.5	2.8, 4.4	1.9, 3.9
	Median	3.5	4.0	3.0	2.5
	Q25, Q75	2.0, 4.0	2.5, 5.0	3.0, 5.0	2.0, 3.5
	Min, Max	2, 5	2, 7	2, 5	2, 5

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Note: T<sub>0</sub> = Time of first product use at single use day.

Appendix 15.3.3.5

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_cosing3.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.10.2 Descriptive Statistics of Exhaled CO (ppm) During Single Use Categorical Measurements - PK Population**

Timepoint	Statistic	Group-1 PK		Group-2 PK	
		THS 2.2 Menthol (N=43)	mCC (N=43)	THS 2.2 Menthol (N=18)	NRT gum (N=18)
15 min < T <sub>0</sub>	CO ≤ 10 ppm - n (%)	43 (100 %)	43 (100 %)	18 (100 %)	18 (100 %)
	CO > 10 ppm - n (%)	0	0	0	0
12:00-01:30 PM	CO ≤ 10 ppm - n (%)	43 (100 %)	43 (100 %)	18 (100 %)	18 (100 %)
	CO > 10 ppm - n (%)	0	0	0	0
04:00-05:30 PM	CO ≤ 10 ppm - n (%)	43 (100 %)	42 ( 97.7%)	18 (100 %)	18 (100 %)
	CO > 10 ppm - n (%)	0	1 ( 2.3%)	0	0
08:00-09:30 PM	CO ≤ 10 ppm - n (%)	43 (100 %)	43 (100 %)	18 (100 %)	18 (100 %)
	CO > 10 ppm - n (%)	0	0	0	0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Note: T<sub>0</sub> = Time of first product use at single use day.

Appendix 15.3.3.5

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_cosingcat.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.11.1 Descriptive Statistics of Exhaled CO (ppm) During Days -1, 0, 2 and 4 Continuous Measurements - PK Population**

Timepoint	Statistic	Group-1 PK		Group-2 PK	
		THS 2.2 Menthol - mCC (N=21)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)
Day -1	n	21	22	9	9
	Mean (SD)	14.3 (5.09)	14.9 (6.68)	14.3 (7.69)	16.4 (8.58)
	95% CI	11.9, 16.6	11.9, 17.9	8.4, 20.3	9.8, 23.1
	Median	15.0	14.0	12.0	14.0
	Q25, Q75	10.0, 18.0	9.0, 20.0	9.0, 16.0	14.0, 20.0
	Min, Max	7, 25	5, 28	7, 29	7, 36
Washout Period 1 (Day 0 08:00-09:30 AM)	n	21	22	9	9
	Mean (SD)	9.6 (3.06)	9.8 (3.10)	8.3 (1.42)	9.1 (2.90)
	95% CI	8.1, 11.0	8.4, 11.2	7.2, 9.5	6.8, 11.4
	Median	9.0	9.0	8.0	7.0
	Q25, Q75	7.0, 10.0	7.0, 12.0	8.0, 9.0	7.0, 11.0
	Min, Max	6, 17	6, 16	6, 11	6, 14

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Appendix 15.3.3.5

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**Table 15.2.4.11.1 Descriptive Statistics of Exhaled CO (ppm) During Days -1, 0, 2 and 4 Continuous Measurements - PK Population**

Timepoint	Statistic	Group-1 PK		Group-2 PK	
		THS 2.2 Menthol - mCC (N=21)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)
Washout Period 1 (Day 0 12:00-01:30 PM)	n	21	22	9	9
	Mean (SD)	6.9 (2.10)	6.8 (2.19)	6.0 (1.81)	6.4 (2.13)
	95% CI	5.9, 7.9	5.8, 7.8	4.6, 7.4	4.8, 8.1
	Median	7.0	6.5	7.0	6.0
	Q25, Q75	5.0, 9.0	5.0, 8.0	5.0, 7.0	5.0, 9.0
	Min, Max	4, 10	4, 11	3, 8	4, 9
Washout Period 1 (Day 0 04:00-05:30 PM)	n	21	22	9	9
	Mean (SD)	5.4 (1.64)	5.5 (1.51)	4.9 (1.37)	4.6 (1.14)
	95% CI	4.6, 6.2	4.8, 6.3	3.8, 6.0	3.6, 5.5
	Median	5.0	6.0	5.0	5.0
	Q25, Q75	4.0, 6.0	4.0, 7.0	4.0, 5.0	4.0, 5.0
	Min, Max	3, 10	3, 8	3, 7	3, 6

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Appendix 15.3.3.5

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_coday.sas

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**Table 15.2.4.11.1 Descriptive Statistics of Exhaled CO (ppm) During Days -1, 0, 2 and 4 Continuous Measurements - PK Population**

Timepoint	Statistic	Group-1 PK		Group-2 PK	
		THS 2.2 Menthol - mCC (N=21)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)
Washout Period 1 (Day 0 08:00-09:30 PM)	n	21	22	9	9
	Mean (SD)	4.7 (1.31)	4.4 (1.44)	4.4 (1.24)	3.6 (1.14)
	95% CI	4.1, 5.4	3.7, 5.1	3.4, 5.4	2.6, 4.5
	Median	5.0	4.0	5.0	4.0
	Q25, Q75	4.0, 5.0	4.0, 5.0	3.0, 5.0	3.0, 4.0
	Min, Max	2, 8	2, 8	3, 6	2, 5
Washout Period 2 (Day 2 08:00-09:30 AM)	n	21	22	9	9
	Mean (SD)	3.5 (1.08)	3.2 (0.91)	3.1 (0.61)	2.9 (1.06)
	95% CI	2.9, 4.0	2.7, 3.6	2.6, 3.6	2.0, 3.7
	Median	3.0	3.0	3.0	3.0
	Q25, Q75	3.0, 4.0	3.0, 4.0	3.0, 3.0	3.0, 3.0
	Min, Max	2, 6	1, 5	2, 4	1, 5

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Appendix 15.3.3.5

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**Table 15.2.4.11.1 Descriptive Statistics of Exhaled CO (ppm) During Days -1, 0, 2 and 4 Continuous Measurements - PK Population**

Timepoint	Statistic	Group-1 PK		Group-2 PK	
		THS 2.2 Menthol - mCC (N=21)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)
Washout Period 2 (Day 2 12:00-01:30 PM)	n	21	22	9	9
	Mean (SD)	3.1 (0.77)	3.0 (0.79)	2.8 (0.84)	2.6 (0.89)
	95% CI	2.7, 3.5	2.6, 3.4	2.1, 3.5	1.8, 3.3
	Median	3.0	3.0	3.0	3.0
	Q25, Q75	3.0, 3.0	3.0, 3.0	2.0, 3.0	2.0, 3.0
	Min, Max	2, 5	2, 5	2, 4	1, 4
Washout Period 2 (Day 2 04:00-05:30 PM)	n	21	22	9	9
	Mean (SD)	3.4 (1.08)	3.0 (0.98)	3.1 (0.61)	2.8 (0.84)
	95% CI	2.8, 3.9	2.5, 3.5	2.6, 3.6	2.1, 3.5
	Median	4.0	3.0	3.0	3.0
	Q25, Q75	2.0, 4.0	2.0, 3.0	3.0, 3.0	2.0, 3.0
	Min, Max	2, 5	2, 6	2, 4	2, 4

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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**Table 15.2.4.11.1 Descriptive Statistics of Exhaled CO (ppm) During Days -1, 0, 2 and 4 Continuous Measurements - PK Population**

Timepoint	Statistic	Group-1 PK		Group-2 PK	
		THS 2.2 Menthol - mCC (N=21)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)
Washout Period 2 (Day 2 08:00-09:30 PM)	n	21	22	9	9
	Mean (SD)	3.5 (1.44)	3.5 (1.41)	3.6 (1.24)	3.2 (1.31)
	95% CI	2.8, 4.2	2.8, 4.1	2.6, 4.6	2.2, 4.3
	Median	4.0	3.0	4.0	4.0
	Q25, Q75	2.0, 5.0	2.0, 4.0	3.0, 4.0	2.0, 4.0
	Min, Max	1, 5	2, 6	1, 5	1, 5
Day 4/Discharge	n	21	22	9	9
	Mean (SD)	3.2 (0.75)	2.8 (0.53)	2.8 (0.67)	2.8 (0.67)
	95% CI	2.8, 3.6	2.5, 3.1	2.2, 3.3	2.2, 3.3
	Median	3.0	3.0	3.0	3.0
	Q25, Q75	3.0, 4.0	2.0, 3.0	2.0, 3.0	2.0, 3.0
	Min, Max	2, 5	2, 4	2, 4	2, 4

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.



**Table 15.2.4.11.1.1 Descriptive Statistics of Exhaled CO (ppm) During Days -1, 0, 2 and 4 by Sex - PK Population**

Timepoint	Statistic	Group-1 PK			
		THS 2.2 Menthol - mCC		mCC - THS 2.2 Menthol	
		Males (N=11)	Females (N=10)	Males (N=12)	Females (N=10)
Day -1	n	11	10	12	10
	Mean (SD)	12.3 (4.30)	16.5 (5.15)	13.5 (6.44)	16.6 (6.90)
	95% CI	9.3, 15.2	12.8, 20.2	9.4, 17.6	11.6, 21.6
	Median	11.0	16.5	13.5	14.0
	Q25, Q75	8.0, 17.0	11.0, 20.0	8.0, 19.0	11.0, 22.0
	Min, Max	7, 18	10, 25	5, 23	9, 28
Washout Period 1 (Day 0 08:00-09:30 AM)	n	11	10	12	10
	Mean (SD)	9.8 (2.14)	9.3 (3.95)	9.5 (2.82)	10.2 (3.53)
	95% CI	8.3, 11.3	6.4, 12.2	7.7, 11.3	7.6, 12.8
	Median	9.0	7.5	9.0	9.5
	Q25, Q75	8.0, 11.0	7.0, 10.0	7.5, 11.0	7.0, 13.0
	Min, Max	7, 14	6, 17	6, 16	6, 15

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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**Table 15.2.4.11.1.1 Descriptive Statistics of Exhaled CO (ppm) During Days -1, 0, 2 and 4 by Sex - PK Population**

Timepoint	Statistic	Group-1 PK			
		THS 2.2 Menthol - mCC		mCC - THS 2.2 Menthol	
		Males (N=11)	Females (N=10)	Males (N=12)	Females (N=10)
Washout Period 1 (Day 0 12:00-01:30 PM)	n	11	10	12	10
	Mean (SD)	7.3 (1.91)	6.5 (2.33)	6.8 (1.96)	6.8 (2.53)
	95% CI	5.9, 8.6	4.8, 8.2	5.5, 8.0	4.9, 8.7
	Median	7.0	6.5	7.0	6.0
	Q25, Q75	6.0, 9.0	4.0, 8.0	5.0, 7.5	5.0, 9.0
	Min, Max	4, 10	4, 10	4, 11	4, 11
Washout Period 1 (Day 0 04:00-05:30 PM)	n	11	10	12	10
	Mean (SD)	5.9 (1.65)	4.9 (1.53)	5.8 (1.41)	5.2 (1.62)
	95% CI	4.8, 7.1	3.8, 6.0	4.9, 6.8	4.0, 6.4
	Median	5.0	4.5	6.0	5.5
	Q25, Q75	5.0, 7.0	4.0, 6.0	4.5, 7.0	4.0, 7.0
	Min, Max	4, 10	3, 7	4, 8	3, 7

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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**Table 15.2.4.11.1.1 Descriptive Statistics of Exhaled CO (ppm) During Days -1, 0, 2 and 4 by Sex - PK Population**

Timepoint	Statistic	Group-1 PK			
		THS 2.2 Menthol - mCC		mCC - THS 2.2 Menthol	
		Males (N=11)	Females (N=10)	Males (N=12)	Females (N=10)
Washout Period 1 (Day 0 08:00-09:30 PM)	n	11	10	12	10
	Mean (SD)	5.1 (1.31)	4.3 (1.26)	4.9 (1.25)	3.8 (1.48)
	95% CI	4.2, 6.0	3.4, 5.2	4.1, 5.8	2.7, 4.9
	Median	5.0	4.0	4.5	4.0
	Q25, Q75	4.0, 5.0	4.0, 5.0	4.0, 5.5	2.0, 5.0
	Min, Max	4, 8	2, 6	4, 8	2, 6
Washout Period 2 (Day 2 08:00-09:30 AM)	n	11	10	12	10
	Mean (SD)	3.8 (1.08)	3.1 (1.00)	3.5 (0.80)	2.8 (0.92)
	95% CI	3.0, 4.6	2.3, 3.9	2.9, 4.1	2.1, 3.5
	Median	4.0	3.0	3.5	3.0
	Q25, Q75	3.0, 4.0	2.0, 4.0	3.0, 4.0	2.0, 3.0
	Min, Max	2, 6	2, 5	2, 5	1, 4

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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**Table 15.2.4.11.1.1 Descriptive Statistics of Exhaled CO (ppm) During Days -1, 0, 2 and 4 by Sex - PK Population**

Timepoint	Statistic	Group-1 PK			
		THS 2.2 Menthol - mCC		mCC - THS 2.2 Menthol	
		Males (N=11)	Females (N=10)	Males (N=12)	Females (N=10)
Washout Period 2 (Day 2 12:00-01:30 PM)	n	11	10	12	10
	Mean (SD)	3.5 (0.69)	2.7 (0.68)	3.3 (0.66)	2.7 (0.83)
	95% CI	2.9, 4.0	2.2, 3.2	2.9, 3.8	2.1, 3.3
	Median	3.0	3.0	3.0	2.5
	Q25, Q75	3.0, 4.0	2.0, 3.0	3.0, 3.5	2.0, 3.0
	Min, Max	3, 5	2, 4	3, 5	2, 4
Washout Period 2 (Day 2 04:00-05:30 PM)	n	11	10	12	10
	Mean (SD)	3.8 (1.08)	2.9 (0.88)	3.2 (1.12)	2.8 (0.79)
	95% CI	3.0, 4.6	2.2, 3.6	2.4, 3.9	2.2, 3.4
	Median	4.0	3.0	3.0	3.0
	Q25, Q75	3.0, 5.0	2.0, 4.0	2.5, 3.5	2.0, 3.0
	Min, Max	2, 5	2, 4	2, 6	2, 4

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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**Table 15.2.4.11.1.1 Descriptive Statistics of Exhaled CO (ppm) During Days -1, 0, 2 and 4 by Sex - PK Population**

Timepoint	Statistic	Group-1 PK			
		THS 2.2 Menthol - mCC		mCC - THS 2.2 Menthol	
		Males (N=11)	Females (N=10)	Males (N=12)	Females (N=10)
Washout Period 2 (Day 2 08:00-09:30 PM)	n	11	10	12	10
	Mean (SD)	3.9 (1.31)	3.0 (1.50)	3.7 (1.56)	3.2 (1.23)
	95% CI	3.0, 4.8	1.9, 4.1	2.6, 4.7	2.3, 4.1
	Median	4.0	3.0	3.0	3.0
	Q25, Q75	2.0, 5.0	2.0, 4.0	2.5, 5.0	2.0, 4.0
	Min, Max	2, 5	1, 5	2, 6	2, 5
Day 4/Discharge	n	11	10	12	10
	Mean (SD)	3.4 (0.51)	3.0 (0.95)	2.8 (0.39)	2.7 (0.68)
	95% CI	3.0, 3.8	2.3, 3.7	2.5, 3.1	2.2, 3.2
	Median	3.0	3.0	3.0	3.0
	Q25, Q75	3.0, 4.0	2.0, 3.0	3.0, 3.0	2.0, 3.0
	Min, Max	3, 4	2, 5	2, 3	2, 4

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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**Table 15.2.4.11.1.1 Descriptive Statistics of Exhaled CO (ppm) During Days -1, 0, 2 and 4 by Sex - PK Population**

Timepoint	Statistic	Group-2 PK			
		THS 2.2 Menthol - NRT gum		NRT gum - THS 2.2 Menthol	
		Males (N=5)	Females (N=4)	Males (N=4)	Females (N=5)
Day -1	n	5	4	4	5
	Mean (SD)	11.2 (3.43)	18.3 (10.25)	10.8 (3.78)	21.0 (8.84)
	95% CI	6.9, 15.5	1.9, 34.6	4.7, 16.8	10.0, 32.0
	Median	12.0	17.5	11.0	20.0
	Q25, Q75	9.0, 12.0	9.5, 27.0	7.5, 14.0	15.0, 20.0
	Min, Max	7, 16	9, 29	7, 14	14, 36
Washout Period 1 (Day 0 08:00-09:30 AM)	n	5	4	4	5
	Mean (SD)	8.0 (1.23)	8.8 (1.71)	9.3 (2.63)	9.0 (3.40)
	95% CI	6.4, 9.6	6.0, 11.5	5.0, 13.5	4.7, 13.3
	Median	8.0	8.5	9.0	7.0
	Q25, Q75	8.0, 9.0	7.5, 10.0	7.0, 11.5	7.0, 11.0
	Min, Max	6, 9	7, 11	7, 12	6, 14

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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**Table 15.2.4.11.1.1 Descriptive Statistics of Exhaled CO (ppm) During Days -1, 0, 2 and 4 by Sex - PK Population**

Timepoint	Statistic	Group-2 PK			
		THS 2.2 Menthol - NRT gum		NRT gum - THS 2.2 Menthol	
		Males (N=5)	Females (N=4)	Males (N=4)	Females (N=5)
Washout Period 1 (Day 0 12:00-01:30 PM)	n	5	4	4	5
	Mean (SD)	6.0 (2.35)	6.0 (1.16)	7.0 (2.31)	6.0 (2.13)
	95% CI	3.0, 9.0	4.1, 7.9	3.3, 10.7	3.3, 8.7
	Median	7.0	6.0	7.0	6.0
	Q25, Q75	4.0, 8.0	5.0, 7.0	5.0, 9.0	4.0, 7.0
	Min, Max	3, 8	5, 7	5, 9	4, 9
Washout Period 1 (Day 0 04:00-05:30 PM)	n	5	4	4	5
	Mean (SD)	5.2 (1.79)	4.5 (0.58)	5.0 (1.42)	4.2 (0.84)
	95% CI	2.9, 7.5	3.5, 5.5	2.7, 7.3	3.1, 5.3
	Median	5.0	4.5	5.5	4.0
	Q25, Q75	4.0, 7.0	4.0, 5.0	4.0, 6.0	4.0, 5.0
	Min, Max	3, 7	4, 5	3, 6	3, 5

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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**Table 15.2.4.11.1.1 Descriptive Statistics of Exhaled CO (ppm) During Days -1, 0, 2 and 4 by Sex - PK Population**

Timepoint	Statistic	Group-2 PK			
		THS 2.2 Menthol - NRT gum		NRT gum - THS 2.2 Menthol	
		Males (N=5)	Females (N=4)	Males (N=4)	Females (N=5)
Washout Period 1 (Day 0 08:00-09:30 PM)	n	5	4	4	5
	Mean (SD)	4.6 (1.15)	4.3 (1.50)	4.3 (0.96)	3.0 (1.00)
	95% CI	3.1, 6.1	1.8, 6.7	2.7, 5.8	1.7, 4.3
	Median	5.0	4.0	4.5	3.0
	Q25, Q75	4.0, 5.0	3.0, 5.5	3.5, 5.0	2.0, 4.0
	Min, Max	3, 6	3, 6	3, 5	2, 4
Washout Period 2 (Day 2 08:00-09:30 AM)	n	5	4	4	5
	Mean (SD)	3.4 (0.55)	2.8 (0.50)	3.3 (1.26)	2.6 (0.90)
	95% CI	2.7, 4.1	1.9, 3.6	1.2, 5.3	1.4, 3.8
	Median	3.0	3.0	3.0	3.0
	Q25, Q75	3.0, 4.0	2.5, 3.0	2.5, 4.0	3.0, 3.0
	Min, Max	3, 4	2, 3	2, 5	1, 3

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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**Table 15.2.4.11.1.1 Descriptive Statistics of Exhaled CO (ppm) During Days -1, 0, 2 and 4 by Sex - PK Population**

Timepoint	Statistic	Group-2 PK			
		THS 2.2 Menthol - NRT gum		NRT gum - THS 2.2 Menthol	
		Males (N=5)	Females (N=4)	Males (N=4)	Females (N=5)
Washout Period 2 (Day 2 12:00-01:30 PM)	n	5	4	4	5
	Mean (SD)	3.4 (0.55)	2.0 (0.00)	3.0 (0.82)	2.2 (0.84)
	95% CI	2.7, 4.1	NC	1.7, 4.3	1.1, 3.3
	Median	3.0	2.0	3.0	2.0
	Q25, Q75	3.0, 4.0	2.0, 2.0	2.5, 3.5	2.0, 3.0
	Min, Max	3, 4	2, 2	2, 4	1, 3
Washout Period 2 (Day 2 04:00-05:30 PM)	n	5	4	4	5
	Mean (SD)	3.4 (0.55)	2.8 (0.50)	3.0 (0.82)	2.6 (0.90)
	95% CI	2.7, 4.1	1.9, 3.6	1.7, 4.3	1.4, 3.8
	Median	3.0	3.0	3.0	2.0
	Q25, Q75	3.0, 4.0	2.5, 3.0	2.5, 3.5	2.0, 3.0
	Min, Max	3, 4	2, 3	2, 4	2, 4

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Note: NC = Not calculated.

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**Table 15.2.4.11.1.1 Descriptive Statistics of Exhaled CO (ppm) During Days -1, 0, 2 and 4 by Sex - PK Population**

Timepoint	Statistic	Group-2 PK			
		THS 2.2 Menthol - NRT gum		NRT gum - THS 2.2 Menthol	
		Males (N=5)	Females (N=4)	Males (N=4)	Females (N=5)
Washout Period 2 (Day 2 08:00-09:30 PM)	n	5	4	4	5
	Mean (SD)	3.8 (0.84)	3.3 (1.71)	3.0 (1.42)	3.4 (1.35)
	95% CI	2.7, 4.9	0.5, 6.0	0.7, 5.3	1.7, 5.1
	Median	4.0	3.5	3.5	4.0
	Q25, Q75	3.0, 4.0	2.0, 4.5	2.0, 4.0	2.0, 4.0
	Min, Max	3, 5	1, 5	1, 4	2, 5
Day 4/Discharge	n	5	4	4	5
	Mean (SD)	3.2 (0.45)	2.3 (0.50)	3.3 (0.50)	2.4 (0.55)
	95% CI	2.6, 3.8	1.4, 3.1	2.4, 4.1	1.7, 3.1
	Median	3.0	2.0	3.0	2.0
	Q25, Q75	3.0, 3.0	2.0, 2.5	3.0, 3.5	2.0, 3.0
	Min, Max	3, 4	2, 3	3, 4	2, 3

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Appendix 15.3.3.5

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_coday2.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.11.1.2 Descriptive Statistics of Exhaled CO (ppm) During Days -1, 0, 2 and 4 by Nicotine Level - PK Population**

Timepoint	Statistic	Group-1 PK			
		THS 2.2 Menthol - mCC		mCC - THS 2.2 Menthol	
		<= 0.6 mg (N=13)	> 0.6 - 1.0 mg (N=8)	<= 0.6 mg (N=12)	> 0.6 - 1.0 mg (N=10)
Day -1	n	13	8	12	10
	Mean (SD)	15.5 (5.46)	12.4 (4.00)	13.8 (6.90)	16.3 (6.49)
	95% CI	12.1, 18.8	9.0, 15.8	9.3, 18.2	11.6, 21.0
	Median	16.0	12.0	12.0	16.0
	Q25, Q75	10.0, 19.0	9.0, 16.5	9.0, 18.0	10.0, 22.0
	Min, Max	8, 25	7, 17	5, 28	7, 27
Washout Period 1 (Day 0 08:00-09:30 AM)	n	13	8	12	10
	Mean (SD)	10.0 (3.47)	8.9 (2.30)	9.9 (3.71)	9.7 (2.36)
	95% CI	7.9, 12.1	6.9, 10.8	7.5, 12.3	8.0, 11.4
	Median	9.0	8.0	9.0	9.5
	Q25, Q75	7.0, 11.0	7.5, 9.5	6.5, 13.0	8.0, 11.0
	Min, Max	6, 17	7, 14	6, 16	7, 15

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Appendix 15.3.3.5

Path: /cvm/projects/prj/development/000000106326/dev/tables/t\_coday3.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.11.1.2 Descriptive Statistics of Exhaled CO (ppm) During Days -1, 0, 2 and 4 by Nicotine Level - PK Population**

Timepoint	Statistic	Group-1 PK			
		THS 2.2 Menthol - mCC		mCC - THS 2.2 Menthol	
		<= 0.6 mg (N=13)	> 0.6 - 1.0 mg (N=8)	<= 0.6 mg (N=12)	> 0.6 - 1.0 mg (N=10)
Washout Period 1 (Day 0 12:00-01:30 PM)	n	13	8	12	10
	Mean (SD)	7.0 (2.45)	6.8 (1.49)	7.1 (2.58)	6.4 (1.65)
	95% CI	5.5, 8.5	5.5, 8.0	5.4, 8.8	5.2, 7.6
	Median	7.0	7.0	7.0	6.0
	Q25, Q75	5.0, 9.0	6.0, 7.5	5.0, 9.0	5.0, 7.0
	Min, Max	4, 10	4, 9	4, 11	4, 9
Washout Period 1 (Day 0 04:00-05:30 PM)	n	13	8	12	10
	Mean (SD)	5.7 (1.85)	5.0 (1.20)	5.6 (1.63)	5.5 (1.44)
	95% CI	4.5, 6.9	4.0, 6.0	4.5, 6.7	4.4, 6.6
	Median	6.0	5.0	6.0	5.0
	Q25, Q75	4.0, 7.0	4.5, 5.5	4.0, 7.0	4.0, 7.0
	Min, Max	3, 10	3, 7	3, 7	4, 8

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Appendix 15.3.3.5

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_coday3.sas

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**Table 15.2.4.11.1.2 Descriptive Statistics of Exhaled CO (ppm) During Days -1, 0, 2 and 4 by Nicotine Level - PK Population**

Timepoint	Statistic	Group-1 PK			
		THS 2.2 Menthol - mCC		mCC - THS 2.2 Menthol	
		<= 0.6 mg (N=13)	> 0.6 - 1.0 mg (N=8)	<= 0.6 mg (N=12)	> 0.6 - 1.0 mg (N=10)
Washout Period 1 (Day 0 08:00-09:30 PM)	n	13	8	12	10
	Mean (SD)	4.7 (1.38)	4.8 (1.29)	4.0 (1.28)	4.9 (1.53)
	95% CI	3.8, 5.6	3.6, 5.9	3.1, 4.9	3.8, 6.0
	Median	5.0	4.5	4.0	5.0
	Q25, Q75	4.0, 5.0	4.0, 5.5	3.5, 4.5	4.0, 5.0
	Min, Max	2, 8	3, 7	2, 6	2, 8
Washout Period 2 (Day 2 08:00-09:30 AM)	n	13	8	12	10
	Mean (SD)	3.5 (1.13)	3.5 (1.07)	3.1 (0.80)	3.3 (1.06)
	95% CI	2.7, 4.2	2.6, 4.4	2.5, 3.6	2.5, 4.1
	Median	3.0	4.0	3.0	3.0
	Q25, Q75	3.0, 4.0	2.5, 4.0	2.5, 4.0	3.0, 4.0
	Min, Max	2, 6	2, 5	2, 4	1, 5

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Appendix 15.3.3.5

Path: /cvm/projects/prj/development/000000106326/dev/tables/t\_coday3.sas

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**Table 15.2.4.11.1.2 Descriptive Statistics of Exhaled CO (ppm) During Days -1, 0, 2 and 4 by Nicotine Level - PK Population**

Timepoint	Statistic	Group-1 PK			
		THS 2.2 Menthol - mCC		mCC - THS 2.2 Menthol	
		<= 0.6 mg (N=13)	> 0.6 - 1.0 mg (N=8)	<= 0.6 mg (N=12)	> 0.6 - 1.0 mg (N=10)
Washout Period 2 (Day 2 12:00-01:30 PM)	n	13	8	12	10
	Mean (SD)	3.0 (0.82)	3.3 (0.71)	2.9 (0.67)	3.2 (0.92)
	95% CI	2.5, 3.5	2.6, 3.9	2.4, 3.4	2.5, 3.9
	Median	3.0	3.0	3.0	3.0
	Q25, Q75	3.0, 3.0	3.0, 4.0	2.5, 3.0	3.0, 4.0
	Min, Max	2, 5	2, 4	2, 4	2, 5
Washout Period 2 (Day 2 04:00-05:30 PM)	n	13	8	12	10
	Mean (SD)	3.5 (1.27)	3.3 (0.71)	2.8 (0.72)	3.2 (1.23)
	95% CI	2.6, 4.3	2.6, 3.9	2.3, 3.3	2.3, 4.1
	Median	4.0	3.0	3.0	3.0
	Q25, Q75	2.0, 4.0	3.0, 4.0	2.0, 3.0	2.0, 4.0
	Min, Max	2, 5	2, 4	2, 4	2, 6

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Appendix 15.3.3.5

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_coday3.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.11.1.2 Descriptive Statistics of Exhaled CO (ppm) During Days -1, 0, 2 and 4 by Nicotine Level - PK Population**

Timepoint	Statistic	Group-1 PK			
		THS 2.2 Menthol - mCC		mCC - THS 2.2 Menthol	
		<= 0.6 mg (N=13)	> 0.6 - 1.0 mg (N=8)	<= 0.6 mg (N=12)	> 0.6 - 1.0 mg (N=10)
Washout Period 2 (Day 2 08:00-09:30 PM)	n	13	8	12	10
	Mean (SD)	3.4 (1.51)	3.6 (1.41)	3.2 (1.27)	3.8 (1.55)
	95% CI	2.4, 4.3	2.4, 4.9	2.3, 4.0	2.6, 5.0
	Median	4.0	4.0	3.0	4.0
	Q25, Q75	2.0, 5.0	2.0, 5.0	2.0, 3.5	2.0, 5.0
	Min, Max	1, 5	2, 5	2, 6	2, 6
Day 4/Discharge	n	13	8	12	10
	Mean (SD)	3.1 (0.87)	3.4 (0.52)	2.8 (0.63)	2.8 (0.43)
	95% CI	2.5, 3.6	2.9, 3.9	2.3, 3.2	2.4, 3.2
	Median	3.0	3.0	3.0	3.0
	Q25, Q75	3.0, 3.0	3.0, 4.0	2.0, 3.0	3.0, 3.0
	Min, Max	2, 5	3, 4	2, 4	2, 3

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Appendix 15.3.3.5

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_coday3.sas

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**Table 15.2.4.11.1.2 Descriptive Statistics of Exhaled CO (ppm) During Days -1, 0, 2 and 4 by Nicotine Level - PK Population**

Timepoint	Statistic	Group-2 PK			
		THS 2.2 Menthol - NRT gum		NRT gum - THS 2.2 Menthol	
		<= 0.6 mg (N=5)	> 0.6 - 1.0 mg (N=4)	<= 0.6 mg (N=5)	> 0.6 - 1.0 mg (N=4)
Day -1	n	5	4	5	4
	Mean (SD)	16.4 (9.94)	11.8 (3.10)	18.6 (9.74)	13.8 (7.23)
	95% CI	4.0, 28.8	6.8, 16.7	6.5, 30.7	2.2, 25.3
	Median	12.0	11.0	14.0	14.0
	Q25, Q75	9.0, 25.0	9.5, 14.0	14.0, 15.0	7.5, 20.0
	Min, Max	7, 29	9, 16	14, 36	7, 20
Washout Period 1 (Day 0 08:00-09:30 AM)	n	5	4	5	4
	Mean (SD)	8.2 (1.79)	8.5 (1.00)	10.8 (2.95)	7.0 (0.00)
	95% CI	5.9, 10.5	6.9, 10.1	7.1, 14.5	NC
	Median	8.0	9.0	11.0	7.0
	Q25, Q75	8.0, 8.0	8.0, 9.0	11.0, 12.0	7.0, 7.0
	Min, Max	6, 11	7, 9	6, 14	7, 7

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Note: NC = Not calculated.

Appendix 15.3.3.5

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_coday3.sas

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**Table 15.2.4.11.1.2 Descriptive Statistics of Exhaled CO (ppm) During Days -1, 0, 2 and 4 by Nicotine Level - PK Population**

Timepoint	Statistic	Group-2 PK			
		THS 2.2 Menthol - NRT gum		NRT gum - THS 2.2 Menthol	
		<= 0.6 mg (N=5)	> 0.6 - 1.0 mg (N=4)	<= 0.6 mg (N=5)	> 0.6 - 1.0 mg (N=4)
Washout Period 1 (Day 0 12:00-01:30 PM)	n	5	4	5	4
	Mean (SD)	5.6 (1.95)	6.5 (1.74)	7.6 (2.20)	5.0 (0.82)
	95% CI	3.1, 8.1	3.7, 9.3	4.8, 10.4	3.7, 6.3
	Median	7.0	6.5	9.0	5.0
	Q25, Q75	4.0, 7.0	5.0, 8.0	7.0, 9.0	4.5, 5.5
	Min, Max	3, 7	5, 8	4, 9	4, 6
Washout Period 1 (Day 0 04:00-05:30 PM)	n	5	4	5	4
	Mean (SD)	4.8 (1.49)	5.0 (1.42)	5.0 (1.00)	4.0 (1.16)
	95% CI	2.9, 6.7	2.7, 7.3	3.7, 6.3	2.1, 5.9
	Median	5.0	4.5	5.0	4.0
	Q25, Q75	4.0, 5.0	4.0, 6.0	4.0, 6.0	3.0, 5.0
	Min, Max	3, 7	4, 7	4, 6	3, 5

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Appendix 15.3.3.5

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**Table 15.2.4.11.1.2 Descriptive Statistics of Exhaled CO (ppm) During Days -1, 0, 2 and 4 by Nicotine Level - PK Population**

Timepoint	Statistic	Group-2 PK			
		THS 2.2 Menthol - NRT gum		NRT gum - THS 2.2 Menthol	
		<= 0.6 mg (N=5)	> 0.6 - 1.0 mg (N=4)	<= 0.6 mg (N=5)	> 0.6 - 1.0 mg (N=4)
Washout Period 1 (Day 0 08:00-09:30 PM)	n	5	4	5	4
	Mean (SD)	4.6 (1.15)	4.3 (1.50)	3.6 (1.15)	3.5 (1.30)
	95% CI	3.1, 6.1	1.8, 6.7	2.1, 5.1	1.4, 5.6
	Median	5.0	4.0	4.0	3.5
	Q25, Q75	4.0, 5.0	3.0, 5.5	3.0, 4.0	2.5, 4.5
	Min, Max	3, 6	3, 6	2, 5	2, 5
Washout Period 2 (Day 2 08:00-09:30 AM)	n	5	4	5	4
	Mean (SD)	3.0 (0.71)	3.3 (0.50)	3.0 (0.00)	2.8 (1.71)
	95% CI	2.1, 3.9	2.4, 4.1	NC	0.0, 5.5
	Median	3.0	3.0	3.0	2.5
	Q25, Q75	3.0, 3.0	3.0, 3.5	3.0, 3.0	1.5, 4.0
	Min, Max	2, 4	3, 4	3, 3	1, 5

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Note: NC = Not calculated.

Appendix 15.3.3.5

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_coday3.sas

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**Table 15.2.4.11.1.2 Descriptive Statistics of Exhaled CO (ppm) During Days -1, 0, 2 and 4 by Nicotine Level - PK Population**

Timepoint	Statistic	Group-2 PK			
		THS 2.2 Menthol - NRT gum		NRT gum - THS 2.2 Menthol	
		<= 0.6 mg (N=5)	> 0.6 - 1.0 mg (N=4)	<= 0.6 mg (N=5)	> 0.6 - 1.0 mg (N=4)
Washout Period 2 (Day 2 12:00-01:30 PM)	n	5	4	5	4
	Mean (SD)	2.8 (0.84)	2.8 (0.96)	2.6 (0.55)	2.5 (1.30)
	95% CI	1.7, 3.9	1.2, 4.3	1.9, 3.3	0.4, 4.6
	Median	3.0	2.5	3.0	2.5
	Q25, Q75	2.0, 3.0	2.0, 3.5	2.0, 3.0	1.5, 3.5
	Min, Max	2, 4	2, 4	2, 3	1, 4
Washout Period 2 (Day 2 04:00-05:30 PM)	n	5	4	5	4
	Mean (SD)	3.0 (0.71)	3.3 (0.50)	2.8 (0.84)	2.8 (0.96)
	95% CI	2.1, 3.9	2.4, 4.1	1.7, 3.9	1.2, 4.3
	Median	3.0	3.0	3.0	2.5
	Q25, Q75	3.0, 3.0	3.0, 3.5	2.0, 3.0	2.0, 3.5
	Min, Max	2, 4	3, 4	2, 4	2, 4

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Appendix 15.3.3.5

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_coday3.sas

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**Table 15.2.4.11.1.2 Descriptive Statistics of Exhaled CO (ppm) During Days -1, 0, 2 and 4 by Nicotine Level - PK Population**

Timepoint	Statistic	Group-2 PK			
		THS 2.2 Menthol - NRT gum		NRT gum - THS 2.2 Menthol	
		<= 0.6 mg (N=5)	> 0.6 - 1.0 mg (N=4)	<= 0.6 mg (N=5)	> 0.6 - 1.0 mg (N=4)
Washout Period 2 (Day 2 08:00-09:30 PM)	n	5	4	5	4
	Mean (SD)	3.4 (1.52)	3.8 (0.96)	3.0 (1.00)	3.5 (1.74)
	95% CI	1.5, 5.3	2.2, 5.3	1.7, 4.3	0.7, 6.3
	Median	4.0	3.5	3.0	4.0
	Q25, Q75	3.0, 4.0	3.0, 4.5	2.0, 4.0	2.5, 4.5
	Min, Max	1, 5	3, 5	2, 4	1, 5
Day 4/Discharge	n	5	4	5	4
	Mean (SD)	2.6 (0.55)	3.0 (0.82)	2.8 (0.45)	2.8 (0.96)
	95% CI	1.9, 3.3	1.7, 4.3	2.2, 3.4	1.2, 4.3
	Median	3.0	3.0	3.0	2.5
	Q25, Q75	2.0, 3.0	2.5, 3.5	3.0, 3.0	2.0, 3.5
	Min, Max	2, 3	2, 4	2, 3	2, 4

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Appendix 15.3.3.5

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_coday3.sas

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**Table 15.2.4.11.2 Descriptive Statistics of Exhaled CO (ppm) During Days -1, 0, 2 and 4 Categorical Measurements - PK Population**

Timepoint	Statistic	Group-1 PK		Group-2 PK	
		THS 2.2 Menthol - mCC (N=21)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)
Day -1	CO <= 10 ppm - n (%)	7 ( 33.3%)	7 ( 31.8%)	4 ( 44.4%)	2 ( 22.2%)
	CO > 10 ppm - n (%)	14 ( 66.7%)	15 ( 68.2%)	5 ( 55.6%)	7 ( 77.8%)
Washout Period 1 (Day 0 08:00-09:30 AM)	CO <= 10 ppm - n (%)	16 ( 76.2%)	14 ( 63.6%)	8 ( 88.9%)	5 ( 55.6%)
	CO > 10 ppm - n (%)	5 ( 23.8%)	8 ( 36.4%)	1 ( 11.1%)	4 ( 44.4%)
Washout Period 1 (Day 0 12:00-01:30 PM)	CO <= 10 ppm - n (%)	21 (100 %)	20 ( 90.9%)	9 (100 %)	9 (100 %)
	CO > 10 ppm - n (%)	0	2 ( 9.1%)	0	0
Washout Period 1 (Day 0 04:00-05:30 PM)	CO <= 10 ppm - n (%)	21 (100 %)	22 (100 %)	9 (100 %)	9 (100 %)
	CO > 10 ppm - n (%)	0	0	0	0
Washout Period 1 (Day 0 08:00-09:30 PM)	CO <= 10 ppm - n (%)	21 (100 %)	22 (100 %)	9 (100 %)	9 (100 %)
	CO > 10 ppm - n (%)	0	0	0	0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Appendix 15.3.3.5

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_codaycat.sas

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**Table 15.2.4.11.2 Descriptive Statistics of Exhaled CO (ppm) During Days -1, 0, 2 and 4 Categorical Measurements - PK Population**

Timepoint	Statistic	Group-1 PK		Group-2 PK	
		THS 2.2 Menthol - mCC (N=21)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)
Washout Period 2 (Day 2 08:00-09:30 AM)	CO <= 10 ppm - n (%)	21 (100 %)	22 (100 %)	9 (100 %)	9 (100 %)
	CO > 10 ppm - n (%)	0	0	0	0
Washout Period 2 (Day 2 12:00-01:30 PM)	CO <= 10 ppm - n (%)	21 (100 %)	22 (100 %)	9 (100 %)	9 (100 %)
	CO > 10 ppm - n (%)	0	0	0	0
Washout Period 2 (Day 2 04:00-05:30 PM)	CO <= 10 ppm - n (%)	21 (100 %)	22 (100 %)	9 (100 %)	9 (100 %)
	CO > 10 ppm - n (%)	0	0	0	0
Washout Period 2 (Day 2 08:00-09:30 PM)	CO <= 10 ppm - n (%)	21 (100 %)	22 (100 %)	9 (100 %)	9 (100 %)
	CO > 10 ppm - n (%)	0	0	0	0
Day 4/Discharge	CO <= 10 ppm - n (%)	21 (100 %)	22 (100 %)	9 (100 %)	9 (100 %)
	CO > 10 ppm - n (%)	0	0	0	0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Appendix 15.3.3.5

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_codaycat.sas

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**Table 15.2.4.12 Analysis of QSU-brief Questionnaire Factors and Total Score - PK Population**

Variable	Time point	Statistic	Group-1 PK		
			THS 2.2 Menthol (N=43)	mCC (N=43)	THS 2.2 Menthol - mCC
Total Score	Overall	n	43	43	
		Mean (SE)	3.36	3.64	-0.28 (0.255)
		95% CI	2.74, 3.97	3.02, 4.26	-0.79, 0.22
	T <sub>0</sub> + 15 min	n	43	43	
		Mean (SE)	2.85	3.26	-0.41 (0.369)
		95% CI	2.13, 3.57	2.54, 3.98	-1.14, 0.31
	T <sub>0</sub> + 30 min	n	43	43	
		Mean (SE)	2.85	3.17	-0.33 (0.360)
		95% CI	2.13, 3.56	2.46, 3.89	-1.04, 0.38
	T <sub>0</sub> + 45 min	n	43	43	
		Mean (SE)	2.96	3.33	-0.38 (0.332)
		95% CI	2.27, 3.64	2.65, 4.02	-1.03, 0.28

Note: mCC = menthol conventional cigarettes; THS = Tobacco Heating System.

Note: QSU-brief scores reported on a 7-point scale. Higher values indicate greater intensity of urge.

Note: Means and 95% CI are the adjusted least squares means and confidence intervals from an ANOVA model.

Note: T<sub>0</sub> = Time of first product-use at single use day

Note: Comparison overall time points is the main comparison.

Appendix 15.4.4.12 and 15.3.6.11

Path: /cvn/projects/prj/development/000000106326/dev/tables/tlf\_anlqsu.sas

Program Run: 28OCT14 cvn\_ahedge Program Status: FINAL

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**Table 15.2.4.12 Analysis of QSU-brief Questionnaire Factors and Total Score - PK Population**

Variable	Time point	Statistic	Group-1 PK		
			THS 2.2 Menthol (N=43)	mCC (N=43)	THS 2.2 Menthol - mCC
Total Score	T <sub>0</sub> + 60 min	n	43	43	
		Mean (SE)	3.07	3.45	-0.38 (0.326)
		95% CI	2.39, 3.75	2.77, 4.13	-1.02, 0.26
	T <sub>0</sub> + 2 h	n	43	43	
		Mean (SE)	3.36	3.57	-0.21 (0.299)
		95% CI	2.70, 4.01	2.91, 4.23	-0.80, 0.38
	T <sub>0</sub> + 4 h	n	43	43	
		Mean (SE)	3.48	3.71	-0.23 (0.229)
		95% CI	2.88, 4.08	3.11, 4.31	-0.68, 0.23
	T <sub>0</sub> + 6 h	n	43	43	
		Mean (SE)	3.80	4.09	-0.28 (0.178)
		95% CI	3.24, 4.37	3.52, 4.65	-0.63, 0.07

Note: mCC = menthol conventional cigarettes; THS = Tobacco Heating System.

Note: QSU-brief scores reported on a 7-point scale. Higher values indicate greater intensity of urge.

Note: Means and 95% CI are the adjusted least squares means and confidence intervals from an ANOVA model.

Note: T<sub>0</sub> = Time of first product-use at single use day

Note: Comparison overall time points is the main comparison.

Appendix 15.4.4.12 and 15.3.6.11

Path: /cvn/projects/prj/development/000000106326/dev/tables/tlf\_anlqsu.sas

Program Run: 28OCT14 cvn\_ahedge Program Status: FINAL

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**Table 15.2.4.12 Analysis of QSU-brief Questionnaire Factors and Total Score - PK Population**

Variable	Time point	Statistic	Group-1 PK		
			THS 2.2 Menthol (N=43)	mCC (N=43)	THS 2.2 Menthol - mCC
Total Score	T <sub>0</sub> + 9 h	n	43	43	
		Mean (SE)	3.90	4.05	-0.15 (0.183)
		95% CI	3.33, 4.47	3.48, 4.61	-0.51, 0.22
	T <sub>0</sub> + 12 h	n	43	43	
		Mean (SE)	3.94	4.12	-0.19 (0.168)
		95% CI	3.38, 4.50	3.56, 4.68	-0.52, 0.15
Factor 1 - Reward	Overall	n	43	43	
		Mean (SE)	4.14	4.46	-0.32 (0.292)
		95% CI	3.47, 4.81	3.79, 5.13	-0.90, 0.26
	T <sub>0</sub> + 15 min	n	43	43	
		Mean (SE)	3.46	3.89	-0.44 (0.426)
		95% CI	2.66, 4.26	3.10, 4.69	-1.28, 0.40

Note: mCC = menthol conventional cigarettes; THS = Tobacco Heating System.

Note: QSU-brief scores reported on a 7-point scale. Higher values indicate greater intensity of urge.

Note: Means and 95% CI are the adjusted least squares means and confidence intervals from an ANOVA model.

Note: T<sub>0</sub> = Time of first product-use at single use day

Note: Comparison overall time points is the main comparison.

Appendix 15.4.4.12 and 15.3.6.11

Path: /cvn/projects/prj/development/000000106326/dev/tables/tlf\_anlqsu.sas

Program Run: 28OCT14 cvn\_ahedge Program Status: FINAL

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**Table 15.2.4.12 Analysis of QSU-brief Questionnaire Factors and Total Score - PK Population**

Variable	Time point	Statistic	Group-1 PK		
			THS 2.2 Menthol (N=43)	mCC (N=43)	THS 2.2 Menthol - mCC
Factor 1 - Reward	T <sub>0</sub> + 30 min	n	43	43	
		Mean (SE)	3.49	3.87	-0.38 (0.416)
		95% CI	2.70, 4.27	3.08, 4.66	-1.20, 0.44
	T <sub>0</sub> + 45 min	n	43	43	
		Mean (SE)	3.68	4.06	-0.39 (0.374)
		95% CI	2.93, 4.42	3.32, 4.81	-1.12, 0.35
	T <sub>0</sub> + 60 min	n	43	43	
		Mean (SE)	3.77	4.20	-0.43 (0.373)
		95% CI	3.03, 4.52	3.46, 4.95	-1.16, 0.31
	T <sub>0</sub> + 2 h	n	43	43	
		Mean (SE)	4.16	4.44	-0.28 (0.343)
		95% CI	3.44, 4.87	3.72, 5.16	-0.96, 0.39

Note: mCC = menthol conventional cigarettes; THS = Tobacco Heating System.

Note: QSU-brief scores reported on a 7-point scale. Higher values indicate greater intensity of urge.

Note: Means and 95% CI are the adjusted least squares means and confidence intervals from an ANOVA model.

Note: T<sub>0</sub> = Time of first product-use at single use day

Note: Comparison overall time points is the main comparison.

Appendix 15.4.4.12 and 15.3.6.11

Path: /cvn/projects/prj/development/000000106326/dev/tables/tlf\_anlqsu.sas

Program Run: 28OCT14 cvn\_ahedge Program Status: FINAL

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**Table 15.2.4.12 Analysis of QSU-brief Questionnaire Factors and Total Score - PK Population**

Variable	Time point	Statistic	Group-1 PK		
			THS 2.2 Menthol (N=43)	mCC (N=43)	THS 2.2 Menthol - mCC
Factor 1 - Reward	T <sub>0</sub> + 4 h	n	43	43	
		Mean (SE)	4.30	4.56	-0.26 (0.266)
		95% CI	3.65, 4.96	3.91, 5.21	-0.78, 0.27
	T <sub>0</sub> + 6 h	n	43	43	
		Mean (SE)	4.69	5.06	-0.38 (0.209)
		95% CI	4.08, 5.29	4.45, 5.67	-0.79, 0.04
	T <sub>0</sub> + 9 h	n	43	43	
		Mean (SE)	4.83	4.97	-0.15 (0.225)
		95% CI	4.21, 5.44	4.35, 5.59	-0.59, 0.30
	T <sub>0</sub> + 12 h	n	43	43	
		Mean (SE)	4.90	5.09	-0.19 (0.193)
		95% CI	4.30, 5.50	4.49, 5.68	-0.57, 0.20

Note: mCC = menthol conventional cigarettes; THS = Tobacco Heating System.

Note: QSU-brief scores reported on a 7-point scale. Higher values indicate greater intensity of urge.

Note: Means and 95% CI are the adjusted least squares means and confidence intervals from an ANOVA model.

Note: T<sub>0</sub> = Time of first product-use at single use day

Note: Comparison overall time points is the main comparison.

Appendix 15.4.4.12 and 15.3.6.11

Path: /cvn/projects/prj/development/000000106326/dev/tables/tlf\_anlqsu.sas

Program Run: 28OCT14 cvn\_ahedge Program Status: FINAL

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**Table 15.2.4.12 Analysis of QSU-brief Questionnaire Factors and Total Score - PK Population**

Variable	Time point	Statistic	Group-1 PK		
			THS 2.2 Menthol (N=43)	mCC (N=43)	THS 2.2 Menthol - mCC
Factor 2 - Relief	Overall	n	43	43	
		Mean (SE)	2.56	2.80	-0.24 (0.279)
		95% CI	2.02, 3.10	2.26, 3.34	-0.80, 0.31
	T <sub>0</sub> + 15 min	n	43	43	
		Mean (SE)	2.22	2.61	-0.39 (0.229)
		95% CI	1.73, 2.72	2.12, 3.10	-0.84, 0.07
	T <sub>0</sub> + 30 min	n	43	43	
		Mean (SE)	2.19	2.46	-0.27 (0.211)
		95% CI	1.72, 2.67	1.99, 2.94	-0.69, 0.15
	T <sub>0</sub> + 45 min	n	43	43	
		Mean (SE)	2.22	2.59	-0.36 (0.238)
		95% CI	1.72, 2.73	2.09, 3.09	-0.83, 0.11

Note: mCC = menthol conventional cigarettes; THS = Tobacco Heating System.

Note: QSU-brief scores reported on a 7-point scale. Higher values indicate greater intensity of urge.

Note: Means and 95% CI are the adjusted least squares means and confidence intervals from an ANOVA model.

Note: T<sub>0</sub> = Time of first product-use at single use day

Note: Comparison overall time points is the main comparison.

Appendix 15.4.4.12 and 15.3.6.11

Path: /cvn/projects/prj/development/000000106326/dev/tables/tlf\_anlqsu.sas

Program Run: 28OCT14 cvn\_ahedge Program Status: FINAL

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**Table 15.2.4.12 Analysis of QSU-brief Questionnaire Factors and Total Score - PK Population**

Variable	Time point	Statistic	Group-1 PK		
			THS 2.2 Menthol (N=43)	mCC (N=43)	THS 2.2 Menthol - mCC
Factor 2 - Relief	T <sub>0</sub> + 60 min	n	43	43	
		Mean (SE)	2.35	2.68	-0.33 (0.273)
		95% CI	1.82, 2.88	2.15, 3.22	-0.87, 0.21
	T <sub>0</sub> + 2 h	n	43	43	
		Mean (SE)	2.54	2.68	-0.14 (0.308)
		95% CI	1.97, 3.11	2.11, 3.25	-0.75, 0.47
	T <sub>0</sub> + 4 h	n	43	43	
		Mean (SE)	2.64	2.84	-0.20 (0.310)
		95% CI	2.07, 3.22	2.27, 3.41	-0.81, 0.42
	T <sub>0</sub> + 6 h	n	43	43	
		Mean (SE)	2.91	3.09	-0.18 (0.362)
		95% CI	2.28, 3.54	2.46, 3.72	-0.90, 0.53

Note: mCC = menthol conventional cigarettes; THS = Tobacco Heating System.

Note: QSU-brief scores reported on a 7-point scale. Higher values indicate greater intensity of urge.

Note: Means and 95% CI are the adjusted least squares means and confidence intervals from an ANOVA model.

Note: T<sub>0</sub> = Time of first product-use at single use day

Note: Comparison overall time points is the main comparison.

Appendix 15.4.4.12 and 15.3.6.11

Path: /cvn/projects/prj/development/000000106326/dev/tables/tlf\_anlqsu.sas

Program Run: 28OCT14 cvn\_ahedge Program Status: FINAL

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**Table 15.2.4.12 Analysis of QSU-brief Questionnaire Factors and Total Score - PK Population**

Variable	Time point	Statistic	Group-1 PK		
			THS 2.2 Menthol (N=43)	mCC (N=43)	THS 2.2 Menthol - mCC
Factor 2 - Relief	T <sub>0</sub> + 9 h	n	43	43	
		Mean (SE)	2.96	3.10	-0.14 (0.370)
		95% CI	2.32, 3.60	2.46, 3.74	-0.87, 0.59
	T <sub>0</sub> + 12 h	n	43	43	
		Mean (SE)	2.96	3.14	-0.18 (0.375)
		95% CI	2.32, 3.60	2.50, 3.79	-0.92, 0.56

Note: mCC = menthol conventional cigarettes; THS = Tobacco Heating System.

Note: QSU-brief scores reported on a 7-point scale. Higher values indicate greater intensity of urge.

Note: Means and 95% CI are the adjusted least squares means and confidence intervals from an ANOVA model.

Note: T<sub>0</sub> = Time of first product-use at single use day

Note: Comparison overall time points is the main comparison.

Appendix 15.4.4.12 and 15.3.6.11

Path: /cvn/projects/prj/development/000000106326/dev/tables/tlf\_anlqsu.sas

Program Run: 28OCT14 cvn\_ahedge Program Status: FINAL

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**Table 15.2.4.12 Analysis of QSU-brief Questionnaire Factors and Total Score - PK Population**

Variable	Time point	Statistic	Group-2 PK		
			THS 2.2 Menthol (N=18)	NRT gum (N=18)	THS 2.2 Menthol - NRT gum
Total Score	Overall	n	18	18	
		Mean (SE)	3.33	3.66	-0.34 (0.267)
		95% CI	2.35, 4.31	2.69, 4.64	-0.87, 0.19
	T <sub>0</sub> + 15/20 min	n	18	18	
		Mean (SE)	2.94	3.71	-0.76 (0.448)
		95% CI	1.85, 4.04	2.61, 4.80	-1.65, 0.12
	T <sub>0</sub> + 30 min	n	18	18	
		Mean (SE)	2.88	3.60	-0.72 (0.418)
		95% CI	1.80, 3.95	2.52, 4.68	-1.55, 0.10
	T <sub>0</sub> + 45 min	n	18	18	
		Mean (SE)	3.02	3.33	-0.32 (0.317)
		95% CI	2.01, 4.02	2.33, 4.34	-0.94, 0.31

Note: NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Note: QSU-brief scores reported on a 7-point scale. Higher values indicate greater intensity of urge.

Note: Means and 95% CI are the adjusted least squares means and confidence intervals from an ANOVA model.

Note: T<sub>0</sub> = Time of first product-use at single use day

Note: In Group-2 PK, the QSU-b values at T<sub>0</sub> + 15/20 min refer to QSU-brief data at time T<sub>0</sub> + 15 min for THS 2.2 Menthol, and to QSU-brief data at time T<sub>0</sub> + 20 min for NRT gum

Note: Comparison overall time points is the main comparison.

Appendix 15.4.4.12 and 15.3.6.11

Path: /cvn/projects/prj/development/000000106326/dev/tables/tlf\_anlqsu.sas

Program Run: 28OCT14 cvn\_ahedge Program Status: FINAL

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**Table 15.2.4.12 Analysis of QSU-brief Questionnaire Factors and Total Score - PK Population**

Variable	Time point	Statistic	Group-2 PK		
			THS 2.2 Menthol (N=18)	NRT gum (N=18)	THS 2.2 Menthol - NRT gum
Total Score	T <sub>0</sub> + 60 min	n	18	18	
		Mean (SE)	2.90	3.50	-0.60 (0.338)
		95% CI	1.88, 3.92	2.48, 4.52	-1.27, 0.07
	T <sub>0</sub> + 2 h	n	18	18	
		Mean (SE)	3.37	3.67	-0.30 (0.276)
		95% CI	2.38, 4.35	2.68, 4.65	-0.85, 0.25
	T <sub>0</sub> + 4 h	n	18	18	
		Mean (SE)	3.49	3.72	-0.23 (0.241)
		95% CI	2.53, 4.46	2.76, 4.69	-0.71, 0.25
	T <sub>0</sub> + 6 h	n	18	18	
		Mean (SE)	3.85	3.82	0.03 (0.224)
		95% CI	2.89, 4.81	2.86, 4.77	-0.41, 0.48

Note: NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Note: QSU-brief scores reported on a 7-point scale. Higher values indicate greater intensity of urge.

Note: Means and 95% CI are the adjusted least squares means and confidence intervals from an ANOVA model.

Note: T<sub>0</sub> = Time of first product-use at single use day

Note: In Group-2 PK, the QSU-b values at T<sub>0</sub> + 15/20 min refer to QSU-brief data at time T<sub>0</sub> + 15 min for THS 2.2 Menthol, and to QSU-brief data at time T<sub>0</sub> + 20 min for NRT gum

Note: Comparison overall time points is the main comparison.

Appendix 15.4.4.12 and 15.3.6.11

Path: /cvn/projects/prj/development/000000106326/dev/tables/tlf\_anlqsu.sas

Program Run: 28OCT14 cvn\_ahedge Program Status: FINAL

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**Table 15.2.4.12 Analysis of QSU-brief Questionnaire Factors and Total Score - PK Population**

Variable	Time point	Statistic	Group-2 PK		
			THS 2.2 Menthol (N=18)	NRT gum (N=18)	THS 2.2 Menthol - NRT gum
Total Score	T <sub>0</sub> + 9 h	n	18	18	
		Mean (SE)	3.71	3.77	-0.07 (0.224)
		95% CI	2.75, 4.66	2.81, 4.73	-0.51, 0.38
	T <sub>0</sub> + 12 h	n	18	18	
		Mean (SE)	3.79	3.86	-0.07 (0.189)
		95% CI	2.85, 4.73	2.91, 4.80	-0.44, 0.31
Factor 1 - Reward	Overall	n	18	18	
		Mean (SE)	4.09	4.38	-0.29 (0.364)
		95% CI	2.97, 5.21	3.26, 5.50	-1.01, 0.43
	T <sub>0</sub> + 15/20 min	n	18	18	
		Mean (SE)	3.61	4.43	-0.82 (0.568)
		95% CI	2.34, 4.88	3.16, 5.71	-1.94, 0.30

Note: NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Note: QSU-brief scores reported on a 7-point scale. Higher values indicate greater intensity of urge.

Note: Means and 95% CI are the adjusted least squares means and confidence intervals from an ANOVA model.

Note: T<sub>0</sub> = Time of first product-use at single use day

Note: In Group-2 PK, the QSU-b values at T<sub>0</sub> + 15/20 min refer to QSU-brief data at time T<sub>0</sub> + 15 min for THS 2.2 Menthol, and to QSU-brief data at time T<sub>0</sub> + 20 min for NRT gum

Note: Comparison overall time points is the main comparison.

Appendix 15.4.4.12 and 15.3.6.11

Path: /cvn/projects/prj/development/000000106326/dev/tables/tlf\_anlqsu.sas

Program Run: 28OCT14 cvn\_ahedge Program Status: FINAL

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**Table 15.2.4.12 Analysis of QSU-brief Questionnaire Factors and Total Score - PK Population**

Variable	Time point	Statistic	Group-2 PK		
			THS 2.2 Menthol (N=18)	NRT gum (N=18)	THS 2.2 Menthol - NRT gum
Factor 1 - Reward	T <sub>0</sub> + 30 min	n	18	18	
		Mean (SE)	3.56	4.28	-0.72 (0.555)
		95% CI	2.29, 4.82	3.02, 5.54	-1.82, 0.37
	T <sub>0</sub> + 45 min	n	18	18	
		Mean (SE)	3.61	3.93	-0.32 (0.431)
		95% CI	2.45, 4.77	2.77, 5.10	-1.17, 0.53
	T <sub>0</sub> + 60 min	n	18	18	
		Mean (SE)	3.66	4.06	-0.40 (0.437)
		95% CI	2.49, 4.82	2.89, 5.22	-1.27, 0.47
	T <sub>0</sub> + 2 h	n	18	18	
		Mean (SE)	4.04	4.38	-0.33 (0.398)
		95% CI	2.90, 5.18	3.24, 5.52	-1.12, 0.45

Note: NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Note: QSU-brief scores reported on a 7-point scale. Higher values indicate greater intensity of urge.

Note: Means and 95% CI are the adjusted least squares means and confidence intervals from an ANOVA model.

Note: T<sub>0</sub> = Time of first product-use at single use day

Note: In Group-2 PK, the QSU-b values at T<sub>0</sub> + 15/20 min refer to QSU-brief data at time T<sub>0</sub> + 15 min for THS 2.2 Menthol, and to QSU-brief data at time T<sub>0</sub> + 20 min for NRT gum

Note: Comparison overall time points is the main comparison.

Appendix 15.4.4.12 and 15.3.6.11

Path: /cvn/projects/prj/development/000000106326/dev/tables/tlf\_anlqsu.sas

Program Run: 28OCT14 cvn\_ahedge Program Status: FINAL

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**Table 15.2.4.12 Analysis of QSU-brief Questionnaire Factors and Total Score - PK Population**

Variable	Time point	Statistic	Group-2 PK		
			THS 2.2 Menthol (N=18)	NRT gum (N=18)	THS 2.2 Menthol - NRT gum
Factor 1 - Reward	T <sub>0</sub> + 4 h	n	18	18	
		Mean (SE)	4.37	4.46	-0.09 (0.330)
		95% CI	3.27, 5.46	3.36, 5.55	-0.74, 0.56
	T <sub>0</sub> + 6 h	n	18	18	
		Mean (SE)	4.80	4.61	0.19 (0.321)
		95% CI	3.71, 5.89	3.52, 5.70	-0.45, 0.82
	T <sub>0</sub> + 9 h	n	18	18	
		Mean (SE)	4.52	4.60	-0.08 (0.302)
		95% CI	3.44, 5.60	3.52, 5.68	-0.68, 0.52
	T <sub>0</sub> + 12 h	n	18	18	
		Mean (SE)	4.66	4.68	-0.02 (0.241)
		95% CI	3.60, 5.71	3.63, 5.73	-0.50, 0.46

Note: NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Note: QSU-brief scores reported on a 7-point scale. Higher values indicate greater intensity of urge.

Note: Means and 95% CI are the adjusted least squares means and confidence intervals from an ANOVA model.

Note: T<sub>0</sub> = Time of first product-use at single use day

Note: In Group-2 PK, the QSU-b values at T<sub>0</sub> + 15/20 min refer to QSU-brief data at time T<sub>0</sub> + 15 min for THS 2.2 Menthol, and to QSU-brief data at time T<sub>0</sub> + 20 min for NRT gum

Note: Comparison overall time points is the main comparison.

Appendix 15.4.4.12 and 15.3.6.11

Path: /cvn/projects/prj/development/000000106326/dev/tables/tlf\_anlqsu.sas

Program Run: 28OCT14 cvn\_ahedge Program Status: FINAL

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**Table 15.2.4.12 Analysis of QSU-brief Questionnaire Factors and Total Score - PK Population**

Variable	Time point	Statistic	Group-2 PK		
			THS 2.2 Menthol (N=18)	NRT gum (N=18)	THS 2.2 Menthol - NRT gum
Factor 2 - Relief	Overall	n	18	18	
		Mean (SE)	2.56	2.95	-0.38 (0.165)
		95% CI	1.66, 3.47	2.04, 3.85	-0.71, -0.05
	T <sub>0</sub> + 15/20 min	n	18	18	
		Mean (SE)	2.28	2.98	-0.70 (0.349)
		95% CI	1.28, 3.28	1.98, 3.98	-1.39, -0.01
	T <sub>0</sub> + 30 min	n	18	18	
		Mean (SE)	2.20	2.92	-0.72 (0.290)
		95% CI	1.24, 3.16	1.96, 3.88	-1.30, -0.15
	T <sub>0</sub> + 45 min	n	18	18	
		Mean (SE)	2.42	2.73	-0.31 (0.183)
		95% CI	1.51, 3.33	1.82, 3.64	-0.68, 0.05

Note: NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Note: QSU-brief scores reported on a 7-point scale. Higher values indicate greater intensity of urge.

Note: Means and 95% CI are the adjusted least squares means and confidence intervals from an ANOVA model.

Note: T<sub>0</sub> = Time of first product-use at single use day

Note: In Group-2 PK, the QSU-b values at T<sub>0</sub> + 15/20 min refer to QSU-brief data at time T<sub>0</sub> + 15 min for THS 2.2 Menthol, and to QSU-brief data at time T<sub>0</sub> + 20 min for NRT gum

Note: Comparison overall time points is the main comparison.

Appendix 15.4.4.12 and 15.3.6.11

Path: /cvn/projects/prj/development/000000106326/dev/tables/tlf\_anlqsu.sas

Program Run: 28OCT14 cvn\_ahedge Program Status: FINAL

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**Table 15.2.4.12 Analysis of QSU-brief Questionnaire Factors and Total Score - PK Population**

Variable	Time point	Statistic	Group-2 PK		
			THS 2.2 Menthol (N=18)	NRT gum (N=18)	THS 2.2 Menthol - NRT gum
Factor 2 - Relief	T <sub>0</sub> + 60 min	n	18	18	
		Mean (SE)	2.14	2.94	-0.80 (0.257)
		95% CI	1.20, 3.09	2.00, 3.89	-1.31, -0.29
	T <sub>0</sub> + 2 h	n	18	18	
		Mean (SE)	2.69	2.96	-0.27 (0.145)
		95% CI	1.79, 3.58	2.06, 3.85	-0.56, 0.02
	T <sub>0</sub> + 4 h	n	18	18	
		Mean (SE)	2.62	2.99	-0.37 (0.152)
		95% CI	1.72, 3.52	2.09, 3.89	-0.67, -0.06
	T <sub>0</sub> + 6 h	n	18	18	
		Mean (SE)	2.90	3.02	-0.12 (0.146)
		95% CI	2.00, 3.80	2.13, 3.92	-0.41, 0.17

Note: NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Note: QSU-brief scores reported on a 7-point scale. Higher values indicate greater intensity of urge.

Note: Means and 95% CI are the adjusted least squares means and confidence intervals from an ANOVA model.

Note: T<sub>0</sub> = Time of first product-use at single use day

Note: In Group-2 PK, the QSU-b values at T<sub>0</sub> + 15/20 min refer to QSU-brief data at time T<sub>0</sub> + 15 min for THS 2.2 Menthol, and to QSU-brief data at time T<sub>0</sub> + 20 min for NRT gum

Note: Comparison overall time points is the main comparison.

Appendix 15.4.4.12 and 15.3.6.11

Path: /cvn/projects/prj/development/000000106326/dev/tables/tlf\_anlqsu.sas

Program Run: 28OCT14 cvn\_ahedge Program Status: FINAL

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**Table 15.2.4.12 Analysis of QSU-brief Questionnaire Factors and Total Score - PK Population**

Variable	Time point	Statistic	Group-2 PK		
			THS 2.2 Menthol (N=18)	NRT gum (N=18)	THS 2.2 Menthol - NRT gum
Factor 2 - Relief	T <sub>0</sub> + 9 h	n	18	18	
		Mean (SE)	2.89	2.94	-0.06 (0.204)
		95% CI	1.97, 3.81	2.03, 3.86	-0.46, 0.35
	T <sub>0</sub> + 12 h	n	18	18	
		Mean (SE)	2.92	3.03	-0.11 (0.204)
		95% CI	2.00, 3.84	2.12, 3.95	-0.52, 0.29

Note: NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Note: QSU-brief scores reported on a 7-point scale. Higher values indicate greater intensity of urge.

Note: Means and 95% CI are the adjusted least squares means and confidence intervals from an ANOVA model.

Note: T<sub>0</sub> = Time of first product-use at single use day

Note: In Group-2 PK, the QSU-b values at T<sub>0</sub> + 15/20 min refer to QSU-brief data at time T<sub>0</sub> + 15 min for THS 2.2 Menthol, and to QSU-brief data at time T<sub>0</sub> + 20 min for NRT gum

Note: Comparison overall time points is the main comparison.

Appendix 15.4.4.12 and 15.3.6.11

Path: /cvn/projects/prj/development/000000106326/dev/tables/tlf\_anlqsu.sas

Program Run: 28OCT14 cvn\_ahedge Program Status: FINAL

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**Table 15.2.4.13 Analysis of QSU-brief Questionnaire Factors and Total Score by Bootstrapping Techniques - PK Population**

Variable	Timepoint	Statistic	Group-1 PK		
			THS 2.2 Menthol (N=43)	mCC (N=43)	THS 2.2 Menthol - mCC
Total Score	Overall	n	43	43	
		Mean (SE)	3.35	3.63	-0.28 (0.256)
		95% CI	2.97, 3.73	3.26, 4.00	-0.47, -0.10
		Median	3.35	3.63	-0.29
Factor 1 - Reward	Overall	n	43	43	
		Mean (SE)	4.13	4.45	-0.32 (0.289)
		95% CI	3.60, 4.65	3.96, 4.90	-0.62, -0.02
		Median	4.13	4.45	-0.32
Factor 2 - Relief	Overall	n	43	43	
		Mean (SE)	2.56	2.81	-0.25 (0.211)
		95% CI	NC	NC	NC
		Median	2.56	2.81	-0.25

Note: mCC = menthol conventional cigarettes; THS = Tobacco Heating System.

Note: Higher scores indicate a higher urge to smoke.

Note: Point and 95% interval estimates by means of the percentile bootstrap technique of the adjusted least squares means from an ANOVA model.

Note: NC = Not calculable, due to greater than 50 bootstrap samples failing to converge.

Appendix 15.4.4.13 and 15.3.6.11

Path: /cvn/projects/prj/development/000000106326/dev/tables/tl\_banlqsu.sas

Program Run: 28OCT14 cvn\_ahedge Program Status: FINAL

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**Table 15.2.4.13 Analysis of QSU-brief Questionnaire Factors and Total Score by Bootstrapping Techniques - PK Population**

Variable	Timepoint	Statistic	Group-2 PK		
			THS 2.2 Menthol (N=18)	NRT gum (N=18)	THS 2.2 Menthol - NRT gum
Total Score	Overall	n	18	18	
		Mean (SE)	3.32	3.65	-0.34 (0.220)
		95% CI	NC	NC	NC
		Median	3.31	3.65	-0.33
Factor 1 - Reward	Overall	n	18	18	
		Mean (SE)	4.08	4.36	-0.29 (0.283)
		95% CI	NC	NC	NC
		Median	4.06	4.36	-0.29
Factor 2 - Relief	Overall	n	18	18	
		Mean (SE)	2.56	2.94	-0.38 (0.173)
		95% CI	NC	NC	NC
		Median	2.54	2.93	-0.37

Note: NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Note: Higher scores indicate a higher urge to smoke.

Note: Point and 95% interval estimates by means of the percentile bootstrap technique of the adjusted least squares means from an ANOVA model.

Note: NC = Not calculable, due to greater than 50 bootstrap samples failing to converge.

Appendix 15.4.4.13 and 15.3.6.11

Path: /cvn/projects/prj/development/000000106326/dev/tables/tl\_banlqsu.sas

Program Run: 28OCT14 cvn\_ahedge Program Status: FINAL

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**Table 15.2.4.14 Descriptive Statistics of QSU-brief Questionnaire Factors and Total Score - PK Population**

Variable	Timepoint	Statistic	Group-1 PK		Group-2 PK	
			THS 2.2 Menthol (N=43)	mCC (N=43)	THS 2.2 Menthol (N=18)	NRT gum (N=18)
Factor 1 - Reward	15 min < T <sub>0</sub>	n	43	43	18	18
		Mean (SD)	5.37 (1.109)	5.40 (1.091)	4.81 (2.133)	5.18 (1.548)
		95% CI	5.02, 5.71	5.05, 5.74	3.75, 5.88	4.40, 5.95
		Median	5.60	5.60	5.40	5.40
		Q25, Q75	4.80, 6.20	4.80, 6.40	2.40, 6.80	4.00, 6.60
		Min, Max	2.2, 7.0	2.0, 7.0	1.4, 7.0	2.0, 7.0
	T <sub>0</sub> + 15/20 min	n	43	43	18	18
		Mean (SD)	3.45 (1.696)	3.89 (1.661)	3.61 (2.188)	4.43 (2.181)
		95% CI	2.92, 3.98	3.38, 4.41	2.52, 4.70	3.34, 5.52
		Median	3.00	3.80	3.50	4.60
		Q25, Q75	2.20, 5.00	2.80, 5.40	1.40, 5.40	2.20, 6.60
		Min, Max	1.0, 7.0	1.0, 6.6	1.0, 7.0	1.0, 7.0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: The QSU-brief values at T<sub>0</sub> + 15/20 min refer to QSU-brief data at time T<sub>0</sub> + 15 min for THS 2.2 Menthol and mCC, and to QSU-brief data at time T<sub>0</sub> + 20 min for NRT gum.

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/000000106326/dev/tables/t\_qsu.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.14 Descriptive Statistics of QSU-brief Questionnaire Factors and Total Score - PK Population**

Variable	Timepoint	Statistic	Group-1 PK		Group-2 PK	
			THS 2.2 Menthol (N=43)	mCC (N=43)	THS 2.2 Menthol (N=18)	NRT gum (N=18)
Factor 1 - Reward	T <sub>0</sub> + 30 min	n	43	43	18	18
		Mean (SD)	3.48 (1.815)	3.87 (1.661)	3.56 (2.250)	4.28 (2.336)
		95% CI	2.92, 4.04	3.35, 4.38	2.43, 4.68	3.11, 5.44
		Median	3.20	3.80	2.80	4.30
		Q25, Q75	2.00, 5.20	2.60, 5.40	1.80, 6.00	2.20, 7.00
		Min, Max	1.0, 7.0	1.0, 6.6	1.0, 7.0	1.0, 7.0
	T <sub>0</sub> + 45 min	n	43	43	18	18
		Mean (SD)	3.67 (1.738)	4.06 (1.600)	3.61 (2.183)	3.93 (2.354)
		95% CI	3.13, 4.21	3.56, 4.56	2.52, 4.70	2.76, 5.11
		Median	3.40	4.00	3.20	3.80
		Q25, Q75	2.20, 5.40	3.20, 5.40	1.60, 5.40	2.00, 6.40
		Min, Max	1.0, 6.6	1.0, 6.6	1.0, 7.0	1.0, 7.0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: The QSU-brief values at T<sub>0</sub> + 15/20 min refer to QSU-brief data at time T<sub>0</sub> + 15 min for THS 2.2 Menthol and mCC, and to QSU-brief data at time T<sub>0</sub> + 20 min for NRT gum.

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/000000106326/dev/tables/t\_qsu.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.14 Descriptive Statistics of QSU-brief Questionnaire Factors and Total Score - PK Population**

Variable	Timepoint	Statistic	Group-1 PK		Group-2 PK	
			THS 2.2 Menthol (N=43)	mCC (N=43)	THS 2.2 Menthol (N=18)	NRT gum (N=18)
Factor 1 - Reward	T <sub>0</sub> + 60 min	n	43	43	18	18
		Mean (SD)	3.77 (1.820)	4.20 (1.680)	3.66 (2.126)	4.06 (2.477)
		95% CI	3.20, 4.33	3.68, 4.72	2.59, 4.72	2.82, 5.29
		Median	4.00	4.20	2.70	3.70
		Q25, Q75	2.00, 5.60	3.00, 5.60	1.80, 5.40	2.00, 6.60
		Min, Max	1.0, 6.8	1.0, 7.0	1.0, 7.0	1.0, 7.0
	T <sub>0</sub> + 2 h	n	43	43	18	18
		Mean (SD)	4.15 (1.794)	4.44 (1.528)	4.04 (2.300)	4.38 (2.182)
		95% CI	3.59, 4.71	3.96, 4.91	2.90, 5.19	3.29, 5.47
		Median	4.40	4.40	3.70	5.00
		Q25, Q75	2.60, 5.60	3.60, 5.40	2.00, 6.80	2.80, 6.40
		Min, Max	1.0, 7.0	1.0, 6.8	1.0, 7.0	1.0, 7.0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: The QSU-brief values at T<sub>0</sub> + 15/20 min refer to QSU-brief data at time T<sub>0</sub> + 15 min for THS 2.2 Menthol and mCC, and to QSU-brief data at time T<sub>0</sub> + 20 min for NRT gum.

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/000000106326/dev/tables/t\_qsu.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.14 Descriptive Statistics of QSU-brief Questionnaire Factors and Total Score - PK Population**

Variable	Timepoint	Statistic	Group-1 PK		Group-2 PK	
			THS 2.2 Menthol (N=43)	mCC (N=43)	THS 2.2 Menthol (N=18)	NRT gum (N=18)
Factor 1 - Reward	T <sub>0</sub> + 4 h	n	43	43	18	18
		Mean (SD)	4.30 (1.557)	4.56 (1.523)	4.37 (2.152)	4.46 (2.326)
		95% CI	3.81, 4.78	4.08, 5.03	3.29, 5.44	3.29, 5.62
		Median	4.20	4.60	4.50	5.60
		Q25, Q75	3.60, 5.60	3.40, 5.80	2.80, 6.80	2.00, 6.80
		Min, Max	1.0, 7.0	1.0, 6.6	1.0, 7.0	1.0, 7.0
	T <sub>0</sub> + 6 h	n	43	43	18	18
		Mean (SD)	4.68 (1.658)	5.06 (1.526)	4.80 (2.073)	4.61 (2.239)
		95% CI	4.16, 5.19	4.59, 5.54	3.76, 5.84	3.49, 5.73
		Median	4.80	5.00	5.50	5.20
		Q25, Q75	3.60, 5.80	4.40, 6.20	3.20, 6.80	2.80, 6.80
		Min, Max	1.0, 7.0	1.0, 7.0	1.0, 7.0	1.0, 7.0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: The QSU-brief values at T<sub>0</sub> + 15/20 min refer to QSU-brief data at time T<sub>0</sub> + 15 min for THS 2.2 Menthol and mCC, and to QSU-brief data at time T<sub>0</sub> + 20 min for NRT gum.

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/000000106326/dev/tables/t\_qsu.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.14 Descriptive Statistics of QSU-brief Questionnaire Factors and Total Score - PK Population**

Variable	Timepoint	Statistic	Group-1 PK		Group-2 PK	
			THS 2.2 Menthol (N=43)	mCC (N=43)	THS 2.2 Menthol (N=18)	NRT gum (N=18)
Factor 1 - Reward	T <sub>0</sub> + 9 h	n	43	43	18	18
		Mean (SD)	4.82 (1.538)	4.97 (1.437)	4.52 (2.123)	4.60 (2.149)
		95% CI	4.34, 5.30	4.52, 5.42	3.46, 5.58	3.53, 5.67
		Median	5.20	5.20	4.30	4.80
		Q25, Q75	4.00, 6.00	4.40, 6.00	3.00, 7.00	2.80, 6.80
		Min, Max	1.0, 6.8	1.0, 7.0	1.0, 7.0	1.0, 7.0
	T <sub>0</sub> + 12 h	n	43	43	18	18
		Mean (SD)	4.89 (1.595)	5.08 (1.446)	4.66 (2.067)	4.68 (2.198)
		95% CI	4.40, 5.39	4.63, 5.53	3.62, 5.69	3.58, 5.78
		Median	5.20	5.20	4.90	5.40
		Q25, Q75	4.00, 6.00	4.40, 6.20	3.20, 6.80	2.60, 6.80
		Min, Max	1.0, 7.0	1.4, 7.0	1.0, 7.0	1.2, 7.0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: The QSU-brief values at T<sub>0</sub> + 15/20 min refer to QSU-brief data at time T<sub>0</sub> + 15 min for THS 2.2 Menthol and mCC, and to QSU-brief data at time T<sub>0</sub> + 20 min for NRT gum.

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/000000106326/dev/tables/t\_qsu.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.14 Descriptive Statistics of QSU-brief Questionnaire Factors and Total Score - PK Population**

Variable	Timepoint	Statistic	Group-1 PK		Group-2 PK	
			THS 2.2 Menthol (N=43)	mCC (N=43)	THS 2.2 Menthol (N=18)	NRT gum (N=18)
Factor 2 - Relief	15 min < T <sub>0</sub>	n	43	43	18	18
		Mean (SD)	3.35 (1.217)	3.51 (1.328)	3.16 (1.856)	3.36 (1.723)
		95% CI	2.97, 3.73	3.09, 3.92	2.23, 4.08	2.49, 4.22
		Median	3.40	3.40	3.10	3.00
		Q25, Q75	2.60, 4.40	2.40, 4.60	1.20, 4.60	2.00, 4.60
		Min, Max	1.0, 5.6	1.0, 6.0	1.0, 6.4	1.0, 7.0
	T <sub>0</sub> + 15/20 min	n	43	43	18	18
		Mean (SD)	2.23 (1.088)	2.62 (1.187)	2.28 (1.751)	2.98 (1.737)
		95% CI	1.89, 2.57	2.25, 2.99	1.40, 3.15	2.11, 3.85
		Median	2.00	2.40	1.30	2.60
		Q25, Q75	1.40, 3.00	1.60, 3.60	1.00, 4.00	1.40, 4.20
		Min, Max	1.0, 4.8	1.0, 4.8	1.0, 6.2	1.0, 6.4

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: The QSU-brief values at T<sub>0</sub> + 15/20 min refer to QSU-brief data at time T<sub>0</sub> + 15 min for THS 2.2 Menthol and mCC, and to QSU-brief data at time T<sub>0</sub> + 20 min for NRT gum.

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/000000106326/dev/tables/t\_qsu.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.14 Descriptive Statistics of QSU-brief Questionnaire Factors and Total Score - PK Population**

Variable	Timepoint	Statistic	Group-1 PK		Group-2 PK	
			THS 2.2 Menthol (N=43)	mCC (N=43)	THS 2.2 Menthol (N=18)	NRT gum (N=18)
Factor 2 - Relief	T <sub>0</sub> + 30 min	n	43	43	18	18
		Mean (SD)	2.20 (1.118)	2.47 (1.144)	2.20 (1.641)	2.92 (1.845)
		95% CI	1.85, 2.55	2.12, 2.83	1.38, 3.02	2.00, 3.84
		Median	2.00	2.20	1.30	2.70
		Q25, Q75	1.00, 3.00	1.60, 3.00	1.00, 3.00	1.00, 4.00
		Min, Max	1.0, 5.0	1.0, 5.2	1.0, 6.0	1.0, 6.8
	T <sub>0</sub> + 45 min	n	43	43	18	18
		Mean (SD)	2.23 (1.098)	2.60 (1.226)	2.42 (1.736)	2.73 (1.821)
		95% CI	1.89, 2.58	2.22, 2.98	1.55, 3.29	1.82, 3.64
		Median	2.20	2.40	1.80	2.20
		Q25, Q75	1.00, 3.00	1.80, 3.20	1.00, 3.40	1.00, 4.00
		Min, Max	1.0, 5.2	1.0, 5.8	1.0, 6.4	1.0, 6.4

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: The QSU-brief values at T<sub>0</sub> + 15/20 min refer to QSU-brief data at time T<sub>0</sub> + 15 min for THS 2.2 Menthol and mCC, and to QSU-brief data at time T<sub>0</sub> + 20 min for NRT gum.

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/000000106326/dev/tables/t\_qsu.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.14 Descriptive Statistics of QSU-brief Questionnaire Factors and Total Score - PK Population**

Variable	Timepoint	Statistic	Group-1 PK		Group-2 PK	
			THS 2.2 Menthol (N=43)	mCC (N=43)	THS 2.2 Menthol (N=18)	NRT gum (N=18)
Factor 2 - Relief	T <sub>0</sub> + 60 min	n	43	43	18	18
		Mean (SD)	2.36 (1.267)	2.69 (1.269)	2.14 (1.534)	2.94 (1.944)
		95% CI	1.96, 2.75	2.30, 3.09	1.38, 2.91	1.97, 3.92
		Median	2.20	2.40	1.60	2.60
		Q25, Q75	1.00, 3.00	1.80, 3.80	1.00, 2.40	1.00, 4.00
		Min, Max	1.0, 5.8	1.0, 5.4	1.0, 6.4	1.0, 6.4
	T <sub>0</sub> + 2 h	n	43	43	18	18
		Mean (SD)	2.55 (1.274)	2.69 (1.206)	2.69 (1.782)	2.96 (1.838)
		95% CI	2.15, 2.95	2.32, 3.07	1.80, 3.58	2.04, 3.87
		Median	2.40	2.60	2.20	2.80
		Q25, Q75	1.40, 3.40	1.80, 3.60	1.20, 3.80	1.20, 4.40
		Min, Max	1.0, 5.2	1.0, 4.8	1.0, 6.4	1.0, 6.4

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: The QSU-brief values at T<sub>0</sub> + 15/20 min refer to QSU-brief data at time T<sub>0</sub> + 15 min for THS 2.2 Menthol and mCC, and to QSU-brief data at time T<sub>0</sub> + 20 min for NRT gum.

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/000000106326/dev/tables/t\_qsu.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.14 Descriptive Statistics of QSU-brief Questionnaire Factors and Total Score - PK Population**

Variable	Timepoint	Statistic	Group-1 PK		Group-2 PK	
			THS 2.2 Menthol (N=43)	mCC (N=43)	THS 2.2 Menthol (N=18)	NRT gum (N=18)
Factor 2 - Relief	T <sub>0</sub> + 4 h	n	43	43	18	18
		Mean (SD)	2.65 (1.149)	2.85 (1.245)	2.62 (1.728)	2.99 (1.882)
		95% CI	2.29, 3.01	2.46, 3.24	1.76, 3.49	2.05, 3.93
		Median	2.80	2.80	2.30	2.80
		Q25, Q75	1.60, 3.20	1.80, 4.00	1.20, 3.00	1.20, 4.60
		Min, Max	1.0, 5.4	1.0, 5.0	1.0, 6.2	1.0, 6.6
	T <sub>0</sub> + 6 h	n	43	43	18	18
		Mean (SD)	2.92 (1.385)	3.10 (1.424)	2.90 (1.762)	3.02 (1.926)
		95% CI	2.49, 3.35	2.66, 3.55	2.02, 3.78	2.06, 3.98
		Median	3.00	3.00	2.40	2.50
		Q25, Q75	2.00, 4.00	2.20, 4.20	1.20, 3.80	1.20, 4.40
		Min, Max	1.0, 6.2	1.0, 6.8	1.0, 6.4	1.0, 6.4

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: The QSU-brief values at T<sub>0</sub> + 15/20 min refer to QSU-brief data at time T<sub>0</sub> + 15 min for THS 2.2 Menthol and mCC, and to QSU-brief data at time T<sub>0</sub> + 20 min for NRT gum.

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/000000106326/dev/tables/t\_qsu.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.14 Descriptive Statistics of QSU-brief Questionnaire Factors and Total Score - PK Population**

Variable	Timepoint	Statistic	Group-1 PK		Group-2 PK	
			THS 2.2 Menthol (N=43)	mCC (N=43)	THS 2.2 Menthol (N=18)	NRT gum (N=18)
Factor 2 - Relief	T <sub>0</sub> + 9 h	n	43	43	18	18
		Mean (SD)	2.97 (1.362)	3.11 (1.363)	2.89 (1.800)	2.94 (1.798)
		95% CI	2.54, 3.39	2.69, 3.54	1.99, 3.79	2.05, 3.84
		Median	2.80	3.40	2.70	2.50
		Q25, Q75	1.80, 4.00	2.20, 4.20	1.20, 3.40	1.20, 4.40
		Min, Max	1.0, 6.0	1.0, 6.0	1.0, 7.0	1.0, 6.4
	T <sub>0</sub> + 12 h	n	43	43	18	18
		Mean (SD)	2.97 (1.411)	3.15 (1.396)	2.92 (1.827)	3.03 (1.915)
		95% CI	2.53, 3.41	2.72, 3.59	2.01, 3.84	2.08, 3.99
		Median	2.80	3.20	2.80	2.80
		Q25, Q75	1.80, 4.00	2.20, 4.20	1.00, 4.20	1.20, 4.60
		Min, Max	1.0, 6.4	1.0, 6.4	1.0, 6.6	1.0, 6.6

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: The QSU-brief values at T<sub>0</sub> + 15/20 min refer to QSU-brief data at time T<sub>0</sub> + 15 min for THS 2.2 Menthol and mCC, and to QSU-brief data at time T<sub>0</sub> + 20 min for NRT gum.

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/000000106326/dev/tables/t\_qsu.sas

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**Table 15.2.4.14 Descriptive Statistics of QSU-brief Questionnaire Factors and Total Score - PK Population**

Variable	Timepoint	Statistic	Group-1 PK		Group-2 PK	
			THS 2.2 Menthol (N=43)	mCC (N=43)	THS 2.2 Menthol (N=18)	NRT gum (N=18)
Total score	15 min < T <sub>0</sub>	n	43	43	18	18
		Mean (SD)	4.36 (1.060)	4.45 (1.131)	3.98 (1.893)	4.27 (1.531)
		95% CI	4.03, 4.69	4.10, 4.80	3.04, 4.93	3.50, 5.03
		Median	4.50	4.50	4.60	4.40
		Q25, Q75	3.70, 5.20	3.70, 5.30	1.70, 5.60	2.80, 5.40
		Min, Max	1.6, 6.3	1.5, 6.3	1.2, 6.7	1.9, 7.0
	T <sub>0</sub> + 15/20 min	n	43	43	18	18
		Mean (SD)	2.84 (1.296)	3.26 (1.366)	2.94 (1.885)	3.71 (1.818)
		95% CI	2.44, 3.25	2.83, 3.68	2.00, 3.89	2.80, 4.61
		Median	2.80	3.30	2.35	4.35
		Q25, Q75	1.70, 3.80	2.20, 4.40	1.20, 4.50	1.80, 4.80
		Min, Max	1.0, 5.6	1.0, 5.5	1.0, 6.6	1.0, 6.6

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: The QSU-brief values at T<sub>0</sub> + 15/20 min refer to QSU-brief data at time T<sub>0</sub> + 15 min for THS 2.2 Menthol and mCC, and to QSU-brief data at time T<sub>0</sub> + 20 min for NRT gum.

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/000000106326/dev/tables/t\_qsu.sas

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**Table 15.2.4.14 Descriptive Statistics of QSU-brief Questionnaire Factors and Total Score - PK Population**

Variable	Timepoint	Statistic	Group-1 PK		Group-2 PK	
			THS 2.2 Menthol (N=43)	mCC (N=43)	THS 2.2 Menthol (N=18)	NRT gum (N=18)
Total score	T <sub>0</sub> + 30 min	n	43	43	18	18
		Mean (SD)	2.84 (1.368)	3.17 (1.324)	2.88 (1.845)	3.60 (1.977)
		95% CI	2.41, 3.27	2.76, 3.58	1.96, 3.80	2.61, 4.59
		Median	2.90	3.00	2.05	4.05
		Q25, Q75	1.70, 3.90	2.20, 4.50	1.40, 4.70	1.70, 5.20
		Min, Max	1.0, 5.7	1.0, 5.6	1.0, 6.5	1.0, 6.9
	T <sub>0</sub> + 45 min	n	43	43	18	18
		Mean (SD)	2.95 (1.306)	3.33 (1.332)	3.02 (1.885)	3.33 (1.999)
		95% CI	2.54, 3.36	2.92, 3.75	2.07, 3.96	2.33, 4.33
		Median	2.90	3.20	2.40	3.40
		Q25, Q75	2.00, 4.10	2.60, 4.40	1.30, 5.00	1.60, 4.70
		Min, Max	1.0, 5.9	1.0, 6.1	1.0, 6.7	1.0, 6.7

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: The QSU-brief values at T<sub>0</sub> + 15/20 min refer to QSU-brief data at time T<sub>0</sub> + 15 min for THS 2.2 Menthol and mCC, and to QSU-brief data at time T<sub>0</sub> + 20 min for NRT gum.

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/000000106326/dev/tables/t\_qsu.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.14 Descriptive Statistics of QSU-brief Questionnaire Factors and Total Score - PK Population**

Variable	Timepoint	Statistic	Group-1 PK		Group-2 PK	
			THS 2.2 Menthol (N=43)	mCC (N=43)	THS 2.2 Menthol (N=18)	NRT gum (N=18)
Total score	T <sub>0</sub> + 60 min	n	43	43	18	18
		Mean (SD)	3.06 (1.423)	3.45 (1.403)	2.90 (1.712)	3.50 (2.136)
		95% CI	2.62, 3.51	3.01, 3.88	2.04, 3.76	2.43, 4.57
		Median	3.20	3.40	2.20	3.40
		Q25, Q75	1.50, 4.20	2.50, 4.60	1.40, 4.00	1.50, 5.20
		Min, Max	1.0, 5.8	1.0, 5.9	1.0, 6.7	1.0, 6.7
	T <sub>0</sub> + 2 h	n	43	43	18	18
		Mean (SD)	3.35 (1.417)	3.57 (1.252)	3.37 (1.951)	3.67 (1.919)
		95% CI	2.91, 3.79	3.18, 3.96	2.39, 4.34	2.71, 4.63
		Median	3.30	3.80	3.25	4.00
		Q25, Q75	2.20, 4.40	2.60, 4.60	1.60, 5.00	2.10, 5.30
		Min, Max	1.0, 5.7	1.0, 5.7	1.0, 6.7	1.0, 6.7

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: The QSU-brief values at T<sub>0</sub> + 15/20 min refer to QSU-brief data at time T<sub>0</sub> + 15 min for THS 2.2 Menthol and mCC, and to QSU-brief data at time T<sub>0</sub> + 20 min for NRT gum.

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/000000106326/dev/tables/t\_qsu.sas

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**Table 15.2.4.14 Descriptive Statistics of QSU-brief Questionnaire Factors and Total Score - PK Population**

Variable	Timepoint	Statistic	Group-1 PK		Group-2 PK	
			THS 2.2 Menthol (N=43)	mCC (N=43)	THS 2.2 Menthol (N=18)	NRT gum (N=18)
Total score	T <sub>0</sub> + 4 h	n	43	43	18	18
		Mean (SD)	3.47 (1.260)	3.70 (1.293)	3.49 (1.832)	3.72 (2.001)
		95% CI	3.08, 3.87	3.30, 4.11	2.58, 4.41	2.72, 4.72
		Median	3.40	3.80	3.25	4.40
		Q25, Q75	2.60, 4.60	2.70, 4.70	2.00, 4.80	1.90, 5.40
		Min, Max	1.0, 6.0	1.0, 5.6	1.0, 6.6	1.0, 6.8
	T <sub>0</sub> + 6 h	n	43	43	18	18
		Mean (SD)	3.80 (1.397)	4.08 (1.382)	3.85 (1.795)	3.82 (1.982)
		95% CI	3.36, 4.23	3.65, 4.51	2.95, 4.75	2.83, 4.81
		Median	4.00	4.30	3.95	4.15
		Q25, Q75	2.80, 4.80	3.10, 5.00	2.40, 5.20	2.10, 5.40
		Min, Max	1.0, 6.4	1.0, 6.7	1.0, 6.7	1.0, 6.7

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: The QSU-brief values at T<sub>0</sub> + 15/20 min refer to QSU-brief data at time T<sub>0</sub> + 15 min for THS 2.2 Menthol and mCC, and to QSU-brief data at time T<sub>0</sub> + 20 min for NRT gum.

Note: QSU-brief scores reported on a 7-point scale. Higher scores indicate greater intensity of urge.

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**Table 15.2.4.14 Descriptive Statistics of QSU-brief Questionnaire Factors and Total Score - PK Population**

Variable	Timepoint	Statistic	Group-1 PK		Group-2 PK	
			THS 2.2 Menthol (N=43)	mCC (N=43)	THS 2.2 Menthol (N=18)	NRT gum (N=18)
Total score	T <sub>0</sub> + 9 h	n	43	43	18	18
		Mean (SD)	3.89 (1.320)	4.04 (1.297)	3.71 (1.864)	3.77 (1.870)
		95% CI	3.48, 4.30	3.64, 4.45	2.77, 4.64	2.84, 4.71
		Median	4.10	4.30	3.70	4.10
		Q25, Q75	3.20, 4.90	3.30, 4.80	2.00, 5.20	2.30, 5.20
		Min, Max	1.0, 6.3	1.0, 6.3	1.0, 7.0	1.0, 6.6
	T <sub>0</sub> + 12 h	n	43	43	18	18
		Mean (SD)	3.93 (1.387)	4.12 (1.319)	3.79 (1.857)	3.86 (1.946)
		95% CI	3.50, 4.36	3.71, 4.53	2.86, 4.72	2.88, 4.83
		Median	4.00	4.30	4.10	4.20
		Q25, Q75	3.20, 4.90	3.30, 5.10	2.40, 5.20	2.00, 5.50
		Min, Max	1.0, 6.7	1.2, 6.7	1.0, 6.7	1.1, 6.7

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: The QSU-brief values at T<sub>0</sub> + 15/20 min refer to QSU-brief data at time T<sub>0</sub> + 15 min for THS 2.2 Menthol and mCC, and to QSU-brief data at time T<sub>0</sub> + 20 min for NRT gum.

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/000000106326/dev/tables/t\_qsu.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.14.1 Descriptive Statistics of QSU-brief Questionnaire Factors and Total Score by Sex - PK Population**

Variable	Timepoint	Statistic	Group-1 PK			
			THS 2.2 Menthol		mCC	
			Males (N=23)	Females (N=20)	Males (N=23)	Females (N=20)
Factor 1 - Reward	15 min < T <sub>0</sub>	n	23	20	23	20
		Mean (SD)	5.20 (1.261)	5.56 (0.896)	5.41 (0.776)	5.38 (1.391)
		95% CI	4.65, 5.75	5.14, 5.98	5.07, 5.75	4.72, 6.04
		Median	5.40	5.80	5.40	5.70
		Q25, Q75	4.60, 5.80	5.10, 6.20	4.80, 6.20	4.60, 6.50
		Min, Max	2.2, 7.0	3.6, 6.8	4.0, 6.6	2.0, 7.0
	T <sub>0</sub> + 15/20 min	n	23	20	23	20
		Mean (SD)	3.20 (1.433)	3.74 (1.954)	3.66 (1.779)	4.16 (1.516)
		95% CI	2.58, 3.82	2.82, 4.66	2.89, 4.43	3.45, 4.87
		Median	3.00	3.40	3.80	4.10
		Q25, Q75	2.20, 4.40	2.10, 5.60	2.20, 5.00	2.90, 5.70
		Min, Max	1.0, 6.4	1.0, 7.0	1.0, 6.6	1.6, 6.6

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: The QSU-brief values at T<sub>0</sub> + 15/20 min refer to QSU-brief data at time T<sub>0</sub> + 15 min for THS 2.2 Menthol and mCC, and to QSU-brief data at time T<sub>0</sub> + 20 min for NRT gum.

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/000000106326/dev/tables/t\_qsu2.sas

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**Table 15.2.4.14.1 Descriptive Statistics of QSU-brief Questionnaire Factors and Total Score by Sex - PK Population**

Variable	Timepoint	Statistic	Group-1 PK			
			THS 2.2 Menthol		mCC	
			Males (N=23)	Females (N=20)	Males (N=23)	Females (N=20)
Factor 1 - Reward	T <sub>0</sub> + 30 min	n	23	20	23	20
		Mean (SD)	3.19 (1.727)	3.81 (1.901)	3.68 (1.668)	4.08 (1.668)
		95% CI	2.44, 3.94	2.92, 4.70	2.95, 4.40	3.29, 4.87
		Median	2.80	3.30	3.80	3.80
		Q25, Q75	1.80, 4.80	2.20, 5.70	2.20, 4.80	2.70, 5.70
		Min, Max	1.0, 6.4	1.0, 7.0	1.0, 6.4	1.6, 6.6
	T <sub>0</sub> + 45 min	n	23	20	23	20
		Mean (SD)	3.48 (1.669)	3.89 (1.832)	3.92 (1.608)	4.22 (1.616)
		95% CI	2.75, 4.20	3.03, 4.75	3.22, 4.62	3.46, 4.98
		Median	3.40	3.80	4.00	4.10
		Q25, Q75	2.20, 5.00	2.40, 5.60	2.60, 5.20	3.20, 5.70
		Min, Max	1.0, 6.6	1.0, 6.6	1.0, 6.2	1.6, 6.6

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: The QSU-brief values at T<sub>0</sub> + 15/20 min refer to QSU-brief data at time T<sub>0</sub> + 15 min for THS 2.2 Menthol and mCC, and to QSU-brief data at time T<sub>0</sub> + 20 min for NRT gum.

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/000000106326/dev/tables/t\_qsu2.sas

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**Table 15.2.4.14.1 Descriptive Statistics of QSU-brief Questionnaire Factors and Total Score by Sex - PK Population**

Variable	Timepoint	Statistic	Group-1 PK			
			THS 2.2 Menthol		mCC	
			Males (N=23)	Females (N=20)	Males (N=23)	Females (N=20)
Factor 1 - Reward	T <sub>0</sub> + 60 min	n	23	20	23	20
		Mean (SD)	3.68 (1.753)	3.87 (1.936)	4.16 (1.706)	4.25 (1.691)
		95% CI	2.92, 4.44	2.96, 4.78	3.41, 4.90	3.45, 5.05
		Median	4.00	3.70	4.40	3.80
		Q25, Q75	2.00, 5.40	1.90, 5.60	3.00, 5.40	3.10, 5.80
		Min, Max	1.0, 6.4	1.0, 6.8	1.0, 6.4	1.2, 7.0
	T <sub>0</sub> + 2 h	n	23	20	23	20
		Mean (SD)	3.97 (1.675)	4.35 (1.945)	4.37 (1.520)	4.52 (1.573)
		95% CI	3.24, 4.70	3.44, 5.26	3.70, 5.03	3.78, 5.26
		Median	4.00	5.10	4.40	4.70
		Q25, Q75	2.60, 5.60	2.90, 5.80	3.80, 5.40	3.20, 5.70
		Min, Max	1.0, 6.6	1.0, 7.0	1.0, 6.6	1.6, 6.8

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: The QSU-brief values at T<sub>0</sub> + 15/20 min refer to QSU-brief data at time T<sub>0</sub> + 15 min for THS 2.2 Menthol and mCC, and to QSU-brief data at time T<sub>0</sub> + 20 min for NRT gum.

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/000000106326/dev/tables/t\_qsu2.sas

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**Table 15.2.4.14.1 Descriptive Statistics of QSU-brief Questionnaire Factors and Total Score by Sex - PK Population**

Variable	Timepoint	Statistic	Group-1 PK			
			THS 2.2 Menthol		mCC	
			Males (N=23)	Females (N=20)	Males (N=23)	Females (N=20)
Factor 1 - Reward	T <sub>0</sub> + 4 h	n	23	20	23	20
		Mean (SD)	4.25 (1.406)	4.35 (1.751)	4.60 (1.466)	4.51 (1.623)
		95% CI	3.64, 4.86	3.53, 5.17	3.96, 5.24	3.75, 5.27
		Median	4.00	4.30	4.40	4.60
		Q25, Q75	3.60, 5.20	3.60, 5.80	4.40, 5.60	3.30, 5.90
		Min, Max	1.0, 6.6	1.0, 7.0	1.0, 6.6	1.4, 6.6
	T <sub>0</sub> + 6 h	n	23	20	23	20
		Mean (SD)	4.61 (1.533)	4.76 (1.828)	4.97 (1.538)	5.17 (1.546)
		95% CI	3.94, 5.28	3.90, 5.62	4.30, 5.63	4.44, 5.90
		Median	4.60	5.10	5.00	5.10
		Q25, Q75	3.40, 5.80	3.90, 6.30	4.20, 6.00	4.50, 6.60
		Min, Max	1.0, 6.8	1.0, 7.0	1.0, 7.0	1.6, 7.0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: The QSU-brief values at T<sub>0</sub> + 15/20 min refer to QSU-brief data at time T<sub>0</sub> + 15 min for THS 2.2 Menthol and mCC, and to QSU-brief data at time T<sub>0</sub> + 20 min for NRT gum.

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/000000106326/dev/tables/t\_qsu2.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.14.1 Descriptive Statistics of QSU-brief Questionnaire Factors and Total Score by Sex - PK Population**

Variable	Timepoint	Statistic	Group-1 PK			
			THS 2.2 Menthol		mCC	
			Males (N=23)	Females (N=20)	Males (N=23)	Females (N=20)
Factor 1 - Reward	T <sub>0</sub> + 9 h	n	23	20	23	20
		Mean (SD)	4.88 (1.245)	4.75 (1.851)	4.91 (1.400)	5.04 (1.512)
		95% CI	4.34, 5.42	3.88, 5.62	4.30, 5.52	4.33, 5.75
		Median	4.60	5.50	5.20	5.10
		Q25, Q75	4.20, 6.00	3.90, 6.20	4.40, 5.60	4.40, 6.10
		Min, Max	2.6, 6.8	1.0, 6.8	1.0, 6.6	1.4, 7.0
	T <sub>0</sub> + 12 h	n	23	20	23	20
		Mean (SD)	4.83 (1.243)	4.96 (1.956)	5.03 (1.334)	5.14 (1.599)
		95% CI	4.29, 5.38	4.04, 5.88	4.45, 5.62	4.39, 5.89
		Median	5.20	5.50	5.20	5.20
		Q25, Q75	4.00, 5.80	4.30, 6.20	4.40, 5.80	4.60, 6.50
		Min, Max	2.0, 6.8	1.0, 7.0	1.6, 7.0	1.4, 7.0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: The QSU-brief values at T<sub>0</sub> + 15/20 min refer to QSU-brief data at time T<sub>0</sub> + 15 min for THS 2.2 Menthol and mCC, and to QSU-brief data at time T<sub>0</sub> + 20 min for NRT gum.

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/000000106326/dev/tables/t\_qsu2.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.14.1 Descriptive Statistics of QSU-brief Questionnaire Factors and Total Score by Sex - PK Population**

Variable	Timepoint	Statistic	Group-1 PK			
			THS 2.2 Menthol		mCC	
			Males (N=23)	Females (N=20)	Males (N=23)	Females (N=20)
Factor 2 - Relief	15 min < T <sub>0</sub>	n	23	20	23	20
		Mean (SD)	3.30 (1.361)	3.41 (1.060)	3.40 (1.208)	3.63 (1.476)
		95% CI	2.71, 3.90	2.91, 3.91	2.87, 3.93	2.93, 4.33
		Median	3.20	3.50	3.40	4.00
		Q25, Q75	2.60, 4.60	2.80, 4.30	2.60, 4.20	2.30, 5.00
		Min, Max	1.0, 5.6	1.2, 4.8	1.2, 6.0	1.0, 5.6
	T <sub>0</sub> + 15/20 min	n	23	20	23	20
		Mean (SD)	2.02 (0.963)	2.48 (1.192)	2.55 (1.222)	2.71 (1.171)
		95% CI	1.60, 2.44	1.92, 3.04	2.01, 3.08	2.16, 3.26
		Median	2.00	2.30	2.40	2.40
		Q25, Q75	1.20, 2.60	1.50, 3.80	1.40, 3.60	2.00, 3.60
		Min, Max	1.0, 4.8	1.0, 4.6	1.0, 4.8	1.0, 4.8

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: The QSU-brief values at T<sub>0</sub> + 15/20 min refer to QSU-brief data at time T<sub>0</sub> + 15 min for THS 2.2 Menthol and mCC, and to QSU-brief data at time T<sub>0</sub> + 20 min for NRT gum.

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/000000106326/dev/tables/t\_qsu2.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.14.1 Descriptive Statistics of QSU-brief Questionnaire Factors and Total Score by Sex - PK Population**

Variable	Timepoint	Statistic	Group-1 PK			
			THS 2.2 Menthol		mCC	
			Males (N=23)	Females (N=20)	Males (N=23)	Females (N=20)
Factor 2 - Relief	T <sub>0</sub> + 30 min	n	23	20	23	20
		Mean (SD)	2.05 (1.116)	2.37 (1.125)	2.39 (1.155)	2.57 (1.153)
		95% CI	1.56, 2.54	1.84, 2.90	1.89, 2.90	2.03, 3.11
		Median	1.80	2.40	2.20	2.40
		Q25, Q75	1.00, 2.40	1.20, 3.10	1.60, 3.00	1.70, 3.40
		Min, Max	1.0, 5.0	1.0, 4.4	1.0, 5.2	1.0, 4.6
	T <sub>0</sub> + 45 min	n	23	20	23	20
		Mean (SD)	2.14 (1.108)	2.34 (1.104)	2.55 (1.223)	2.66 (1.257)
		95% CI	1.66, 2.62	1.82, 2.86	2.01, 3.08	2.07, 3.25
		Median	2.00	2.20	2.60	2.30
		Q25, Q75	1.00, 2.60	1.30, 3.00	1.60, 3.00	1.90, 3.60
		Min, Max	1.0, 5.2	1.0, 4.4	1.0, 5.8	1.0, 5.6

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: The QSU-brief values at T<sub>0</sub> + 15/20 min refer to QSU-brief data at time T<sub>0</sub> + 15 min for THS 2.2 Menthol and mCC, and to QSU-brief data at time T<sub>0</sub> + 20 min for NRT gum.

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/000000106326/dev/tables/t\_qsu2.sas

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**Table 15.2.4.14.1 Descriptive Statistics of QSU-brief Questionnaire Factors and Total Score by Sex - PK Population**

Variable	Timepoint	Statistic	Group-1 PK			
			THS 2.2 Menthol		mCC	
			Males (N=23)	Females (N=20)	Males (N=23)	Females (N=20)
Factor 2 - Relief	T <sub>0</sub> + 60 min	n	23	20	23	20
		Mean (SD)	2.25 (1.178)	2.48 (1.384)	2.80 (1.339)	2.57 (1.206)
		95% CI	1.74, 2.77	1.83, 3.13	2.22, 3.38	2.00, 3.14
		Median	2.00	2.20	2.80	2.20
		Q25, Q75	1.00, 3.00	1.10, 3.40	1.80, 3.80	1.70, 3.70
		Min, Max	1.0, 5.0	1.0, 5.8	1.0, 5.4	1.0, 4.6
	T <sub>0</sub> + 2 h	n	23	20	23	20
		Mean (SD)	2.43 (1.240)	2.68 (1.333)	2.73 (1.240)	2.65 (1.195)
		95% CI	1.89, 2.98	2.05, 3.31	2.19, 3.27	2.09, 3.21
		Median	2.00	2.80	3.00	2.60
		Q25, Q75	1.20, 3.20	1.50, 3.70	1.80, 3.40	1.70, 3.80
		Min, Max	1.0, 4.8	1.0, 5.2	1.0, 4.8	1.0, 4.4

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: The QSU-brief values at T<sub>0</sub> + 15/20 min refer to QSU-brief data at time T<sub>0</sub> + 15 min for THS 2.2 Menthol and mCC, and to QSU-brief data at time T<sub>0</sub> + 20 min for NRT gum.

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/000000106326/dev/tables/t\_qsu2.sas

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**Table 15.2.4.14.1 Descriptive Statistics of QSU-brief Questionnaire Factors and Total Score by Sex - PK Population**

Variable	Timepoint	Statistic	Group-1 PK			
			THS 2.2 Menthol		mCC	
			Males (N=23)	Females (N=20)	Males (N=23)	Females (N=20)
Factor 2 - Relief	T <sub>0</sub> + 4 h	n	23	20	23	20
		Mean (SD)	2.61 (1.207)	2.70 (1.108)	2.82 (1.260)	2.89 (1.260)
		95% CI	2.08, 3.14	2.18, 3.22	2.27, 3.37	2.30, 3.48
		Median	2.60	2.90	2.60	3.10
		Q25, Q75	1.60, 3.40	1.80, 3.10	1.60, 4.00	2.00, 4.00
		Min, Max	1.0, 5.4	1.0, 4.6	1.0, 5.0	1.0, 4.8
	T <sub>0</sub> + 6 h	n	23	20	23	20
		Mean (SD)	2.88 (1.373)	2.96 (1.433)	3.03 (1.361)	3.19 (1.524)
		95% CI	2.28, 3.48	2.28, 3.64	2.43, 3.62	2.47, 3.91
		Median	2.80	3.00	2.80	3.40
		Q25, Q75	1.60, 4.00	2.00, 4.00	2.20, 4.20	1.80, 4.10
		Min, Max	1.0, 5.6	1.0, 6.2	1.0, 5.6	1.0, 6.8

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: The QSU-brief values at T<sub>0</sub> + 15/20 min refer to QSU-brief data at time T<sub>0</sub> + 15 min for THS 2.2 Menthol and mCC, and to QSU-brief data at time T<sub>0</sub> + 20 min for NRT gum.

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/000000106326/dev/tables/t\_qsu2.sas

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**Table 15.2.4.14.1 Descriptive Statistics of QSU-brief Questionnaire Factors and Total Score by Sex - PK Population**

Variable	Timepoint	Statistic	Group-1 PK			
			THS 2.2 Menthol		mCC	
			Males (N=23)	Females (N=20)	Males (N=23)	Females (N=20)
Factor 2 - Relief	T <sub>0</sub> + 9 h	n	23	20	23	20
		Mean (SD)	2.93 (1.404)	3.01 (1.347)	3.16 (1.433)	3.06 (1.315)
		95% CI	2.32, 3.54	2.37, 3.65	2.53, 3.78	2.44, 3.68
		Median	2.80	2.90	3.40	3.20
		Q25, Q75	1.60, 4.20	2.10, 3.80	2.20, 4.20	2.30, 4.20
		Min, Max	1.0, 5.6	1.0, 6.0	1.0, 6.0	1.0, 5.4
	T <sub>0</sub> + 12 h	n	23	20	23	20
		Mean (SD)	2.88 (1.404)	3.07 (1.449)	3.18 (1.455)	3.12 (1.362)
		95% CI	2.27, 3.49	2.39, 3.75	2.55, 3.82	2.48, 3.76
		Median	2.80	2.90	3.20	3.30
		Q25, Q75	1.80, 3.80	2.00, 4.00	2.20, 4.20	2.20, 4.10
		Min, Max	1.0, 5.8	1.0, 6.4	1.0, 6.4	1.0, 5.2

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: The QSU-brief values at T<sub>0</sub> + 15/20 min refer to QSU-brief data at time T<sub>0</sub> + 15 min for THS 2.2 Menthol and mCC, and to QSU-brief data at time T<sub>0</sub> + 20 min for NRT gum.

Note: QSU-brief scores reported on a 7-point scale. Higher scores indicate greater intensity of urge.

Appendix 15.3.6.11

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**Table 15.2.4.14.1 Descriptive Statistics of QSU-brief Questionnaire Factors and Total Score by Sex - PK Population**

Variable	Timepoint	Statistic	Group-1 PK			
			THS 2.2 Menthol		mCC	
			Males (N=23)	Females (N=20)	Males (N=23)	Females (N=20)
Total score	15 min < T <sub>0</sub>	n	23	20	23	20
		Mean (SD)	4.25 (1.209)	4.49 (0.873)	4.40 (0.906)	4.51 (1.368)
		95% CI	3.72, 4.78	4.07, 4.90	4.01, 4.80	3.86, 5.15
		Median	4.20	4.75	4.30	4.55
		Q25, Q75	3.40, 5.20	3.95, 5.15	3.70, 5.20	3.60, 5.50
		Min, Max	1.6, 6.3	2.9, 5.6	2.7, 6.1	1.5, 6.3
	T <sub>0</sub> + 15/20 min	n	23	20	23	20
		Mean (SD)	2.61 (1.124)	3.11 (1.453)	3.10 (1.471)	3.44 (1.247)
		95% CI	2.12, 3.10	2.43, 3.79	2.46, 3.75	2.85, 4.02
		Median	2.50	3.05	3.30	3.35
		Q25, Q75	1.70, 3.30	2.00, 4.60	2.00, 4.40	2.55, 4.35
		Min, Max	1.0, 5.6	1.0, 5.1	1.0, 5.5	1.3, 5.5

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: The QSU-brief values at T<sub>0</sub> + 15/20 min refer to QSU-brief data at time T<sub>0</sub> + 15 min for THS 2.2 Menthol and mCC, and to QSU-brief data at time T<sub>0</sub> + 20 min for NRT gum.

Note: QSU-brief scores reported on a 7-point scale. Higher scores indicate greater intensity of urge.

Appendix 15.3.6.11

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**Table 15.2.4.14.1 Descriptive Statistics of QSU-brief Questionnaire Factors and Total Score by Sex - PK Population**

Variable	Timepoint	Statistic	Group-1 PK			
			THS 2.2 Menthol		mCC	
			Males (N=23)	Females (N=20)	Males (N=23)	Females (N=20)
Total score	T <sub>0</sub> + 30 min	n	23	20	23	20
		Mean (SD)	2.62 (1.338)	3.09 (1.394)	3.03 (1.350)	3.33 (1.311)
		95% CI	2.04, 3.20	2.43, 3.75	2.45, 3.62	2.71, 3.94
		Median	2.40	3.25	3.10	3.00
		Q25, Q75	1.40, 3.40	1.85, 4.50	1.90, 4.00	2.40, 4.60
		Min, Max	1.0, 5.7	1.0, 5.2	1.0, 5.6	1.3, 5.5
	T <sub>0</sub> + 45 min	n	23	20	23	20
		Mean (SD)	2.81 (1.280)	3.12 (1.349)	3.23 (1.352)	3.44 (1.336)
		95% CI	2.25, 3.37	2.48, 3.75	2.65, 3.82	2.81, 4.07
		Median	2.90	2.95	3.30	3.15
		Q25, Q75	2.00, 3.40	2.05, 4.35	2.30, 4.20	2.60, 4.55
		Min, Max	1.0, 5.9	1.0, 5.2	1.0, 5.9	1.3, 6.1

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: The QSU-brief values at T<sub>0</sub> + 15/20 min refer to QSU-brief data at time T<sub>0</sub> + 15 min for THS 2.2 Menthol and mCC, and to QSU-brief data at time T<sub>0</sub> + 20 min for NRT gum.

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/000000106326/dev/tables/t\_qsu2.sas

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**Table 15.2.4.14.1 Descriptive Statistics of QSU-brief Questionnaire Factors and Total Score by Sex - PK Population**

Variable	Timepoint	Statistic	Group-1 PK			
			THS 2.2 Menthol		mCC	
			Males (N=23)	Females (N=20)	Males (N=23)	Females (N=20)
Total score	T <sub>0</sub> + 60 min	n	23	20	23	20
		Mean (SD)	2.97 (1.353)	3.18 (1.527)	3.48 (1.475)	3.41 (1.352)
		95% CI	2.38, 3.56	2.46, 3.89	2.84, 4.12	2.77, 4.05
		Median	3.20	3.30	3.40	3.10
		Q25, Q75	1.70, 3.80	1.50, 4.50	2.50, 4.60	2.45, 4.60
		Min, Max	1.0, 5.7	1.0, 5.8	1.0, 5.9	1.1, 5.6
	T <sub>0</sub> + 2 h	n	23	20	23	20
		Mean (SD)	3.20 (1.351)	3.52 (1.506)	3.55 (1.303)	3.59 (1.223)
		95% CI	2.62, 3.79	2.81, 4.22	2.98, 4.12	3.01, 4.16
		Median	3.30	3.70	3.50	4.20
		Q25, Q75	2.20, 4.20	2.25, 4.70	2.60, 4.60	2.60, 4.50
		Min, Max	1.0, 5.7	1.0, 5.6	1.0, 5.7	1.3, 5.2

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: The QSU-brief values at T<sub>0</sub> + 15/20 min refer to QSU-brief data at time T<sub>0</sub> + 15 min for THS 2.2 Menthol and mCC, and to QSU-brief data at time T<sub>0</sub> + 20 min for NRT gum.

Note: QSU-brief scores reported on a 7-point scale. Higher scores indicate greater intensity of urge.

Appendix 15.3.6.11

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**Table 15.2.4.14.1 Descriptive Statistics of QSU-brief Questionnaire Factors and Total Score by Sex - PK Population**

Variable	Timepoint	Statistic	Group-1 PK			
			THS 2.2 Menthol		mCC	
			Males (N=23)	Females (N=20)	Males (N=23)	Females (N=20)
Total score	T <sub>0</sub> + 4 h	n	23	20	23	20
		Mean (SD)	3.43 (1.209)	3.53 (1.345)	3.71 (1.249)	3.70 (1.374)
		95% CI	2.90, 3.96	2.89, 4.16	3.16, 4.25	3.05, 4.35
		Median	3.30	3.65	3.60	4.10
		Q25, Q75	2.60, 4.10	2.65, 4.85	2.80, 4.60	2.50, 4.90
		Min, Max	1.0, 6.0	1.0, 5.2	1.0, 5.6	1.2, 5.6
	T <sub>0</sub> + 6 h	n	23	20	23	20
		Mean (SD)	3.74 (1.328)	3.86 (1.505)	4.00 (1.368)	4.18 (1.425)
		95% CI	3.16, 4.32	3.15, 4.57	3.40, 4.59	3.51, 4.85
		Median	3.60	4.05	4.00	4.55
		Q25, Q75	2.70, 4.80	3.00, 4.85	3.20, 5.00	3.05, 5.10
		Min, Max	1.0, 6.2	1.0, 6.4	1.0, 6.2	1.3, 6.7

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: The QSU-brief values at T<sub>0</sub> + 15/20 min refer to QSU-brief data at time T<sub>0</sub> + 15 min for THS 2.2 Menthol and mCC, and to QSU-brief data at time T<sub>0</sub> + 20 min for NRT gum.

Note: QSU-brief scores reported on a 7-point scale. Higher scores indicate greater intensity of urge.

Appendix 15.3.6.11

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**Table 15.2.4.14.1 Descriptive Statistics of QSU-brief Questionnaire Factors and Total Score by Sex - PK Population**

Variable	Timepoint	Statistic	Group-1 PK			
			THS 2.2 Menthol		mCC	
			Males (N=23)	Females (N=20)	Males (N=23)	Females (N=20)
Total score	T <sub>0</sub> + 9 h	n	23	20	23	20
		Mean (SD)	3.90 (1.187)	3.88 (1.490)	4.03 (1.321)	4.05 (1.303)
		95% CI	3.39, 4.42	3.18, 4.58	3.46, 4.61	3.44, 4.66
		Median	3.70	4.10	4.20	4.50
		Q25, Q75	3.10, 4.90	3.35, 4.90	3.20, 4.90	3.35, 4.75
		Min, Max	2.0, 5.8	1.0, 6.3	1.0, 6.3	1.2, 6.2
	T <sub>0</sub> + 12 h	n	23	20	23	20
		Mean (SD)	3.86 (1.214)	4.02 (1.591)	4.11 (1.302)	4.13 (1.373)
		95% CI	3.33, 4.39	3.27, 4.76	3.54, 4.68	3.48, 4.78
		Median	4.00	4.30	4.30	4.35
		Q25, Q75	3.10, 4.60	3.55, 5.05	3.30, 4.90	3.40, 5.15
		Min, Max	1.5, 6.3	1.0, 6.7	1.3, 6.7	1.2, 5.9

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: The QSU-brief values at T<sub>0</sub> + 15/20 min refer to QSU-brief data at time T<sub>0</sub> + 15 min for THS 2.2 Menthol and mCC, and to QSU-brief data at time T<sub>0</sub> + 20 min for NRT gum.

Note: QSU-brief scores reported on a 7-point scale. Higher scores indicate greater intensity of urge.

Appendix 15.3.6.11

Path: /cvm/projects/prj/development/000000106326/dev/tables/t\_qsu2.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.14.1 Descriptive Statistics of QSU-brief Questionnaire Factors and Total Score by Sex - PK Population**

Variable	Timepoint	Statistic	Group-2 PK			
			THS 2.2 Menthol		NRT gum	
			Males (N=9)	Females (N=9)	Males (N=9)	Females (N=9)
Factor 1 - Reward	15 min < T <sub>0</sub>	n	9	9	9	9
		Mean (SD)	4.44 (2.280)	5.18 (2.041)	4.96 (1.682)	5.40 (1.467)
		95% CI	2.69, 6.20	3.60, 6.75	3.66, 6.25	4.27, 6.53
		Median	4.80	5.40	5.00	5.40
		Q25, Q75	2.40, 6.80	4.20, 7.00	4.00, 6.40	5.00, 6.60
		Min, Max	1.4, 7.0	1.8, 7.0	2.0, 7.0	2.4, 7.0
	T <sub>0</sub> + 15/20 min	n	9	9	9	9
		Mean (SD)	3.07 (1.963)	4.16 (2.377)	3.78 (2.215)	5.09 (2.057)
		95% CI	1.55, 4.58	2.32, 5.99	2.07, 5.49	3.50, 6.68
		Median	2.00	4.20	2.80	6.00
		Q25, Q75	1.80, 4.60	1.40, 6.00	2.00, 5.40	3.40, 7.00
		Min, Max	1.0, 6.6	1.2, 7.0	1.0, 6.8	1.6, 7.0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: The QSU-brief values at T<sub>0</sub> + 15/20 min refer to QSU-brief data at time T<sub>0</sub> + 15 min for THS 2.2 Menthol and mCC, and to QSU-brief data at time T<sub>0</sub> + 20 min for NRT gum.

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/000000106326/dev/tables/t\_qsu2.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.14.1 Descriptive Statistics of QSU-brief Questionnaire Factors and Total Score by Sex - PK Population**

Variable	Timepoint	Statistic	Group-2 PK			
			THS 2.2 Menthol		NRT gum	
			Males (N=9)	Females (N=9)	Males (N=9)	Females (N=9)
Factor 1 - Reward	T <sub>0</sub> + 30 min	n	9	9	9	9
		Mean (SD)	3.16 (1.952)	3.96 (2.567)	3.58 (2.310)	4.98 (2.271)
		95% CI	1.65, 4.66	1.98, 5.93	1.80, 5.36	3.23, 6.73
		Median	2.80	2.80	2.60	5.80
		Q25, Q75	1.40, 5.20	1.80, 6.40	1.80, 5.00	3.60, 7.00
		Min, Max	1.0, 6.2	1.0, 7.0	1.0, 7.0	1.2, 7.0
	T <sub>0</sub> + 45 min	n	9	9	9	9
		Mean (SD)	3.16 (2.071)	4.07 (2.318)	3.38 (2.208)	4.49 (2.491)
		95% CI	1.56, 4.75	2.28, 5.85	1.68, 5.08	2.57, 6.41
		Median	2.20	4.00	2.40	5.60
		Q25, Q75	1.40, 5.20	2.00, 6.00	2.00, 4.40	2.40, 7.00
		Min, Max	1.0, 6.4	1.2, 7.0	1.0, 6.8	1.0, 7.0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: The QSU-brief values at T<sub>0</sub> + 15/20 min refer to QSU-brief data at time T<sub>0</sub> + 15 min for THS 2.2 Menthol and mCC, and to QSU-brief data at time T<sub>0</sub> + 20 min for NRT gum.

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/000000106326/dev/tables/t\_qsu2.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.14.1 Descriptive Statistics of QSU-brief Questionnaire Factors and Total Score by Sex - PK Population**

Variable	Timepoint	Statistic	Group-2 PK			
			THS 2.2 Menthol		NRT gum	
			Males (N=9)	Females (N=9)	Males (N=9)	Females (N=9)
Factor 1 - Reward	T <sub>0</sub> + 60 min	n	9	9	9	9
		Mean (SD)	3.02 (1.791)	4.29 (2.344)	3.69 (2.403)	4.42 (2.638)
		95% CI	1.64, 4.40	2.48, 6.10	1.84, 5.54	2.39, 6.45
		Median	2.40	5.40	2.60	5.80
		Q25, Q75	1.40, 5.00	2.00, 6.00	2.00, 6.40	2.00, 7.00
		Min, Max	1.0, 5.4	1.4, 7.0	1.0, 6.8	1.0, 7.0
	T <sub>0</sub> + 2 h	n	9	9	9	9
		Mean (SD)	3.56 (2.209)	4.53 (2.413)	3.93 (2.210)	4.82 (2.188)
		95% CI	1.85, 5.26	2.67, 6.39	2.23, 5.64	3.14, 6.51
		Median	3.20	5.40	3.40	5.40
		Q25, Q75	1.60, 4.60	2.20, 7.00	2.80, 5.80	3.20, 7.00
		Min, Max	1.0, 6.8	1.8, 7.0	1.0, 6.8	1.0, 7.0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: The QSU-brief values at T<sub>0</sub> + 15/20 min refer to QSU-brief data at time T<sub>0</sub> + 15 min for THS 2.2 Menthol and mCC, and to QSU-brief data at time T<sub>0</sub> + 20 min for NRT gum.

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/000000106326/dev/tables/t\_qsu2.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.14.1 Descriptive Statistics of QSU-brief Questionnaire Factors and Total Score by Sex - PK Population**

Variable	Timepoint	Statistic	Group-2 PK			
			THS 2.2 Menthol		NRT gum	
			Males (N=9)	Females (N=9)	Males (N=9)	Females (N=9)
Factor 1 - Reward	T <sub>0</sub> + 4 h	n	9	9	9	9
		Mean (SD)	3.89 (2.038)	4.84 (2.274)	3.93 (2.152)	4.98 (2.499)
		95% CI	2.32, 5.46	3.09, 6.60	2.27, 5.59	3.05, 6.90
		Median	3.20	5.00	2.60	5.80
		Q25, Q75	2.60, 4.80	2.80, 7.00	2.00, 5.80	3.00, 7.00
		Min, Max	1.2, 7.0	1.0, 7.0	1.6, 6.8	1.0, 7.0
	T <sub>0</sub> + 6 h	n	9	9	9	9
		Mean (SD)	4.09 (2.129)	5.51 (1.861)	4.33 (2.119)	4.89 (2.446)
		95% CI	2.45, 5.73	4.08, 6.95	2.70, 5.97	3.00, 6.77
		Median	3.20	5.80	3.60	5.60
		Q25, Q75	3.00, 5.60	4.80, 7.00	2.80, 6.60	3.00, 7.00
		Min, Max	1.0, 6.8	1.6, 7.0	1.8, 6.8	1.0, 7.0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: The QSU-brief values at T<sub>0</sub> + 15/20 min refer to QSU-brief data at time T<sub>0</sub> + 15 min for THS 2.2 Menthol and mCC, and to QSU-brief data at time T<sub>0</sub> + 20 min for NRT gum.

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/000000106326/dev/tables/t\_qsu2.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.14.1 Descriptive Statistics of QSU-brief Questionnaire Factors and Total Score by Sex - PK Population**

Variable	Timepoint	Statistic	Group-2 PK			
			THS 2.2 Menthol		NRT gum	
			Males (N=9)	Females (N=9)	Males (N=9)	Females (N=9)
Factor 1 - Reward	T <sub>0</sub> + 9 h	n	9	9	9	9
		Mean (SD)	4.09 (2.026)	4.96 (2.247)	4.29 (1.890)	4.91 (2.454)
		95% CI	2.53, 5.65	3.22, 6.69	2.83, 5.75	3.02, 6.80
		Median	3.80	5.80	3.60	6.00
		Q25, Q75	3.00, 5.00	3.00, 7.00	3.00, 6.40	2.80, 7.00
		Min, Max	1.0, 7.0	1.4, 7.0	1.8, 6.8	1.0, 7.0
	T <sub>0</sub> + 12 h	n	9	9	9	9
		Mean (SD)	4.38 (1.984)	4.93 (2.230)	4.40 (2.038)	4.96 (2.437)
		95% CI	2.85, 5.91	3.21, 6.65	2.83, 5.97	3.08, 6.83
		Median	4.80	5.60	4.40	5.80
		Q25, Q75	3.20, 5.60	3.40, 7.00	2.60, 6.40	2.80, 7.00
		Min, Max	1.0, 6.8	1.2, 7.0	1.6, 6.8	1.2, 7.0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: The QSU-brief values at T<sub>0</sub> + 15/20 min refer to QSU-brief data at time T<sub>0</sub> + 15 min for THS 2.2 Menthol and mCC, and to QSU-brief data at time T<sub>0</sub> + 20 min for NRT gum.

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/000000106326/dev/tables/t\_qsu2.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.14.1 Descriptive Statistics of QSU-brief Questionnaire Factors and Total Score by Sex - PK Population**

Variable	Timepoint	Statistic	Group-2 PK			
			THS 2.2 Menthol		NRT gum	
			Males (N=9)	Females (N=9)	Males (N=9)	Females (N=9)
Factor 2 - Relief	15 min < T <sub>0</sub>	n	9	9	9	9
		Mean (SD)	3.40 (2.067)	2.91 (1.707)	3.80 (2.110)	2.91 (1.188)
		95% CI	1.81, 4.99	1.59, 4.23	2.17, 5.43	1.99, 3.83
		Median	3.60	3.00	4.00	2.80
		Q25, Q75	1.60, 5.00	1.20, 3.60	2.00, 4.80	2.40, 3.40
		Min, Max	1.0, 6.4	1.0, 6.0	1.0, 7.0	1.4, 5.2
	T <sub>0</sub> + 15/20 min	n	9	9	9	9
		Mean (SD)	2.29 (1.759)	2.27 (1.850)	3.02 (1.930)	2.93 (1.638)
		95% CI	0.93, 3.65	0.84, 3.69	1.53, 4.51	1.67, 4.20
		Median	1.20	1.40	2.80	2.40
		Q25, Q75	1.00, 4.00	1.00, 3.00	1.40, 4.20	2.00, 3.40
		Min, Max	1.0, 5.0	1.0, 6.2	1.0, 6.4	1.2, 6.2

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: The QSU-brief values at T<sub>0</sub> + 15/20 min refer to QSU-brief data at time T<sub>0</sub> + 15 min for THS 2.2 Menthol and mCC, and to QSU-brief data at time T<sub>0</sub> + 20 min for NRT gum.

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/000000106326/dev/tables/t\_qsu2.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.14.1 Descriptive Statistics of QSU-brief Questionnaire Factors and Total Score by Sex - PK Population**

Variable	Timepoint	Statistic	Group-2 PK			
			THS 2.2 Menthol		NRT gum	
			Males (N=9)	Females (N=9)	Males (N=9)	Females (N=9)
Factor 2 - Relief	T <sub>0</sub> + 30 min	n	9	9	9	9
		Mean (SD)	2.31 (1.747)	2.09 (1.626)	2.93 (2.069)	2.91 (1.718)
		95% CI	0.96, 3.66	0.83, 3.34	1.34, 4.53	1.59, 4.24
		Median	1.20	1.40	2.20	3.20
		Q25, Q75	1.00, 4.20	1.00, 2.40	1.00, 4.00	2.00, 3.40
		Min, Max	1.0, 5.2	1.0, 6.0	1.0, 6.8	1.0, 6.6
	T <sub>0</sub> + 45 min	n	9	9	9	9
		Mean (SD)	2.53 (1.844)	2.31 (1.724)	2.84 (2.005)	2.62 (1.731)
		95% CI	1.11, 3.96	0.98, 3.64	1.30, 4.39	1.29, 3.96
		Median	1.20	2.00	1.80	2.40
		Q25, Q75	1.00, 4.60	1.00, 2.40	1.20, 4.40	1.00, 3.40
		Min, Max	1.0, 5.2	1.0, 6.4	1.0, 6.2	1.0, 6.4

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: The QSU-brief values at T<sub>0</sub> + 15/20 min refer to QSU-brief data at time T<sub>0</sub> + 15 min for THS 2.2 Menthol and mCC, and to QSU-brief data at time T<sub>0</sub> + 20 min for NRT gum.

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/000000106326/dev/tables/t\_qsu2.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.14.1 Descriptive Statistics of QSU-brief Questionnaire Factors and Total Score by Sex - PK Population**

Variable	Timepoint	Statistic	Group-2 PK			
			THS 2.2 Menthol		NRT gum	
			Males (N=9)	Females (N=9)	Males (N=9)	Females (N=9)
Factor 2 - Relief	T <sub>0</sub> + 60 min	n	9	9	9	9
		Mean (SD)	2.16 (1.407)	2.13 (1.738)	3.16 (2.223)	2.73 (1.730)
		95% CI	1.07, 3.24	0.79, 3.47	1.44, 4.87	1.40, 4.07
		Median	1.60	1.60	2.00	3.00
		Q25, Q75	1.00, 2.40	1.00, 2.40	1.20, 4.80	1.00, 3.40
		Min, Max	1.0, 4.6	1.0, 6.4	1.0, 6.4	1.0, 6.4
	T <sub>0</sub> + 2 h	n	9	9	9	9
		Mean (SD)	2.87 (1.945)	2.51 (1.701)	3.11 (2.113)	2.80 (1.631)
		95% CI	1.37, 4.37	1.20, 3.82	1.48, 4.74	1.54, 4.06
		Median	2.20	2.20	2.20	2.80
		Q25, Q75	1.20, 4.40	1.40, 3.00	1.20, 5.00	1.40, 3.20
		Min, Max	1.0, 5.8	1.0, 6.4	1.0, 6.2	1.0, 6.4

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: The QSU-brief values at T<sub>0</sub> + 15/20 min refer to QSU-brief data at time T<sub>0</sub> + 15 min for THS 2.2 Menthol and mCC, and to QSU-brief data at time T<sub>0</sub> + 20 min for NRT gum.

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/000000106326/dev/tables/t\_qsu2.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.14.1 Descriptive Statistics of QSU-brief Questionnaire Factors and Total Score by Sex - PK Population**

Variable	Timepoint	Statistic	Group-2 PK			
			THS 2.2 Menthol		NRT gum	
			Males (N=9)	Females (N=9)	Males (N=9)	Females (N=9)
Factor 2 - Relief	T <sub>0</sub> + 4 h	n	9	9	9	9
		Mean (SD)	2.78 (1.928)	2.47 (1.604)	3.18 (2.078)	2.80 (1.770)
		95% CI	1.29, 4.26	1.23, 3.70	1.58, 4.78	1.44, 4.16
		Median	2.00	2.60	2.00	2.80
		Q25, Q75	1.20, 4.40	1.40, 2.80	1.40, 5.00	1.00, 3.40
		Min, Max	1.0, 6.0	1.0, 6.2	1.0, 6.2	1.0, 6.6
	T <sub>0</sub> + 6 h	n	9	9	9	9
		Mean (SD)	3.09 (1.958)	2.71 (1.638)	3.27 (2.173)	2.78 (1.740)
		95% CI	1.58, 4.60	1.45, 3.97	1.59, 4.94	1.44, 4.12
		Median	2.20	2.60	2.20	2.60
		Q25, Q75	1.20, 4.80	1.80, 3.00	1.40, 5.00	1.20, 3.80
		Min, Max	1.0, 6.2	1.0, 6.4	1.0, 6.4	1.0, 6.4

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: The QSU-brief values at T<sub>0</sub> + 15/20 min refer to QSU-brief data at time T<sub>0</sub> + 15 min for THS 2.2 Menthol and mCC, and to QSU-brief data at time T<sub>0</sub> + 20 min for NRT gum.

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/000000106326/dev/tables/t\_qsu2.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.14.1 Descriptive Statistics of QSU-brief Questionnaire Factors and Total Score by Sex - PK Population**

Variable	Timepoint	Statistic	Group-2 PK			
			THS 2.2 Menthol		NRT gum	
			Males (N=9)	Females (N=9)	Males (N=9)	Females (N=9)
Factor 2 - Relief	T <sub>0</sub> + 9 h	n	9	9	9	9
		Mean (SD)	3.07 (2.013)	2.71 (1.662)	3.20 (2.010)	2.69 (1.638)
		95% CI	1.51, 4.62	1.43, 3.99	1.65, 4.75	1.43, 3.95
		Median	3.00	2.60	2.20	2.60
		Q25, Q75	1.20, 3.40	1.20, 3.40	1.60, 4.60	1.20, 3.40
		Min, Max	1.0, 7.0	1.0, 6.2	1.0, 6.4	1.0, 6.2
	T <sub>0</sub> + 12 h	n	9	9	9	9
		Mean (SD)	3.18 (2.031)	2.67 (1.680)	3.33 (2.162)	2.73 (1.709)
		95% CI	1.61, 4.74	1.37, 3.96	1.67, 5.00	1.41, 4.05
		Median	3.00	2.80	3.00	2.60
		Q25, Q75	1.20, 4.80	1.00, 3.20	1.20, 5.20	1.20, 3.20
		Min, Max	1.0, 6.6	1.0, 6.0	1.0, 6.6	1.0, 6.4

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: The QSU-brief values at T<sub>0</sub> + 15/20 min refer to QSU-brief data at time T<sub>0</sub> + 15 min for THS 2.2 Menthol and mCC, and to QSU-brief data at time T<sub>0</sub> + 20 min for NRT gum.

Note: QSU-brief scores reported on a 7-point scale. Higher scores indicate greater intensity of urge.

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Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_qsu2.sas

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**Table 15.2.4.14.1 Descriptive Statistics of QSU-brief Questionnaire Factors and Total Score by Sex - PK Population**

Variable	Timepoint	Statistic	Group-2 PK			
			THS 2.2 Menthol		NRT gum	
			Males (N=9)	Females (N=9)	Males (N=9)	Females (N=9)
Total score	15 min < T <sub>0</sub>	n	9	9	9	9
		Mean (SD)	3.92 (2.131)	4.04 (1.751)	4.38 (1.829)	4.16 (1.267)
		95% CI	2.28, 5.56	2.69, 5.39	2.97, 5.79	3.18, 5.13
		Median	4.70	4.50	4.70	4.30
		Q25, Q75	1.70, 5.60	2.60, 5.00	2.80, 5.80	3.80, 5.00
		Min, Max	1.2, 6.7	1.4, 6.3	2.0, 7.0	1.9, 5.9
	T <sub>0</sub> + 15/20 min	n	9	9	9	9
		Mean (SD)	2.68 (1.832)	3.21 (2.008)	3.40 (2.049)	4.01 (1.616)
		95% CI	1.26, 4.09	1.66, 4.76	1.82, 4.98	2.76, 5.26
		Median	1.70	2.60	2.80	4.40
		Q25, Q75	1.50, 4.30	1.20, 4.50	1.70, 4.80	2.70, 4.70
		Min, Max	1.0, 5.8	1.1, 6.6	1.0, 6.5	1.4, 6.6

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: The QSU-brief values at T<sub>0</sub> + 15/20 min refer to QSU-brief data at time T<sub>0</sub> + 15 min for THS 2.2 Menthol and mCC, and to QSU-brief data at time T<sub>0</sub> + 20 min for NRT gum.

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/000000106326/dev/tables/t\_qsu2.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.14.1 Descriptive Statistics of QSU-brief Questionnaire Factors and Total Score by Sex - PK Population**

Variable	Timepoint	Statistic	Group-2 PK			
			THS 2.2 Menthol		NRT gum	
			Males (N=9)	Females (N=9)	Males (N=9)	Females (N=9)
Total score	T <sub>0</sub> + 30 min	n	9	9	9	9
		Mean (SD)	2.73 (1.818)	3.02 (1.969)	3.26 (2.176)	3.94 (1.818)
		95% CI	1.33, 4.14	1.50, 4.54	1.58, 4.93	2.54, 5.35
		Median	2.20	1.90	2.40	4.50
		Q25, Q75	1.30, 4.70	1.50, 4.40	1.40, 4.50	2.80, 5.20
		Min, Max	1.0, 5.7	1.0, 6.5	1.0, 6.9	1.1, 6.8
	T <sub>0</sub> + 45 min	n	9	9	9	9
		Mean (SD)	2.84 (1.951)	3.19 (1.917)	3.11 (2.093)	3.56 (2.000)
		95% CI	1.34, 4.35	1.71, 4.67	1.50, 4.72	2.01, 5.10
		Median	1.70	3.00	2.10	4.30
		Q25, Q75	1.30, 5.00	1.80, 4.20	1.60, 4.40	1.70, 4.70
		Min, Max	1.0, 5.8	1.1, 6.7	1.0, 6.5	1.0, 6.7

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: The QSU-brief values at T<sub>0</sub> + 15/20 min refer to QSU-brief data at time T<sub>0</sub> + 15 min for THS 2.2 Menthol and mCC, and to QSU-brief data at time T<sub>0</sub> + 20 min for NRT gum.

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/000000106326/dev/tables/t\_qsu2.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.14.1 Descriptive Statistics of QSU-brief Questionnaire Factors and Total Score by Sex - PK Population**

Variable	Timepoint	Statistic	Group-2 PK			
			THS 2.2 Menthol		NRT gum	
			Males (N=9)	Females (N=9)	Males (N=9)	Females (N=9)
Total score	T <sub>0</sub> + 60 min	n	9	9	9	9
		Mean (SD)	2.59 (1.551)	3.21 (1.898)	3.42 (2.301)	3.58 (2.094)
		95% CI	1.39, 3.79	1.75, 4.67	1.65, 5.20	1.96, 5.19
		Median	2.00	3.60	2.30	4.50
		Q25, Q75	1.20, 3.90	1.50, 4.00	1.60, 5.60	1.50, 5.00
		Min, Max	1.0, 5.0	1.2, 6.7	1.0, 6.5	1.0, 6.7
	T <sub>0</sub> + 2 h	n	9	9	9	9
		Mean (SD)	3.21 (2.064)	3.52 (1.943)	3.52 (2.125)	3.81 (1.807)
		95% CI	1.62, 4.80	2.02, 5.02	1.88, 5.16	2.42, 5.20
		Median	2.70	3.80	2.80	4.20
		Q25, Q75	1.40, 4.50	1.70, 5.00	2.10, 5.40	2.20, 4.90
		Min, Max	1.0, 6.3	1.5, 6.7	1.0, 6.5	1.0, 6.7

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: The QSU-brief values at T<sub>0</sub> + 15/20 min refer to QSU-brief data at time T<sub>0</sub> + 15 min for THS 2.2 Menthol and mCC, and to QSU-brief data at time T<sub>0</sub> + 20 min for NRT gum.

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/000000106326/dev/tables/t\_qsu2.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.14.1 Descriptive Statistics of QSU-brief Questionnaire Factors and Total Score by Sex - PK Population**

Variable	Timepoint	Statistic	Group-2 PK			
			THS 2.2 Menthol		NRT gum	
			Males (N=9)	Females (N=9)	Males (N=9)	Females (N=9)
Total score	T <sub>0</sub> + 4 h	n	9	9	9	9
		Mean (SD)	3.33 (1.952)	3.66 (1.807)	3.56 (2.095)	3.89 (2.014)
		95% CI	1.83, 4.84	2.26, 5.05	1.94, 5.17	2.34, 5.44
		Median	2.60	3.80	2.20	4.60
		Q25, Q75	2.00, 4.60	2.20, 4.80	1.90, 5.50	2.00, 5.10
		Min, Max	1.2, 6.5	1.0, 6.6	1.5, 6.5	1.0, 6.8
	T <sub>0</sub> + 6 h	n	9	9	9	9
		Mean (SD)	3.59 (2.021)	4.11 (1.614)	3.80 (2.115)	3.83 (1.969)
		95% CI	2.03, 5.15	2.87, 5.36	2.17, 5.43	2.32, 5.35
		Median	2.70	4.10	2.70	4.70
		Q25, Q75	2.20, 5.20	3.50, 5.00	2.30, 5.50	2.10, 4.90
		Min, Max	1.0, 6.5	1.3, 6.7	1.4, 6.6	1.0, 6.7

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: The QSU-brief values at T<sub>0</sub> + 15/20 min refer to QSU-brief data at time T<sub>0</sub> + 15 min for THS 2.2 Menthol and mCC, and to QSU-brief data at time T<sub>0</sub> + 20 min for NRT gum.

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/000000106326/dev/tables/t\_qsu2.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.14.1 Descriptive Statistics of QSU-brief Questionnaire Factors and Total Score by Sex - PK Population**

Variable	Timepoint	Statistic	Group-2 PK			
			THS 2.2 Menthol		NRT gum	
			Males (N=9)	Females (N=9)	Males (N=9)	Females (N=9)
Total score	T <sub>0</sub> + 9 h	n	9	9	9	9
		Mean (SD)	3.58 (1.981)	3.83 (1.849)	3.74 (1.919)	3.80 (1.936)
		95% CI	2.05, 5.11	2.41, 5.26	2.26, 5.22	2.31, 5.29
		Median	3.40	4.30	2.60	4.70
		Q25, Q75	2.30, 4.20	2.00, 5.20	2.30, 5.40	2.00, 5.00
		Min, Max	1.0, 7.0	1.3, 6.6	1.4, 6.6	1.0, 6.6
	T <sub>0</sub> + 12 h	n	9	9	9	9
		Mean (SD)	3.78 (1.961)	3.80 (1.866)	3.87 (2.058)	3.84 (1.952)
		95% CI	2.27, 5.29	2.36, 5.24	2.28, 5.45	2.34, 5.35
		Median	4.00	4.20	3.70	4.50
		Q25, Q75	2.60, 5.20	2.40, 5.10	2.30, 5.50	2.00, 5.00
		Min, Max	1.0, 6.7	1.1, 6.5	1.3, 6.7	1.1, 6.7

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: The QSU-brief values at T<sub>0</sub> + 15/20 min refer to QSU-brief data at time T<sub>0</sub> + 15 min for THS 2.2 Menthol and mCC, and to QSU-brief data at time T<sub>0</sub> + 20 min for NRT gum.

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/000000106326/dev/tables/t\_qsu2.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.14.2 Descriptive Statistics of QSU-brief Questionnaire Factors and Total Score by Nicotine Level - PK Population**

Variable	Timepoint	Statistic	Group-1 PK			
			THS 2.2 Menthol		mCC	
			<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)	<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)
Factor 1 - Reward	15 min < T <sub>0</sub>	n	25	18	25	18
		Mean (SD)	5.37 (1.222)	5.37 (0.964)	5.42 (1.299)	5.37 (0.746)
		95% CI	4.86, 5.88	4.88, 5.85	4.87, 5.96	4.99, 5.74
		Median	5.80	5.40	5.80	5.20
		Q25, Q75	4.60, 6.20	4.80, 5.80	4.60, 6.40	4.80, 5.80
		Min, Max	2.2, 7.0	2.8, 7.0	2.0, 7.0	4.2, 7.0
	T <sub>0</sub> + 15/20 min	n	25	18	25	18
		Mean (SD)	3.38 (1.791)	3.54 (1.600)	4.07 (1.800)	3.64 (1.460)
		95% CI	2.64, 4.13	2.74, 4.34	3.32, 4.82	2.91, 4.38
		Median	3.00	3.50	4.20	3.80
		Q25, Q75	2.20, 4.60	2.40, 5.00	2.80, 5.80	2.80, 4.60
		Min, Max	1.0, 7.0	1.2, 6.4	1.0, 6.6	1.0, 6.2

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: The QSU-brief values at T<sub>0</sub> + 15/20 min refer to QSU-brief data at time T<sub>0</sub> + 15 min for THS 2.2 Menthol and mCC, and to QSU-brief data at time T<sub>0</sub> + 20 min for NRT gum.

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/000000106326/dev/tables/t\_qsu3.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.14.2 Descriptive Statistics of QSU-brief Questionnaire Factors and Total Score by Nicotine Level - PK Population**

Variable	Timepoint	Statistic	Group-1 PK			
			THS 2.2 Menthol		mCC	
			<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)	<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)
Factor 1 - Reward	T <sub>0</sub> + 30 min	n	25	18	25	18
		Mean (SD)	3.25 (1.897)	3.80 (1.693)	3.97 (1.841)	3.72 (1.410)
		95% CI	2.46, 4.04	2.95, 4.65	3.20, 4.73	3.02, 4.43
		Median	2.80	4.00	3.60	4.10
		Q25, Q75	2.00, 4.40	2.40, 5.20	2.60, 6.00	2.60, 4.60
		Min, Max	1.0, 7.0	1.0, 6.4	1.0, 6.6	1.2, 6.0
	T <sub>0</sub> + 45 min	n	25	18	25	18
		Mean (SD)	3.31 (1.853)	4.17 (1.472)	4.10 (1.738)	4.01 (1.432)
		95% CI	2.54, 4.08	3.43, 4.90	3.37, 4.82	3.29, 4.73
		Median	3.00	4.30	4.00	4.10
		Q25, Q75	1.80, 5.00	3.00, 5.40	3.20, 5.60	3.20, 5.00
		Min, Max	1.0, 6.6	1.0, 6.6	1.0, 6.6	1.0, 6.6

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: The QSU-brief values at T<sub>0</sub> + 15/20 min refer to QSU-brief data at time T<sub>0</sub> + 15 min for THS 2.2 Menthol and mCC, and to QSU-brief data at time T<sub>0</sub> + 20 min for NRT gum.

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/000000106326/dev/tables/t\_qsu3.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.14.2 Descriptive Statistics of QSU-brief Questionnaire Factors and Total Score by Nicotine Level - PK Population**

Variable	Timepoint	Statistic	Group-1 PK			
			THS 2.2 Menthol		mCC	
			<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)	<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)
Factor 1 - Reward	T <sub>0</sub> + 60 min	n	25	18	25	18
		Mean (SD)	3.52 (1.956)	4.11 (1.604)	4.15 (1.837)	4.27 (1.483)
		95% CI	2.71, 4.33	3.31, 4.91	3.39, 4.91	3.52, 5.01
		Median	3.40	4.10	4.20	4.20
		Q25, Q75	1.80, 5.20	2.80, 5.60	3.00, 6.00	3.20, 5.40
		Min, Max	1.0, 6.8	1.0, 6.4	1.0, 7.0	1.0, 6.6
	T <sub>0</sub> + 2 h	n	25	18	25	18
		Mean (SD)	3.89 (1.979)	4.51 (1.477)	4.43 (1.602)	4.44 (1.465)
		95% CI	3.07, 4.71	3.77, 5.25	3.77, 5.10	3.71, 5.18
		Median	4.40	4.20	4.80	4.20
		Q25, Q75	2.00, 5.60	3.80, 5.60	3.80, 5.40	3.60, 5.40
		Min, Max	1.0, 7.0	1.0, 6.6	1.0, 6.8	1.0, 6.6

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: The QSU-brief values at T<sub>0</sub> + 15/20 min refer to QSU-brief data at time T<sub>0</sub> + 15 min for THS 2.2 Menthol and mCC, and to QSU-brief data at time T<sub>0</sub> + 20 min for NRT gum.

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/000000106326/dev/tables/t\_qsu3.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.14.2 Descriptive Statistics of QSU-brief Questionnaire Factors and Total Score by Nicotine Level - PK Population**

Variable	Timepoint	Statistic	Group-1 PK			
			THS 2.2 Menthol		mCC	
			<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)	<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)
Factor 1 - Reward	T <sub>0</sub> + 4 h	n	25	18	25	18
		Mean (SD)	4.14 (1.688)	4.52 (1.369)	4.54 (1.634)	4.58 (1.401)
		95% CI	3.43, 4.84	3.84, 5.21	3.86, 5.22	3.88, 5.28
		Median	4.00	4.30	4.60	4.40
		Q25, Q75	2.80, 5.60	4.00, 5.60	3.40, 5.80	4.20, 5.60
		Min, Max	1.0, 7.0	1.0, 6.6	1.4, 6.6	1.0, 6.6
	T <sub>0</sub> + 6 h	n	25	18	25	18
		Mean (SD)	4.59 (1.884)	4.80 (1.324)	5.09 (1.655)	5.02 (1.374)
		95% CI	3.81, 5.37	4.14, 5.46	4.40, 5.78	4.33, 5.71
		Median	5.00	4.60	5.80	4.90
		Q25, Q75	2.80, 6.60	4.20, 5.60	4.20, 6.60	4.40, 6.00
		Min, Max	1.0, 7.0	1.0, 6.8	1.6, 7.0	1.0, 7.0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: The QSU-brief values at T<sub>0</sub> + 15/20 min refer to QSU-brief data at time T<sub>0</sub> + 15 min for THS 2.2 Menthol and mCC, and to QSU-brief data at time T<sub>0</sub> + 20 min for NRT gum.

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/000000106326/dev/tables/t\_qsu3.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.14.2 Descriptive Statistics of QSU-brief Questionnaire Factors and Total Score by Nicotine Level - PK Population**

Variable	Timepoint	Statistic	Group-1 PK			
			THS 2.2 Menthol		mCC	
			<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)	<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)
Factor 1 - Reward	T <sub>0</sub> + 9 h	n	25	18	25	18
		Mean (SD)	4.70 (1.890)	4.99 (0.864)	4.94 (1.536)	5.02 (1.329)
		95% CI	3.91, 5.48	4.55, 5.42	4.30, 5.57	4.36, 5.69
		Median	5.40	5.00	5.20	5.10
		Q25, Q75	3.40, 6.40	4.40, 5.60	4.20, 6.20	4.60, 5.60
		Min, Max	1.0, 6.8	3.0, 6.4	1.4, 6.6	1.0, 7.0
	T <sub>0</sub> + 12 h	n	25	18	25	18
		Mean (SD)	4.78 (1.866)	5.06 (1.150)	5.05 (1.616)	5.13 (1.214)
		95% CI	4.00, 5.55	4.48, 5.63	4.38, 5.72	4.52, 5.74
		Median	5.40	5.20	5.60	5.20
		Q25, Q75	3.40, 6.00	4.40, 5.80	4.40, 6.20	4.60, 5.80
		Min, Max	1.0, 7.0	2.0, 6.8	1.4, 7.0	1.6, 7.0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: The QSU-brief values at T<sub>0</sub> + 15/20 min refer to QSU-brief data at time T<sub>0</sub> + 15 min for THS 2.2 Menthol and mCC, and to QSU-brief data at time T<sub>0</sub> + 20 min for NRT gum.

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/000000106326/dev/tables/t\_qsu3.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.14.2 Descriptive Statistics of QSU-brief Questionnaire Factors and Total Score by Nicotine Level - PK Population**

Variable	Timepoint	Statistic	Group-1 PK			
			THS 2.2 Menthol		mCC	
			<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)	<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)
Factor 2 - Relief	15 min < T <sub>0</sub>	n	25	18	25	18
		Mean (SD)	3.30 (1.247)	3.43 (1.205)	3.54 (1.406)	3.46 (1.250)
		95% CI	2.78, 3.82	2.83, 4.04	2.96, 4.13	2.83, 4.08
		Median	3.40	3.40	3.40	3.30
		Q25, Q75	2.80, 4.20	2.60, 4.40	2.40, 4.40	2.60, 4.80
		Min, Max	1.0, 5.6	1.0, 5.2	1.0, 6.0	1.2, 5.4
	T <sub>0</sub> + 15/20 min	n	25	18	25	18
		Mean (SD)	2.18 (1.041)	2.30 (1.177)	2.65 (1.228)	2.59 (1.162)
		95% CI	1.75, 2.62	1.71, 2.89	2.14, 3.16	2.01, 3.17
		Median	2.00	2.10	2.40	2.40
		Q25, Q75	1.40, 2.60	1.20, 3.00	1.60, 3.60	1.80, 3.00
		Min, Max	1.0, 4.6	1.0, 4.8	1.0, 4.8	1.0, 4.8

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: The QSU-brief values at T<sub>0</sub> + 15/20 min refer to QSU-brief data at time T<sub>0</sub> + 15 min for THS 2.2 Menthol and mCC, and to QSU-brief data at time T<sub>0</sub> + 20 min for NRT gum.

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/000000106326/dev/tables/t\_qsu3.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.14.2 Descriptive Statistics of QSU-brief Questionnaire Factors and Total Score by Nicotine Level - PK Population**

Variable	Timepoint	Statistic	Group-1 PK			
			THS 2.2 Menthol		mCC	
			<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)	<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)
Factor 2 - Relief	T <sub>0</sub> + 30 min	n	25	18	25	18
		Mean (SD)	2.08 (0.992)	2.37 (1.284)	2.44 (1.184)	2.52 (1.118)
		95% CI	1.67, 2.49	1.72, 3.01	1.95, 2.93	1.96, 3.08
		Median	2.00	2.20	2.20	2.30
		Q25, Q75	1.20, 2.60	1.00, 3.20	1.60, 2.80	1.80, 3.20
		Min, Max	1.0, 4.4	1.0, 5.0	1.0, 5.2	1.0, 4.4
	T <sub>0</sub> + 45 min	n	25	18	25	18
		Mean (SD)	2.16 (0.997)	2.33 (1.247)	2.58 (1.335)	2.63 (1.092)
		95% CI	1.74, 2.58	1.71, 2.96	2.02, 3.13	2.09, 3.18
		Median	2.00	2.20	2.20	2.70
		Q25, Q75	1.20, 2.80	1.00, 3.00	1.60, 3.00	1.80, 3.20
		Min, Max	1.0, 4.4	1.0, 5.2	1.0, 5.8	1.0, 4.8

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: The QSU-brief values at T<sub>0</sub> + 15/20 min refer to QSU-brief data at time T<sub>0</sub> + 15 min for THS 2.2 Menthol and mCC, and to QSU-brief data at time T<sub>0</sub> + 20 min for NRT gum.

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/000000106326/dev/tables/t\_qsu3.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.14.2 Descriptive Statistics of QSU-brief Questionnaire Factors and Total Score by Nicotine Level - PK Population**

Variable	Timepoint	Statistic	Group-1 PK			
			THS 2.2 Menthol		mCC	
			<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)	<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)
Factor 2 - Relief	T <sub>0</sub> + 60 min	n	25	18	25	18
		Mean (SD)	2.34 (1.244)	2.39 (1.335)	2.59 (1.288)	2.83 (1.266)
		95% CI	1.82, 2.85	1.72, 3.06	2.06, 3.13	2.20, 3.47
		Median	2.20	2.20	2.40	2.80
		Q25, Q75	1.40, 3.00	1.00, 3.40	1.80, 3.00	1.80, 4.00
		Min, Max	1.0, 5.8	1.0, 5.0	1.0, 5.4	1.0, 4.8
	T <sub>0</sub> + 2 h	n	25	18	25	18
		Mean (SD)	2.47 (1.317)	2.66 (1.243)	2.69 (1.223)	2.70 (1.217)
		95% CI	1.92, 3.02	2.03, 3.28	2.18, 3.20	2.09, 3.31
		Median	2.20	2.50	2.60	2.50
		Q25, Q75	1.20, 3.00	1.80, 3.60	1.80, 3.60	1.80, 4.00
		Min, Max	1.0, 5.2	1.0, 4.8	1.0, 4.8	1.0, 4.8

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: The QSU-brief values at T<sub>0</sub> + 15/20 min refer to QSU-brief data at time T<sub>0</sub> + 15 min for THS 2.2 Menthol and mCC, and to QSU-brief data at time T<sub>0</sub> + 20 min for NRT gum.

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/000000106326/dev/tables/t\_qsu3.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.14.2 Descriptive Statistics of QSU-brief Questionnaire Factors and Total Score by Nicotine Level - PK Population**

Variable	Timepoint	Statistic	Group-1 PK			
			THS 2.2 Menthol		mCC	
			<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)	<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)
Factor 2 - Relief	T <sub>0</sub> + 4 h	n	25	18	25	18
		Mean (SD)	2.64 (1.138)	2.67 (1.199)	2.92 (1.295)	2.76 (1.202)
		95% CI	2.17, 3.11	2.07, 3.27	2.38, 3.46	2.15, 3.36
		Median	3.00	2.70	2.80	2.50
		Q25, Q75	1.60, 3.20	1.60, 3.20	2.00, 4.00	1.80, 4.00
		Min, Max	1.0, 4.6	1.0, 5.4	1.0, 5.0	1.0, 4.8
	T <sub>0</sub> + 6 h	n	25	18	25	18
		Mean (SD)	2.92 (1.509)	2.91 (1.234)	3.15 (1.533)	3.03 (1.299)
		95% CI	2.29, 3.55	2.29, 3.53	2.51, 3.79	2.38, 3.68
		Median	3.00	3.00	2.80	3.30
		Q25, Q75	1.60, 4.00	2.20, 3.60	2.20, 4.20	2.20, 3.80
		Min, Max	1.0, 6.2	1.0, 5.6	1.0, 6.8	1.0, 5.6

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: The QSU-brief values at T<sub>0</sub> + 15/20 min refer to QSU-brief data at time T<sub>0</sub> + 15 min for THS 2.2 Menthol and mCC, and to QSU-brief data at time T<sub>0</sub> + 20 min for NRT gum.

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/000000106326/dev/tables/t\_qsu3.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.14.2 Descriptive Statistics of QSU-brief Questionnaire Factors and Total Score by Nicotine Level - PK Population**

Variable	Timepoint	Statistic	Group-1 PK			
			THS 2.2 Menthol		mCC	
			<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)	<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)
Factor 2 - Relief	T <sub>0</sub> + 9 h	n	25	18	25	18
		Mean (SD)	2.99 (1.568)	2.93 (1.052)	3.08 (1.443)	3.16 (1.285)
		95% CI	2.34, 3.64	2.41, 3.46	2.48, 3.68	2.51, 3.80
		Median	2.60	2.80	3.00	3.40
		Q25, Q75	1.60, 4.40	2.60, 3.60	2.20, 4.20	2.40, 3.60
		Min, Max	1.0, 6.0	1.0, 5.0	1.0, 6.0	1.0, 5.4
	T <sub>0</sub> + 12 h	n	25	18	25	18
		Mean (SD)	3.02 (1.525)	2.90 (1.277)	3.13 (1.527)	3.19 (1.234)
		95% CI	2.38, 3.65	2.26, 3.54	2.49, 3.76	2.57, 3.81
		Median	2.80	2.80	3.20	3.40
		Q25, Q75	1.80, 3.80	2.20, 4.00	2.20, 4.20	2.40, 4.00
		Min, Max	1.0, 6.4	1.0, 5.8	1.0, 6.4	1.0, 5.2

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: The QSU-brief values at T<sub>0</sub> + 15/20 min refer to QSU-brief data at time T<sub>0</sub> + 15 min for THS 2.2 Menthol and mCC, and to QSU-brief data at time T<sub>0</sub> + 20 min for NRT gum.

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/000000106326/dev/tables/t\_qsu3.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.14.2 Descriptive Statistics of QSU-brief Questionnaire Factors and Total Score by Nicotine Level - PK Population**

Variable	Timepoint	Statistic	Group-1 PK			
			THS 2.2 Menthol		mCC	
			<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)	<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)
Total score	15 min < T <sub>0</sub>	n	25	18	25	18
		Mean (SD)	4.33 (1.113)	4.40 (1.013)	4.48 (1.281)	4.41 (0.918)
		95% CI	3.87, 4.80	3.89, 4.91	3.95, 5.01	3.95, 4.87
		Median	4.60	4.45	4.60	4.35
		Q25, Q75	3.70, 5.20	4.10, 5.20	3.70, 5.30	3.70, 5.20
		Min, Max	1.6, 6.3	2.2, 6.1	1.5, 6.3	3.2, 6.2
	T <sub>0</sub> + 15/20 min	n	25	18	25	18
		Mean (SD)	2.78 (1.298)	2.92 (1.327)	3.36 (1.441)	3.12 (1.281)
		95% CI	2.24, 3.32	2.26, 3.59	2.76, 3.96	2.47, 3.76
		Median	2.50	2.90	3.40	3.30
		Q25, Q75	1.70, 3.80	1.80, 3.70	2.20, 4.50	2.20, 3.80
		Min, Max	1.0, 5.1	1.3, 5.6	1.0, 5.5	1.0, 5.5

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: The QSU-brief values at T<sub>0</sub> + 15/20 min refer to QSU-brief data at time T<sub>0</sub> + 15 min for THS 2.2 Menthol and mCC, and to QSU-brief data at time T<sub>0</sub> + 20 min for NRT gum.

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/000000106326/dev/tables/t\_qsu3.sas

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**Table 15.2.4.14.2 Descriptive Statistics of QSU-brief Questionnaire Factors and Total Score by Nicotine Level - PK Population**

Variable	Timepoint	Statistic	Group-1 PK			
			THS 2.2 Menthol		mCC	
			<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)	<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)
Total score	T <sub>0</sub> + 30 min	n	25	18	25	18
		Mean (SD)	2.66 (1.348)	3.08 (1.396)	3.20 (1.420)	3.12 (1.216)
		95% CI	2.10, 3.23	2.38, 3.78	2.61, 3.80	2.51, 3.73
		Median	2.40	3.20	3.00	3.00
		Q25, Q75	1.60, 3.60	1.90, 4.20	2.40, 4.50	2.20, 4.00
		Min, Max	1.0, 5.2	1.0, 5.7	1.0, 5.6	1.1, 5.1
	T <sub>0</sub> + 45 min	n	25	18	25	18
		Mean (SD)	2.74 (1.346)	3.25 (1.222)	3.34 (1.433)	3.32 (1.219)
		95% CI	2.18, 3.30	2.64, 3.86	2.74, 3.93	2.71, 3.93
		Median	2.60	3.15	3.30	3.20
		Q25, Q75	1.60, 3.80	2.60, 4.10	2.60, 4.50	2.60, 4.20
		Min, Max	1.0, 5.2	1.0, 5.9	1.0, 6.1	1.0, 5.5

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: The QSU-brief values at T<sub>0</sub> + 15/20 min refer to QSU-brief data at time T<sub>0</sub> + 15 min for THS 2.2 Menthol and mCC, and to QSU-brief data at time T<sub>0</sub> + 20 min for NRT gum.

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/000000106326/dev/tables/t\_qsu3.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.14.2 Descriptive Statistics of QSU-brief Questionnaire Factors and Total Score by Nicotine Level - PK Population**

Variable	Timepoint	Statistic	Group-1 PK			
			THS 2.2 Menthol		mCC	
			<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)	<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)
Total score	T <sub>0</sub> + 60 min	n	25	18	25	18
		Mean (SD)	2.93 (1.492)	3.25 (1.341)	3.37 (1.466)	3.55 (1.344)
		95% CI	2.31, 3.55	2.58, 3.92	2.76, 3.98	2.88, 4.22
		Median	3.10	3.30	3.20	3.40
		Q25, Q75	1.50, 4.20	2.00, 3.90	2.50, 4.60	2.60, 4.70
		Min, Max	1.0, 5.8	1.0, 5.7	1.0, 5.9	1.0, 5.6
	T <sub>0</sub> + 2 h	n	25	18	25	18
		Mean (SD)	3.18 (1.552)	3.58 (1.209)	3.56 (1.286)	3.57 (1.240)
		95% CI	2.53, 3.83	2.98, 4.19	3.02, 4.10	2.95, 4.19
		Median	3.30	3.40	3.90	3.40
		Q25, Q75	1.60, 4.40	3.00, 4.20	2.60, 4.50	2.60, 4.60
		Min, Max	1.0, 5.6	1.0, 5.7	1.0, 5.2	1.0, 5.7

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: The QSU-brief values at T<sub>0</sub> + 15/20 min refer to QSU-brief data at time T<sub>0</sub> + 15 min for THS 2.2 Menthol and mCC, and to QSU-brief data at time T<sub>0</sub> + 20 min for NRT gum.

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/000000106326/dev/tables/t\_qsu3.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.14.2 Descriptive Statistics of QSU-brief Questionnaire Factors and Total Score by Nicotine Level - PK Population**

Variable	Timepoint	Statistic	Group-1 PK			
			THS 2.2 Menthol		mCC	
			<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)	<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)
Total score	T <sub>0</sub> + 4 h	n	25	18	25	18
		Mean (SD)	3.39 (1.322)	3.59 (1.194)	3.73 (1.374)	3.67 (1.209)
		95% CI	2.84, 3.94	3.00, 4.19	3.16, 4.30	3.06, 4.27
		Median	3.50	3.35	4.20	3.55
		Q25, Q75	2.50, 4.60	3.00, 4.10	2.70, 4.70	2.80, 4.60
		Min, Max	1.0, 5.2	1.0, 6.0	1.2, 5.6	1.0, 5.6
	T <sub>0</sub> + 6 h	n	25	18	25	18
		Mean (SD)	3.76 (1.572)	3.86 (1.152)	4.12 (1.483)	4.03 (1.267)
		95% CI	3.10, 4.41	3.28, 4.43	3.50, 4.74	3.39, 4.66
		Median	4.50	3.80	4.50	4.15
		Q25, Q75	2.30, 4.90	3.30, 4.60	3.10, 5.10	3.20, 4.80
		Min, Max	1.0, 6.4	1.0, 6.2	1.3, 6.7	1.0, 6.2

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: The QSU-brief values at T<sub>0</sub> + 15/20 min refer to QSU-brief data at time T<sub>0</sub> + 15 min for THS 2.2 Menthol and mCC, and to QSU-brief data at time T<sub>0</sub> + 20 min for NRT gum.

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/000000106326/dev/tables/t\_qsu3.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.14.2 Descriptive Statistics of QSU-brief Questionnaire Factors and Total Score by Nicotine Level - PK Population**

Variable	Timepoint	Statistic	Group-1 PK			
			THS 2.2 Menthol		mCC	
			<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)	<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)
Total score	T <sub>0</sub> + 9 h	n	25	18	25	18
		Mean (SD)	3.84 (1.603)	3.96 (0.819)	4.01 (1.372)	4.09 (1.222)
		95% CI	3.18, 4.51	3.55, 4.37	3.44, 4.58	3.48, 4.70
		Median	4.40	3.85	4.50	4.20
		Q25, Q75	2.20, 5.20	3.50, 4.20	3.30, 4.90	3.40, 4.70
		Min, Max	1.0, 6.3	2.0, 5.7	1.2, 6.3	1.0, 6.2
	T <sub>0</sub> + 12 h	n	25	18	25	18
		Mean (SD)	3.90 (1.592)	3.98 (1.081)	4.09 (1.458)	4.16 (1.137)
		95% CI	3.23, 4.56	3.44, 4.52	3.48, 4.69	3.59, 4.73
		Median	4.20	3.90	4.40	4.15
		Q25, Q75	2.60, 4.90	3.40, 4.60	3.30, 5.10	3.40, 4.80
		Min, Max	1.0, 6.7	1.5, 6.3	1.2, 6.7	1.3, 5.8

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: The QSU-brief values at T<sub>0</sub> + 15/20 min refer to QSU-brief data at time T<sub>0</sub> + 15 min for THS 2.2 Menthol and mCC, and to QSU-brief data at time T<sub>0</sub> + 20 min for NRT gum.

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/000000106326/dev/tables/t\_qsu3.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.14.2 Descriptive Statistics of QSU-brief Questionnaire Factors and Total Score by Nicotine Level - PK Population**

Variable	Timepoint	Statistic	Group-2 PK			
			THS 2.2 Menthol		NRT gum	
			<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)	<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)
Factor 1 - Reward	15 min < T <sub>0</sub>	n	10	8	10	8
		Mean (SD)	5.06 (1.958)	4.50 (2.434)	5.52 (1.348)	4.75 (1.763)
		95% CI	3.65, 6.47	2.46, 6.54	4.55, 6.49	3.27, 6.23
		Median	5.40	4.50	5.40	4.50
		Q25, Q75	4.20, 6.80	2.30, 7.00	5.00, 6.60	3.60, 6.40
		Min, Max	1.8, 7.0	1.4, 7.0	2.4, 7.0	2.0, 7.0
	T <sub>0</sub> + 15/20 min	n	10	8	10	8
		Mean (SD)	3.78 (2.124)	3.40 (2.393)	4.88 (1.851)	3.88 (2.550)
		95% CI	2.26, 5.30	1.39, 5.41	3.55, 6.21	1.74, 6.01
		Median	4.00	2.50	5.30	2.70
		Q25, Q75	1.80, 5.20	1.40, 5.70	3.40, 6.60	2.00, 6.80
		Min, Max	1.2, 7.0	1.0, 7.0	1.6, 7.0	1.0, 7.0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: The QSU-brief values at T<sub>0</sub> + 15/20 min refer to QSU-brief data at time T<sub>0</sub> + 15 min for THS 2.2 Menthol and mCC, and to QSU-brief data at time T<sub>0</sub> + 20 min for NRT gum.

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/000000106326/dev/tables/t\_qsu3.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.14.2 Descriptive Statistics of QSU-brief Questionnaire Factors and Total Score by Nicotine Level - PK Population**

Variable	Timepoint	Statistic	Group-2 PK			
			THS 2.2 Menthol		NRT gum	
			<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)	<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)
Factor 1 - Reward	T <sub>0</sub> + 30 min	n	10	8	10	8
		Mean (SD)	3.56 (2.140)	3.55 (2.532)	4.74 (1.949)	3.70 (2.772)
		95% CI	2.02, 5.10	1.43, 5.67	3.34, 6.14	1.38, 6.02
		Median	2.80	2.60	4.90	2.30
		Q25, Q75	1.80, 5.20	1.40, 6.20	3.60, 6.60	1.50, 7.00
		Min, Max	1.0, 7.0	1.0, 7.0	1.2, 7.0	1.0, 7.0
	T <sub>0</sub> + 45 min	n	10	8	10	8
		Mean (SD)	3.98 (1.979)	3.15 (2.470)	4.28 (2.220)	3.50 (2.595)
		95% CI	2.56, 5.40	1.08, 5.22	2.69, 5.87	1.33, 5.67
		Median	3.90	1.60	4.40	2.20
		Q25, Q75	2.20, 5.40	1.40, 5.60	2.40, 6.40	1.50, 6.30
		Min, Max	1.2, 7.0	1.0, 7.0	1.0, 7.0	1.0, 7.0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: The QSU-brief values at T<sub>0</sub> + 15/20 min refer to QSU-brief data at time T<sub>0</sub> + 15 min for THS 2.2 Menthol and mCC, and to QSU-brief data at time T<sub>0</sub> + 20 min for NRT gum.

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/000000106326/dev/tables/t\_qsu3.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.14.2 Descriptive Statistics of QSU-brief Questionnaire Factors and Total Score by Nicotine Level - PK Population**

Variable	Timepoint	Statistic	Group-2 PK			
			THS 2.2 Menthol		NRT gum	
			<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)	<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)
Factor 1 - Reward	T <sub>0</sub> + 60 min	n	10	8	10	8
		Mean (SD)	3.90 (1.988)	3.35 (2.388)	4.48 (2.408)	3.53 (2.619)
		95% CI	2.47, 5.33	1.35, 5.35	2.75, 6.21	1.33, 5.72
		Median	4.00	2.30	5.20	2.20
		Q25, Q75	2.00, 5.40	1.40, 5.70	2.60, 6.60	1.50, 6.40
		Min, Max	1.4, 7.0	1.0, 7.0	1.0, 7.0	1.0, 7.0
	T <sub>0</sub> + 2 h	n	10	8	10	8
		Mean (SD)	4.40 (2.060)	3.60 (2.643)	4.84 (1.982)	3.80 (2.412)
		95% CI	2.92, 5.88	1.39, 5.81	3.42, 6.26	1.78, 5.82
		Median	4.40	2.30	5.40	3.10
		Q25, Q75	2.20, 6.80	1.50, 6.60	3.40, 6.40	1.90, 6.20
		Min, Max	1.8, 7.0	1.0, 7.0	1.0, 7.0	1.0, 7.0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: The QSU-brief values at T<sub>0</sub> + 15/20 min refer to QSU-brief data at time T<sub>0</sub> + 15 min for THS 2.2 Menthol and mCC, and to QSU-brief data at time T<sub>0</sub> + 20 min for NRT gum.

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/000000106326/dev/tables/t\_qsu3.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.14.2 Descriptive Statistics of QSU-brief Questionnaire Factors and Total Score by Nicotine Level - PK Population**

Variable	Timepoint	Statistic	Group-2 PK			
			THS 2.2 Menthol		NRT gum	
			<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)	<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)
Factor 1 - Reward	T <sub>0</sub> + 4 h	n	10	8	10	8
		Mean (SD)	4.50 (2.030)	4.20 (2.427)	4.80 (2.249)	4.03 (2.499)
		95% CI	3.04, 5.96	2.17, 6.23	3.19, 6.41	1.93, 6.12
		Median	4.80	3.60	5.70	2.80
		Q25, Q75	2.80, 6.60	2.20, 6.90	2.60, 6.40	2.00, 7.00
		Min, Max	1.0, 7.0	1.2, 7.0	1.0, 7.0	1.6, 7.0
	T <sub>0</sub> + 6 h	n	10	8	10	8
		Mean (SD)	5.06 (1.752)	4.48 (2.505)	4.90 (2.224)	4.25 (2.354)
		95% CI	3.80, 6.32	2.38, 6.57	3.30, 6.50	2.28, 6.22
		Median	5.50	4.50	5.60	3.30
		Q25, Q75	3.80, 6.80	2.40, 7.00	3.20, 6.80	2.30, 7.00
		Min, Max	1.6, 7.0	1.0, 7.0	1.0, 7.0	1.8, 7.0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: The QSU-brief values at T<sub>0</sub> + 15/20 min refer to QSU-brief data at time T<sub>0</sub> + 15 min for THS 2.2 Menthol and mCC, and to QSU-brief data at time T<sub>0</sub> + 20 min for NRT gum.

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/000000106326/dev/tables/t\_qsu3.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.14.2 Descriptive Statistics of QSU-brief Questionnaire Factors and Total Score by Nicotine Level - PK Population**

Variable	Timepoint	Statistic	Group-2 PK			
			THS 2.2 Menthol		NRT gum	
			<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)	<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)
Factor 1 - Reward	T <sub>0</sub> + 9 h	n	10	8	10	8
		Mean (SD)	4.82 (1.885)	4.15 (2.468)	4.80 (2.165)	4.35 (2.250)
		95% CI	3.47, 6.17	2.08, 6.22	3.25, 6.35	2.46, 6.24
		Median	4.90	3.20	5.50	3.30
		Q25, Q75	3.60, 6.80	2.40, 7.00	3.00, 6.40	2.70, 7.00
		Min, Max	1.4, 7.0	1.0, 7.0	1.0, 7.0	1.8, 7.0
	T <sub>0</sub> + 12 h	n	10	8	10	8
		Mean (SD)	4.88 (1.940)	4.38 (2.319)	5.04 (2.070)	4.23 (2.409)
		95% CI	3.49, 6.27	2.43, 6.32	3.55, 6.53	2.21, 6.24
		Median	5.30	3.90	5.70	3.40
		Q25, Q75	3.40, 6.80	2.70, 6.90	4.40, 6.40	2.20, 7.00
		Min, Max	1.2, 7.0	1.0, 7.0	1.2, 7.0	1.6, 7.0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: The QSU-brief values at T<sub>0</sub> + 15/20 min refer to QSU-brief data at time T<sub>0</sub> + 15 min for THS 2.2 Menthol and mCC, and to QSU-brief data at time T<sub>0</sub> + 20 min for NRT gum.

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/000000106326/dev/tables/t\_qsu3.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.14.2 Descriptive Statistics of QSU-brief Questionnaire Factors and Total Score by Nicotine Level - PK Population**

Variable	Timepoint	Statistic	Group-2 PK			
			THS 2.2 Menthol		NRT gum	
			<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)	<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)
Factor 2 - Relief	15 min < T <sub>0</sub>	n	10	8	10	8
		Mean (SD)	3.78 (2.043)	2.38 (1.320)	4.16 (1.791)	2.35 (1.002)
		95% CI	2.31, 5.25	1.27, 3.48	2.87, 5.45	1.51, 3.19
		Median	4.00	2.30	4.30	2.40
		Q25, Q75	1.60, 5.60	1.10, 3.30	2.60, 5.20	1.50, 3.10
		Min, Max	1.0, 6.4	1.0, 4.6	1.4, 7.0	1.0, 3.8
	T <sub>0</sub> + 15/20 min	n	10	8	10	8
		Mean (SD)	2.70 (2.054)	1.75 (1.204)	3.96 (1.676)	1.75 (0.784)
		95% CI	1.23, 4.17	0.74, 2.76	2.76, 5.16	1.09, 2.41
		Median	1.40	1.10	4.10	1.50
		Q25, Q75	1.00, 4.80	1.00, 2.30	2.80, 4.80	1.10, 2.30
		Min, Max	1.0, 6.2	1.0, 4.2	1.2, 6.4	1.0, 3.2

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: The QSU-brief values at T<sub>0</sub> + 15/20 min refer to QSU-brief data at time T<sub>0</sub> + 15 min for THS 2.2 Menthol and mCC, and to QSU-brief data at time T<sub>0</sub> + 20 min for NRT gum.

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/000000106326/dev/tables/t\_qsu3.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.14.2 Descriptive Statistics of QSU-brief Questionnaire Factors and Total Score by Nicotine Level - PK Population**

Variable	Timepoint	Statistic	Group-2 PK			
			THS 2.2 Menthol		NRT gum	
			<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)	<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)
Factor 2 - Relief	T <sub>0</sub> + 30 min	n	10	8	10	8
		Mean (SD)	2.72 (1.985)	1.55 (0.777)	3.82 (1.885)	1.80 (1.053)
		95% CI	1.30, 4.14	0.90, 2.20	2.47, 5.17	0.91, 2.69
		Median	1.50	1.10	3.80	1.30
		Q25, Q75	1.20, 4.40	1.00, 2.10	2.20, 4.80	1.00, 2.70
		Min, Max	1.0, 6.0	1.0, 3.0	1.0, 6.8	1.0, 3.4
	T <sub>0</sub> + 45 min	n	10	8	10	8
		Mean (SD)	3.06 (2.012)	1.63 (0.898)	3.50 (2.003)	1.78 (1.000)
		95% CI	1.62, 4.50	0.87, 2.38	2.06, 4.94	0.93, 2.62
		Median	2.40	1.10	3.70	1.20
		Q25, Q75	1.20, 4.80	1.00, 2.20	1.80, 4.80	1.00, 2.70
		Min, Max	1.0, 6.4	1.0, 3.4	1.0, 6.4	1.0, 3.4

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: The QSU-brief values at T<sub>0</sub> + 15/20 min refer to QSU-brief data at time T<sub>0</sub> + 15 min for THS 2.2 Menthol and mCC, and to QSU-brief data at time T<sub>0</sub> + 20 min for NRT gum.

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/000000106326/dev/tables/t\_qsu3.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.14.2 Descriptive Statistics of QSU-brief Questionnaire Factors and Total Score by Nicotine Level - PK Population**

Variable	Timepoint	Statistic	Group-2 PK			
			THS 2.2 Menthol		NRT gum	
			<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)	<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)
Factor 2 - Relief	T <sub>0</sub> + 60 min	n	10	8	10	8
		Mean (SD)	2.58 (1.887)	1.60 (0.726)	3.74 (2.159)	1.95 (1.074)
		95% CI	1.23, 3.93	0.99, 2.21	2.19, 5.29	1.05, 2.85
		Median	1.70	1.30	3.70	1.50
		Q25, Q75	1.00, 4.40	1.00, 2.20	2.00, 6.20	1.00, 3.10
		Min, Max	1.0, 6.4	1.0, 2.8	1.0, 6.4	1.0, 3.4
	T <sub>0</sub> + 2 h	n	10	8	10	8
		Mean (SD)	3.36 (2.018)	1.85 (1.013)	3.80 (1.956)	1.90 (0.998)
		95% CI	1.91, 4.81	1.00, 2.70	2.40, 5.20	1.06, 2.74
		Median	3.00	1.30	3.80	1.40
		Q25, Q75	1.40, 5.20	1.10, 2.70	2.20, 5.40	1.10, 2.90
		Min, Max	1.0, 6.4	1.0, 3.6	1.0, 6.4	1.0, 3.4

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: The QSU-brief values at T<sub>0</sub> + 15/20 min refer to QSU-brief data at time T<sub>0</sub> + 15 min for THS 2.2 Menthol and mCC, and to QSU-brief data at time T<sub>0</sub> + 20 min for NRT gum.

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/000000106326/dev/tables/t\_qsu3.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.14.2 Descriptive Statistics of QSU-brief Questionnaire Factors and Total Score by Nicotine Level - PK Population**

Variable	Timepoint	Statistic	Group-2 PK			
			THS 2.2 Menthol		NRT gum	
			<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)	<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)
Factor 2 - Relief	T <sub>0</sub> + 4 h	n	10	8	10	8
		Mean (SD)	3.26 (2.018)	1.83 (0.838)	3.78 (2.084)	2.00 (1.003)
		95% CI	1.81, 4.71	1.12, 2.53	2.28, 5.28	1.16, 2.84
		Median	2.60	1.50	4.00	1.70
		Q25, Q75	1.60, 5.20	1.10, 2.70	1.80, 5.40	1.10, 3.00
		Min, Max	1.0, 6.2	1.0, 3.0	1.0, 6.6	1.0, 3.4
	T <sub>0</sub> + 6 h	n	10	8	10	8
		Mean (SD)	3.54 (2.029)	2.10 (0.963)	3.86 (2.117)	1.98 (0.994)
		95% CI	2.08, 5.00	1.29, 2.91	2.34, 5.38	1.14, 2.81
		Median	3.20	2.00	4.10	1.60
		Q25, Q75	2.20, 5.20	1.20, 2.90	2.20, 6.00	1.20, 2.70
		Min, Max	1.0, 6.4	1.0, 3.6	1.0, 6.4	1.0, 3.8

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: The QSU-brief values at T<sub>0</sub> + 15/20 min refer to QSU-brief data at time T<sub>0</sub> + 15 min for THS 2.2 Menthol and mCC, and to QSU-brief data at time T<sub>0</sub> + 20 min for NRT gum.

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/000000106326/dev/tables/t\_qsu3.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.14.2 Descriptive Statistics of QSU-brief Questionnaire Factors and Total Score by Nicotine Level - PK Population**

Variable	Timepoint	Statistic	Group-2 PK			
			THS 2.2 Menthol		NRT gum	
			<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)	<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)
Factor 2 - Relief	T <sub>0</sub> + 9 h	n	10	8	10	8
		Mean (SD)	3.54 (2.001)	2.08 (1.166)	3.70 (2.010)	2.00 (0.908)
		95% CI	2.10, 4.98	1.10, 3.05	2.26, 5.14	1.24, 2.76
		Median	3.10	1.90	3.90	1.80
		Q25, Q75	2.40, 5.20	1.00, 3.00	2.20, 5.40	1.20, 2.80
		Min, Max	1.0, 7.0	1.0, 3.8	1.0, 6.4	1.0, 3.4
	T <sub>0</sub> + 12 h	n	10	8	10	8
		Mean (SD)	3.60 (1.996)	2.08 (1.228)	3.90 (2.049)	1.95 (1.047)
		95% CI	2.17, 5.03	1.04, 3.11	2.43, 5.37	1.07, 2.83
		Median	3.30	1.70	3.90	1.60
		Q25, Q75	2.00, 5.20	1.00, 3.00	2.60, 5.40	1.10, 2.70
		Min, Max	1.0, 6.6	1.0, 4.2	1.0, 6.6	1.0, 3.8

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: The QSU-brief values at T<sub>0</sub> + 15/20 min refer to QSU-brief data at time T<sub>0</sub> + 15 min for THS 2.2 Menthol and mCC, and to QSU-brief data at time T<sub>0</sub> + 20 min for NRT gum.

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/000000106326/dev/tables/t\_qsu3.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.14.2 Descriptive Statistics of QSU-brief Questionnaire Factors and Total Score by Nicotine Level - PK Population**

Variable	Timepoint	Statistic	Group-2 PK			
			THS 2.2 Menthol		NRT gum	
			<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)	<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)
Total score	15 min < T <sub>0</sub>	n	10	8	10	8
		Mean (SD)	4.42 (1.926)	3.44 (1.822)	4.84 (1.494)	3.55 (1.328)
		95% CI	3.04, 5.80	1.91, 4.97	3.77, 5.91	2.43, 4.67
		Median	4.70	3.65	4.70	3.35
		Q25, Q75	2.60, 6.20	1.70, 4.90	3.90, 5.90	2.40, 4.75
		Min, Max	1.4, 6.7	1.2, 5.8	1.9, 7.0	2.0, 5.4
	T <sub>0</sub> + 15/20 min	n	10	8	10	8
		Mean (SD)	3.24 (2.029)	2.58 (1.747)	4.42 (1.699)	2.81 (1.635)
		95% CI	1.78, 4.70	1.11, 4.04	3.20, 5.64	1.44, 4.18
		Median	2.60	1.80	4.65	2.00
		Q25, Q75	1.50, 5.00	1.20, 4.00	2.80, 5.80	1.65, 4.55
		Min, Max	1.1, 6.6	1.0, 5.6	1.4, 6.6	1.0, 5.1

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: The QSU-brief values at T<sub>0</sub> + 15/20 min refer to QSU-brief data at time T<sub>0</sub> + 15 min for THS 2.2 Menthol and mCC, and to QSU-brief data at time T<sub>0</sub> + 20 min for NRT gum.

Note: QSU-brief scores reported on a 7-point scale. Higher scores indicate greater intensity of urge.

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Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_qsu3.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.14.2 Descriptive Statistics of QSU-brief Questionnaire Factors and Total Score by Nicotine Level - PK Population**

Variable	Timepoint	Statistic	Group-2 PK			
			THS 2.2 Menthol		NRT gum	
			<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)	<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)
Total score	T <sub>0</sub> + 30 min	n	10	8	10	8
		Mean (SD)	3.14 (2.049)	2.55 (1.625)	4.28 (1.872)	2.75 (1.871)
		95% CI	1.67, 4.61	1.19, 3.91	2.94, 5.62	1.18, 4.32
		Median	2.05	1.80	4.45	1.80
		Q25, Q75	1.50, 4.80	1.25, 4.15	2.80, 5.70	1.25, 4.85
		Min, Max	1.0, 6.5	1.0, 5.0	1.1, 6.9	1.0, 5.2
	T <sub>0</sub> + 45 min	n	10	8	10	8
		Mean (SD)	3.52 (1.975)	2.39 (1.672)	3.89 (2.081)	2.64 (1.772)
		95% CI	2.10, 4.94	0.99, 3.79	2.40, 5.38	1.15, 4.12
		Median	3.15	1.30	4.30	1.65
		Q25, Q75	1.80, 5.00	1.25, 3.90	2.10, 5.60	1.30, 4.50
		Min, Max	1.1, 6.7	1.0, 5.2	1.0, 6.7	1.0, 5.2

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: The QSU-brief values at T<sub>0</sub> + 15/20 min refer to QSU-brief data at time T<sub>0</sub> + 15 min for THS 2.2 Menthol and mCC, and to QSU-brief data at time T<sub>0</sub> + 20 min for NRT gum.

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/000000106326/dev/tables/t\_qsu3.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.14.2 Descriptive Statistics of QSU-brief Questionnaire Factors and Total Score by Nicotine Level - PK Population**

Variable	Timepoint	Statistic	Group-2 PK			
			THS 2.2 Menthol		NRT gum	
			<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)	<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)
Total score	T <sub>0</sub> + 60 min	n	10	8	10	8
		Mean (SD)	3.24 (1.844)	2.48 (1.540)	4.11 (2.252)	2.74 (1.835)
		95% CI	1.92, 4.56	1.18, 3.77	2.49, 5.73	1.20, 4.28
		Median	3.00	1.80	4.45	1.85
		Q25, Q75	1.50, 4.70	1.20, 3.95	2.30, 6.50	1.25, 4.75
		Min, Max	1.2, 6.7	1.0, 4.9	1.0, 6.7	1.0, 5.2
	T <sub>0</sub> + 2 h	n	10	8	10	8
		Mean (SD)	3.88 (1.993)	2.73 (1.815)	4.32 (1.920)	2.85 (1.682)
		95% CI	2.45, 5.31	1.20, 4.25	2.94, 5.70	1.44, 4.26
		Median	3.90	1.80	4.85	2.20
		Q25, Q75	1.70, 6.00	1.30, 4.65	2.80, 5.40	1.55, 4.55
		Min, Max	1.5, 6.7	1.0, 5.3	1.0, 6.7	1.0, 5.2

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: The QSU-brief values at T<sub>0</sub> + 15/20 min refer to QSU-brief data at time T<sub>0</sub> + 15 min for THS 2.2 Menthol and mCC, and to QSU-brief data at time T<sub>0</sub> + 20 min for NRT gum.

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/000000106326/dev/tables/t\_qsu3.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.14.2 Descriptive Statistics of QSU-brief Questionnaire Factors and Total Score by Nicotine Level - PK Population**

Variable	Timepoint	Statistic	Group-2 PK			
			THS 2.2 Menthol		NRT gum	
			<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)	<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)
Total score	T <sub>0</sub> + 4 h	n	10	8	10	8
		Mean (SD)	3.88 (1.989)	3.01 (1.610)	4.29 (2.114)	3.01 (1.715)
		95% CI	2.45, 5.31	1.66, 4.36	2.77, 5.81	1.57, 4.45
		Median	3.75	2.45	5.00	2.00
		Q25, Q75	2.20, 5.90	1.70, 4.80	2.20, 5.50	1.70, 5.00
		Min, Max	1.0, 6.6	1.2, 5.0	1.0, 6.8	1.5, 5.2
	T <sub>0</sub> + 6 h	n	10	8	10	8
		Mean (SD)	4.30 (1.834)	3.29 (1.688)	4.38 (2.133)	3.11 (1.634)
		95% CI	2.98, 5.62	1.87, 4.70	2.85, 5.91	1.74, 4.48
		Median	4.35	3.20	5.00	2.35
		Q25, Q75	2.70, 6.00	1.85, 4.95	2.70, 6.40	1.85, 4.85
		Min, Max	1.3, 6.7	1.0, 5.3	1.0, 6.7	1.4, 5.4

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: The QSU-brief values at T<sub>0</sub> + 15/20 min refer to QSU-brief data at time T<sub>0</sub> + 15 min for THS 2.2 Menthol and mCC, and to QSU-brief data at time T<sub>0</sub> + 20 min for NRT gum.

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/000000106326/dev/tables/t\_qsu3.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.14.2 Descriptive Statistics of QSU-brief Questionnaire Factors and Total Score by Nicotine Level - PK Population**

Variable	Timepoint	Statistic	Group-2 PK			
			THS 2.2 Menthol		NRT gum	
			<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)	<= 0.6 mg (N=10)	> 0.6 - 1.0 mg (N=8)
Total score	T <sub>0</sub> + 9 h	n	10	8	10	8
		Mean (SD)	4.18 (1.892)	3.11 (1.762)	4.25 (2.044)	3.18 (1.547)
		95% CI	2.82, 5.54	1.63, 4.59	2.78, 5.72	1.88, 4.47
		Median	4.10	2.55	4.75	2.35
		Q25, Q75	3.00, 6.00	1.70, 5.00	2.60, 5.90	2.15, 4.90
		Min, Max	1.3, 7.0	1.0, 5.4	1.0, 6.6	1.4, 5.2
	T <sub>0</sub> + 12 h	n	10	8	10	8
		Mean (SD)	4.24 (1.928)	3.23 (1.715)	4.47 (2.005)	3.09 (1.678)
		95% CI	2.86, 5.62	1.79, 4.66	3.03, 5.91	1.68, 4.49
		Median	4.20	2.65	5.00	2.45
		Q25, Q75	2.70, 6.00	2.00, 4.95	3.70, 5.80	1.70, 4.85
		Min, Max	1.1, 6.7	1.0, 5.6	1.1, 6.7	1.3, 5.4

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

Note: T<sub>0</sub> = Time of first product use at single use day.

Note: The QSU-brief values at T<sub>0</sub> + 15/20 min refer to QSU-brief data at time T<sub>0</sub> + 15 min for THS 2.2 Menthol and mCC, and to QSU-brief data at time T<sub>0</sub> + 20 min for NRT gum.

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/000000106326/dev/tables/t\_qsu3.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.15 Analysis of MCEQ Subscales - PK Population**

Subscales	Statistic	Group-1 PK		
		THS 2.2 Menthol (N=43)	mCC (N=43)	THS 2.2 Menthol - mCC
Aversion Subscale	n	43	43	
	Mean (SE)	2.37	2.46	-0.10 (0.248)
	95% CI	1.99, 2.74	2.09, 2.84	-0.60, 0.41
Craving Reduction Subscale	n	43	43	
	Mean (SE)	3.30	3.44	-0.14 (0.196)
	95% CI	2.90, 3.71	3.04, 3.85	-0.54, 0.26
Enjoyment of Respiratory Tract Sensation Subscale	n	43	43	
	Mean (SE)	3.34	3.94	-0.59 (0.223)
	95% CI	2.88, 3.81	3.47, 4.40	-1.05, -0.14

Note: mCC = menthol conventional cigarettes; THS = Tobacco Heating System.

Note: Higher scores indicate greater intensity on that scale.

Note: Means and 95% CI are the adjusted least squares means and confidence intervals from an ANOVA model.

Appendix 15.4.4.15 and 15.3.6.12

Path: /cvn/projects/prj/development/000000106326/dev/tables/tl\_anlmceq.sas

Program Run: 28OCT14 cvn\_ahedge Program Status: FINAL

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**Table 15.2.4.15 Analysis of MCEQ Subscales - PK Population**

Subscales	Statistic	Group-1 PK		
		THS 2.2 Menthol (N=43)	mCC (N=43)	THS 2.2 Menthol - mCC
Psychological Reward Subscale	n	43	43	
	Mean (SE)	2.77	2.96	-0.19 (0.105)
	95% CI	2.49, 3.06	2.68, 3.24	-0.40, 0.03
Smoking Satisfaction Subscale	n	43	43	
	Mean (SE)	3.19	4.31	-1.12 (0.198)
	95% CI	2.77, 3.60	3.90, 4.73	-1.53, -0.72

Note: mCC = menthol conventional cigarettes; THS = Tobacco Heating System.

Note: Higher scores indicate greater intensity on that scale.

Note: Means and 95% CI are the adjusted least squares means and confidence intervals from an ANOVA model.

Appendix 15.4.4.15 and 15.3.6.12

Path: /cvn/projects/prj/development/000000106326/dev/tables/tl\_anlmceq.sas

Program Run: 28OCT14 cvn\_ahedge Program Status: FINAL

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**Table 15.2.4.16 Analysis of MCEQ Subscales by Bootstrapping Techniques - PK Population**

Subscales	Statistic	Group-1 PK		
		THS 2.2 Menthol (N=43)	mCC (N=43)	THS 2.2 Menthol - mCC
Aversion Subscale	n	43	43	
	Mean (SE)	2.37	2.47	-0.10 (0.236)
	95% CI	2.02, 2.77	2.15, 2.81	-0.58, 0.38
	Median	2.36	2.47	-0.10
Craving Reduction Subscale	n	43	43	
	Mean (SE)	3.30	3.44	-0.14 (0.190)
	95% CI	2.94, 3.66	3.06, 3.85	-0.54, 0.24
	Median	3.30	3.44	-0.14
Enjoyment of Respiratory Tract Sensation Subscale	n	43	43	
	Mean (SE)	3.34	3.94	-0.60 (0.217)
	95% CI	2.90, 3.77	3.49, 4.39	-1.03, -0.19
	Median	3.35	3.94	-0.59

Note: mCC = menthol conventional cigarettes; THS = Tobacco Heating System.

Note: Higher scores indicate greater intensity on that scale

Note: Point and 95% interval estimates by means of the percentile bootstrap technique of the adjusted least squares means from an ANOVA model.

Appendix 15.4.4.16 and 15.3.6.12

Path: /cvn/projects/prj/development/000000106326/dev/tables/tl\_banlmceq.sas

Program Run: 28OCT14 cvn\_ahedge Program Status: FINAL

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**Table 15.2.4.16 Analysis of MCEQ Subscales by Bootstrapping Techniques - PK Population**

Subscales	Statistic	Group-1 PK		
		THS 2.2 Menthol (N=43)	mCC (N=43)	THS 2.2 Menthol - mCC
Psychological Reward Subscale	n	43	43	
	Mean (SE)	2.77	2.96	-0.19 (0.103)
	95% CI	2.51, 3.06	2.70, 3.24	-0.39, 0.03
	Median	2.77	2.96	-0.19
Smoking Satisfaction Subscale	n	43	43	
	Mean (SE)	3.19	4.31	-1.12 (0.193)
	95% CI	2.81, 3.56	3.89, 4.74	-1.52, -0.76
	Median	3.19	4.31	-1.12

Note: mCC = menthol conventional cigarettes; THS = Tobacco Heating System.

Note: Higher scores indicate greater intensity on that scale

Note: Point and 95% interval estimates by means of the percentile bootstrap technique of the adjusted least squares means from an ANOVA model.

Appendix 15.4.4.16 and 15.3.6.12

Path: /cvn/projects/prj/development/000000106326/dev/tables/tl\_banlmceq.sas

Program Run: 28OCT14 cvn\_ahedge Program Status: FINAL

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**Table 15.2.4.17 Descriptive Statistics of MCEQ Subscales - PK Population**

Subscale	Statistic	Group-1 PK	
		THS 2.2 Menthol (N=43)	mCC (N=43)
Aversion subscale	n	43	43
	Mean (SD)	2.36 (1.302)	2.47 (1.099)
	95% CI	1.95, 2.77	2.12, 2.81
	Median	2.00	2.50
	Q25, Q75	1.50, 3.00	1.50, 3.00
	Min, Max	1.0, 7.0	1.0, 7.0
Craving reduction subscale	n	43	43
	Mean (SD)	3.30 (1.226)	3.44 (1.333)
	95% CI	2.92, 3.68	3.03, 3.86
	Median	3.00	3.00
	Q25, Q75	3.00, 4.00	3.00, 4.00
	Min, Max	1.0, 7.0	1.0, 7.0

Note: mCC = menthol conventional cigarettes; THS = Tobacco Heating System.

Appendix 15.3.6.12

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_mceq.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.17 Descriptive Statistics of MCEQ Subscales - PK Population**

Subscale	Statistic	Group-1 PK	
		THS 2.2 Menthol (N=43)	mCC (N=43)
Enjoyment of respiratory tract sensation subscale	n	43	43
	Mean (SD)	3.35 (1.446)	3.93 (1.534)
	95% CI	2.90, 3.80	3.45, 4.41
	Median	3.00	4.00
	Q25, Q75	2.00, 4.00	3.00, 5.00
	Min, Max	1.0, 7.0	1.0, 7.0
Psychological reward subscale	n	43	43
	Mean (SD)	2.77 (0.879)	2.95 (0.934)
	95% CI	2.50, 3.05	2.66, 3.25
	Median	2.60	2.80
	Q25, Q75	2.20, 3.20	2.40, 3.60
	Min, Max	1.6, 5.4	1.2, 4.8

Note: mCC = menthol conventional cigarettes; THS = Tobacco Heating System.

Appendix 15.3.6.12

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_mceq.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.17 Descriptive Statistics of MCEQ Subscales - PK Population**

Subscale	Statistic	Group-1 PK	
		THS 2.2 Menthol (N=43)	mCC (N=43)
Smoking satisfaction subscale	n	43	43
	Mean (SD)	3.20 (1.290)	4.31 (1.393)
	95% CI	2.79, 3.60	3.87, 4.74
	Median	3.00	4.00
	Q25, Q75	2.30, 4.00	3.70, 5.30
	Min, Max	1.0, 6.3	1.3, 7.0

Note: mCC = menthol conventional cigarettes; THS = Tobacco Heating System.

Appendix 15.3.6.12

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_mceq.sas

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Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

**Table 15.2.4.17.1 Descriptive Statistics of MCEQ Subscales by Sex - PK Population**

Subscale	Statistic	Group-1 PK			
		THS 2.2 Menthol		mCC	
		Males (N=23)	Females (N=20)	Males (N=23)	Females (N=20)
Aversion subscale	n	23	20	23	20
	Mean (SD)	2.17 (1.164)	2.58 (1.445)	2.76 (1.242)	2.13 (0.810)
	95% CI	1.67, 2.68	1.89, 3.26	2.22, 3.30	1.74, 2.51
	Median	2.00	2.25	2.50	1.75
	Q25, Q75	1.50, 3.00	1.50, 3.00	2.00, 3.50	1.50, 3.00
	Min, Max	1.0, 6.0	1.0, 7.0	1.0, 7.0	1.0, 4.0
Craving reduction subscale	n	23	20	23	20
	Mean (SD)	3.39 (1.470)	3.20 (0.895)	3.83 (1.403)	3.00 (1.124)
	95% CI	2.75, 4.03	2.78, 3.62	3.21, 4.44	2.47, 3.53
	Median	3.00	3.00	4.00	3.00
	Q25, Q75	3.00, 4.00	3.00, 4.00	3.00, 4.00	2.00, 3.50
	Min, Max	1.0, 7.0	1.0, 5.0	1.0, 7.0	2.0, 6.0

Note: mCC = menthol conventional cigarettes; THS = Tobacco Heating System.

Appendix 15.3.6.12

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_mceq2.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.17.1 Descriptive Statistics of MCEQ Subscales by Sex - PK Population**

Subscale	Statistic	Group-1 PK			
		THS 2.2 Menthol		mCC	
		Males (N=23)	Females (N=20)	Males (N=23)	Females (N=20)
Enjoyment of respiratory tract sensation subscale	n	23	20	23	20
	Mean (SD)	3.65 (1.496)	3.00 (1.338)	3.91 (1.475)	3.95 (1.638)
	95% CI	3.00, 4.30	2.37, 3.63	3.27, 4.56	3.18, 4.72
	Median	4.00	3.00	4.00	4.00
	Q25, Q75	2.00, 4.00	2.00, 4.00	3.00, 5.00	2.00, 5.00
	Min, Max	1.0, 7.0	1.0, 6.0	1.0, 7.0	2.0, 7.0
Psychological reward subscale	n	23	20	23	20
	Mean (SD)	3.01 (1.054)	2.50 (0.526)	2.93 (0.989)	2.98 (0.892)
	95% CI	2.55, 3.47	2.25, 2.75	2.50, 3.36	2.56, 3.40
	Median	2.80	2.40	2.80	2.70
	Q25, Q75	2.20, 3.40	2.20, 2.90	2.00, 3.60	2.40, 3.60
	Min, Max	1.6, 5.4	1.8, 3.6	1.2, 4.8	1.6, 4.6

Note: mCC = menthol conventional cigarettes; THS = Tobacco Heating System.

Appendix 15.3.6.12

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_mceq2.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.17.1 Descriptive Statistics of MCEQ Subscales by Sex - PK Population**

Subscale	Statistic	Group-1 PK			
		THS 2.2 Menthol		mCC	
		Males (N=23)	Females (N=20)	Males (N=23)	Females (N=20)
Smoking satisfaction subscale	n	23	20	23	20
	Mean (SD)	3.63 (1.286)	2.69 (1.123)	4.24 (1.326)	4.38 (1.498)
	95% CI	3.07, 4.20	2.16, 3.22	3.67, 4.82	3.67, 5.09
	Median	3.30	2.30	4.00	4.30
	Q25, Q75	2.70, 4.30	2.00, 3.50	3.70, 5.00	3.35, 5.70
	Min, Max	2.0, 6.3	1.0, 5.0	1.3, 6.7	1.7, 7.0

Note: mCC = menthol conventional cigarettes; THS = Tobacco Heating System.

Appendix 15.3.6.12

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_mceq2.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.17.2 Descriptive Statistics of MCEQ Subscales by Nicotine Level - PK Population**

Subscale	Statistic	Group-1 PK			
		THS 2.2 Menthol		mCC	
		<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)	<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)
Aversion subscale	n	25	18	25	18
	Mean (SD)	2.64 (1.518)	1.97 (0.813)	2.16 (0.826)	2.89 (1.302)
	95% CI	2.01, 3.27	1.56, 2.38	1.81, 2.51	2.24, 3.54
	Median	2.50	2.00	2.00	2.75
	Q25, Q75	1.50, 3.00	1.50, 2.50	1.50, 2.50	2.50, 3.00
	Min, Max	1.0, 7.0	1.0, 3.5	1.0, 3.5	1.0, 7.0
Craving reduction subscale	n	25	18	25	18
	Mean (SD)	3.32 (1.216)	3.28 (1.275)	3.36 (1.381)	3.56 (1.294)
	95% CI	2.81, 3.83	2.64, 3.92	2.79, 3.93	2.91, 4.20
	Median	3.00	3.00	3.00	3.50
	Q25, Q75	3.00, 4.00	3.00, 4.00	2.00, 4.00	3.00, 4.00
	Min, Max	1.0, 6.0	1.0, 7.0	2.0, 7.0	1.0, 6.0

Note: mCC = menthol conventional cigarettes; THS = Tobacco Heating System.

Appendix 15.3.6.12

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_mceq3.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.17.2 Descriptive Statistics of MCEQ Subscales by Nicotine Level - PK Population**

Subscale	Statistic	Group-1 PK			
		THS 2.2 Menthol		mCC	
		<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)	<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)
Enjoyment of respiratory tract sensation subscale	n	25	18	25	18
	Mean (SD)	3.56 (1.387)	3.06 (1.514)	4.04 (1.696)	3.78 (1.309)
	95% CI	2.98, 4.14	2.30, 3.81	3.34, 4.74	3.12, 4.43
	Median	3.00	3.00	4.00	4.00
	Q25, Q75	3.00, 4.00	2.00, 4.00	2.00, 5.00	3.00, 4.00
	Min, Max	1.0, 6.0	1.0, 7.0	1.0, 7.0	2.0, 7.0
Psychological reward subscale	n	25	18	25	18
	Mean (SD)	2.79 (0.797)	2.74 (1.006)	2.92 (0.928)	3.00 (0.968)
	95% CI	2.46, 3.13	2.24, 3.25	2.53, 3.31	2.51, 3.49
	Median	2.60	2.40	2.60	2.80
	Q25, Q75	2.20, 3.40	2.00, 3.00	2.40, 3.60	2.40, 3.60
	Min, Max	1.8, 5.2	1.6, 5.4	1.4, 4.8	1.2, 4.6

Note: mCC = menthol conventional cigarettes; THS = Tobacco Heating System.

Appendix 15.3.6.12

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_mceq3.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.17.2 Descriptive Statistics of MCEQ Subscales by Nicotine Level - PK Population**

Subscale	Statistic	Group-1 PK			
		THS 2.2 Menthol		mCC	
		<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)	<= 0.6 mg (N=25)	> 0.6 - 1.0 mg (N=18)
Smoking satisfaction subscale	n	25	18	25	18
	Mean (SD)	3.08 (1.366)	3.35 (1.198)	4.48 (1.371)	4.07 (1.428)
	95% CI	2.52, 3.65	2.75, 3.95	3.91, 5.05	3.36, 4.79
	Median	2.70	3.15	4.70	3.70
	Q25, Q75	2.00, 4.00	2.70, 3.70	3.70, 5.70	3.70, 4.70
	Min, Max	1.0, 6.3	2.0, 6.0	1.7, 6.7	1.3, 7.0

Note: mCC = menthol conventional cigarettes; THS = Tobacco Heating System.

Appendix 15.3.6.12

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_mceq3.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.4.18 Descriptive Statistics of Sex, Age and Nicotine Level - PK Population**

Variable	Statistic	Group-1 PK		Group-2 PK	
		THS 2.2 Menthol - mCC (N=21)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)
Sex					
Male	n (%)	11 (52.4%)	12 (54.5%)	5 (55.6%)	4 (44.4%)
Female	n (%)	10 (47.6%)	10 (45.5%)	4 (44.4%)	5 (55.6%)
Nicotine level					
<= 0.6 mg	n (%)	13 (61.9%)	12 (54.5%)	5 (55.6%)	5 (55.6%)
> 0.6 - 1.0 mg	n (%)	8 (38.1%)	10 (45.5%)	4 (44.4%)	4 (44.4%)
Age (years)					
	n	21	22	9	9
	Mean (SD)	35.5 (12.04)	31.4 (7.38)	29.3 (7.50)	32.1 (8.29)
	Median	31.0	31.0	28.0	29.0
	Min, Max	23, 61	23, 44	23, 44	23, 44

Note: mCC = menthol conventional cigarettes; NRT gum= Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Appendix 15.3.1.6

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_sanica.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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## **15.2.5 Compliance**

**Table 15.2.5.1 Descriptive Statistics of Compliance Based on CO Breath Test - All Randomized Subjects**

Visit/Study Day	Statistic	Sequence				Overall Randomized (N=62)
		THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)	
Single-use/Day 1						
Compliant	n (%)	NA	NA	NA	9 (100 %)	NA
Wash-out/Day 2						
Compliant	n (%)	21 ( 95.5%)	22 (100 %)	9 (100 %)	9 (100 %)	61 ( 98.4%)
Missing	n (%)	1 ( 4.5%)	0	0	0	1 ( 1.6%)
Single-use/Day 3						
Compliant	n (%)	NA	NA	9 (100 %)	NA	NA
Missing	n (%)	NA	NA	0	NA	NA

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

Note: Compliance assessed on Days 1 and 3 for subjects randomized to NRT gum and for all subjects on Wash-out Day 2.

Note: NA = Not assessed.

Appendix 15.3.3.5

Path: /cvn/projects/prj/development/000000106326/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**

	Sequence															Overall Safety (N=73)		
	THS 2.2 Menthol - mCC (N=22)			mCC - THS 2.2 Menthol (N=22)			THS 2.2 Menthol - NRT gum (N=9)			NRT gum - THS 2.2 Menthol (N=9)			Exposed Not Randomized (N=11)					
	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events
Adverse events (AE)	1	( 4.5%)	1	2	( 9.1%)	2	0			1	( 11.1%)	1	0			4	( 5.5%)	4
AE serious																		
Yes	0			0						0						0		
No	1	( 4.5%)	1	2	( 9.1%)	2				1	( 11.1%)	1				4	( 5.5%)	4
AE related to IP																		
Not related	1	( 4.5%)	1	2	( 9.1%)	2				1	( 11.1%)	1				4	( 5.5%)	4
Related	0			0						0						0		

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System

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

Note: IP = Investigational Product (THS 2.2 Menthol / mCC).

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/000000106326/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**

	Sequence															Overall Safety (N=73)		
	THS 2.2 Menthol - mCC (N=22)			mCC - THS 2.2 Menthol (N=22)			THS 2.2 Menthol - NRT gum (N=9)			NRT gum - THS 2.2 Menthol (N=9)			Exposed Not Randomized (N=11)					
	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events			
AE related to NRT gum																		
Not related	1	( 4.5%)	1	2	( 9.1%)	2				1	( 11.1%)	1				4	( 5.5%)	4
Related	0			0						0						0		
AE related to study procedure																		
Not related	1	( 4.5%)	1	2	( 9.1%)	2				1	( 11.1%)	1				4	( 5.5%)	4
Related	0			0						0						0		
AE severity																		
Mild	1	( 4.5%)	1	2	( 9.1%)	2				1	( 11.1%)	1				4	( 5.5%)	4
Moderate	0			0						0						0		
Severe	0			0						0						0		

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System

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

Note: IP = Investigational Product (THS 2.2 Menthol / mCC).

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/000000106326/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**

	Sequence															Overall Safety (N=73)	
	THS 2.2 Menthol - mCC (N=22)			mCC - THS 2.2 Menthol (N=22)			THS 2.2 Menthol - NRT gum (N=9)			NRT gum - THS 2.2 Menthol (N=9)			Exposed Not Randomized (N=11)				
	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events		
Action taken due to AE																	
Discontinuation from study	0			0						0						0	
Related to product use	0			0						0						0	
Treatment given																	
Yes	0			0						0						0	
No	1	( 4.5%)	1	2	( 9.1%)	2				1	( 11.1%)	1				4	( 5.5%) 4
Other action taken	0			0						0						0	

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System

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

Note: IP = Investigational Product (THS 2.2 Menthol / mCC).

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/000000106326/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	Sequence															Overall Safety (N=73) n (%) Events		
	THS 2.2 Menthol - mCC (N=22)			mCC - THS 2.2 Menthol (N=22)			THS 2.2 Menthol - NRT gum (N=9)			NRT gum - THS 2.2 Menthol (N=9)			Exposed Not Randomized (N=11)					
	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events			
Any adverse events	1	( 4.5%)	1	2	( 9.1%)	2	0			1	( 11.1%)	1	0			4	( 5.5%)	4
Investigations	1	( 4.5%)	1	2	( 9.1%)	2				1	( 11.1%)	1				4	( 5.5%)	4
Bilirubin conjugated increased	0			1	( 4.5%)	1				0						1	( 1.4%)	1
Blood bilirubin increased	0			1	( 4.5%)	1				0						1	( 1.4%)	1
Haemoglobin decreased	0			0						1	( 11.1%)	1				1	( 1.4%)	1
Lymphocyte count increased	1	( 4.5%)	1	0						0						1	( 1.4%)	1

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System

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

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

Appendix 15.3.6.1.1

Path: /cvm/projects/prj/development/000000106326/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	Sequence																		Overall Safety (N=73)																		
	THS 2.2 Menthol - mCC (N=22)						mCC - THS 2.2 Menthol (N=22)						THS 2.2 Menthol - NRT gum (N=9)							NRT gum - THS 2.2 Menthol (N=9)						Exposed Not Randomized (N=11)											
	n			(%)			Events			n			(%)			Events				n			(%)			Events			n			(%)			Events		
	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events		n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events						
Any adverse events	1	( 4.5%)	1	2	( 9.1%)	2	0					1	( 11.1%)	1	0									4	( 5.5%)	4											
Investigations	1	( 4.5%)	1	2	( 9.1%)	2							1	( 11.1%)	1									4	( 5.5%)	4											

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System

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

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/000000106326/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	Sequence															Overall Safety (N=73)		
	THS 2.2 Menthol - mCC (N=22)			mCC - THS 2.2 Menthol (N=22)			THS 2.2 Menthol - NRT gum (N=9)			NRT gum - THS 2.2 Menthol (N=9)			Exposed Not Randomized (N=11)					
	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events
Any adverse events	1	( 4.5%)	1	2	( 9.1%)	2	0			1	( 11.1%)	1	0			4	( 5.5%)	4
Bilirubin conjugated increased	0			1	( 4.5%)	1				0						1	( 1.4%)	1
Blood bilirubin increased	0			1	( 4.5%)	1				0						1	( 1.4%)	1
Haemoglobin decreased	0			0						1	( 11.1%)	1				1	( 1.4%)	1
Lymphocyte count increased	1	( 4.5%)	1	0						0						1	( 1.4%)	1

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System

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

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/000000106326/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 and Preferred Term and Relationship to Study Product Exposure for Investigational Product (THS 2.2 Menthol or mCC) and Reference Point Product (NRT gum) - Safety Population**

	Sequence															Overall Safety (N=73)	
	THS 2.2 Menthol - mCC (N=22)			mCC - THS 2.2 Menthol (N=22)			THS 2.2 Menthol - NRT gum (N=9)			NRT gum - THS 2.2 Menthol (N=9)			Exposed Not Randomized (N=11)				
	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events		
Adverse events (AE)	1	( 4.5%)	1	2	( 9.1%)	2	0			1	( 11.1%)	1	0		4	( 5.5%)	4
Related to IP	0			0						0					0		
Related to NRT gum	0			0						0					0		
Investigations	1	( 4.5%)	1	2	( 9.1%)	2				1	( 11.1%)	1			4	( 5.5%)	4
(Bilirubin	0			1	( 4.5%)	1				0					1	( 1.4%)	1
conjugated increased)																	
Related to IP				0											0		
Related to NRT gum				0											0		

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System

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

Note: IP = Investigational Product (THS 2.2 Menthol / mCC).

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/000000106326/dev/tables/t\_adv6.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 and Preferred Term and Relationship to Study Product Exposure for Investigational Product (THS 2.2 Menthol or mCC) and Reference Point Product (NRT gum) - Safety Population**

	Sequence															Overall Safety (N=73)
	THS 2.2 Menthol - mCC (N=22)			mCC - THS 2.2 Menthol (N=22)			THS 2.2 Menthol - NRT gum (N=9)			NRT gum - THS 2.2 Menthol (N=9)			Exposed Not Randomized (N=11)			
	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events	
(Blood bilirubin increased)	0			1 ( 4.5%)	1					0					1 ( 1.4%)	1
Related to IP				0											0	
Related to NRT gum				0											0	
(Haemoglobin decreased)	0			0						1 ( 11.1%)	1				1 ( 1.4%)	1
Related to IP										0					0	
Related to NRT gum										0					0	
(Lymphocyte count increased)	1 ( 4.5%)	1		0						0					1 ( 1.4%)	1
Related to IP	0														0	
Related to NRT gum	0														0	

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System

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

Note: IP = Investigational Product (THS 2.2 Menthol / mCC).

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/000000106326/dev/tables/t\_adv6.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.6.4 Summary of Adverse Events Leading to Study Discontinuation by System Organ Class and Preferred Term - Safety Population**

System Organ Class	Sequence															Overall Safety (N=73)
	THS 2.2 Menthol - mCC (N=22)			mCC - THS 2.2 Menthol (N=22)			THS 2.2 Menthol - NRT gum (N=9)			NRT gum - THS 2.2 Menthol (N=9)			Exposed Not Randomized (N=11)			
	Preferred Term	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	

No adverse events leading to study discontinuation were reported

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System

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

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/000000106326/dev/tables/t\_adv5.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.6.5 Summary of Adverse Events by System Organ Class, Preferred Term and Severity - Safety Population**

System Organ Class Preferred Term	Sequence															Overall Safety (N=73)  Events		
	THS 2.2 Menthol - mCC (N=22)			mCC - THS 2.2 Menthol (N=22)			THS 2.2 Menthol - NRT gum (N=9)			NRT gum - THS 2.2 Menthol (N=9)			Exposed Not Randomized (N=11)					
	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events			
Adverse events	1	( 4.5%)	1	2	( 9.1%)	2	0				1	( 11.1%)	1	0		4	( 5.5%)	4
Mild	1	( 4.5%)	1	2	( 9.1%)	2					1	( 11.1%)	1			4	( 5.5%)	4
Moderate	0			0							0					0		
Severe	0			0							0					0		
Investigations	1	( 4.5%)	1	2	( 9.1%)	2					1	( 11.1%)	1			4	( 5.5%)	4
Bilirubin conjugated increased	0			1	( 4.5%)	1					0					1	( 1.4%)	1
Mild				1	( 4.5%)	1										1	( 1.4%)	1
Moderate				0												0		
Severe				0												0		

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

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/000000106326/dev/tables/t\_adv7.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.6.5 Summary of Adverse Events by System Organ Class, Preferred Term and Severity - Safety Population**

System Organ Class	Sequence															Overall Safety (N=73)
	THS 2.2 Menthol - mCC (N=22)			mCC - THS 2.2 Menthol (N=22)			THS 2.2 Menthol - NRT gum (N=9)			NRT gum - THS 2.2 Menthol (N=9)			Exposed Not Randomized (N=11)			
	Preferred Term	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	
Blood bilirubin increased	0			1 ( 4.5%)	1					0					1 ( 1.4%)	1
Mild				1 ( 4.5%)	1										1 ( 1.4%)	1
Moderate				0											0	
Severe				0											0	
Haemoglobin decreased	0			0						1 ( 11.1%)	1				1 ( 1.4%)	1
Mild										1 ( 11.1%)	1				1 ( 1.4%)	1
Moderate										0					0	
Severe										0					0	

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

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/000000106326/dev/tables/t\_adv7.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.6.5 Summary of Adverse Events by System Organ Class, Preferred Term and Severity - Safety Population**

System Organ Class	Sequence															Overall Safety (N=73)	
	THS 2.2 Menthol - mCC (N=22)			mCC - THS 2.2 Menthol (N=22)			THS 2.2 Menthol - NRT gum (N=9)			NRT gum - THS 2.2 Menthol (N=9)			Exposed Not Randomized (N=11)				
	Preferred Term	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)		Events
Lymphocyte count increased	1	( 4.5%)	1	0						0					1	( 1.4%)	1
Mild	1	( 4.5%)	1												1	( 1.4%)	1
Moderate	0														0		
Severe	0														0		

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

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/000000106326/dev/tables/t\_adv7.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	Sequence																Overall Safety (N=73)
	THS 2.2 Menthol - mCC (N=22)			mCC - THS 2.2 Menthol (N=22)			THS 2.2 Menthol - NRT gum (N=9)			NRT gum - THS 2.2 Menthol (N=9)			Exposed Not Randomized (N=11)				
	Preferred Term	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events	

No adverse events related to study procedure were reported

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System

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

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/000000106326/dev/tables/t\_adv8.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.6.7 Summary of THS 2.2 Menthol Device Events and Malfunctions - Safety Population**

Sequence																	
THS 2.2 Menthol - mCC (N=22)			mCC - THS 2.2 Menthol (N=22)			THS 2.2 Menthol - NRT gum (N=9)			NRT gum - THS 2.2 Menthol (N=9)			Enrolled Not Randomized (N=11)			Overall Safety (N=73)		
n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events

No device events were recorded

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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/000000106326/dev/tables/t\_device.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.6.8.1 Summary of Prior Medication by Anatomical Therapeutic Classes (ATC) 1 and 2 - Safety Population**

ATC1 ATC2	Sequence															Overall Safety (N=73)
	THS 2.2 Menthol - mCC (N=22)			mCC - THS 2.2 Menthol (N=22)			THS 2.2 Menthol - NRT gum (N=9)			NRT gum - THS 2.2 Menthol (N=9)			Exposed Not Randomized (N=11)			
	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events	

No prior medication reported

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

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/000000106326/dev/tables/t\_priatc.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.6.8.2 Summary of Prior Medication by Preferred Drug Name - Safety Population**

Sequence													
THS 2.2 Menthol - mCC (N=22)		mCC - THS 2.2 Menthol (N=22)		THS 2.2 Menthol - NRT gum (N=9)		NRT gum - THS 2.2 (N=9)		Exposed Not Randomized (N=11)		Overall Safety (N=73)			
Preferred Term	n (%)	Events	n (%)	Events	n (%)	Events	n (%)	Events	n (%)	Events	n (%)	Events	n (%)

No prior medication reported

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

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/000000106326/dev/tables/t\_primed.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.6.9.1 Summary of Concomitant Medication by Anatomical Therapeutic Classes (ATC) 1 and 2 - Safety Population**

ATC1 ATC2		Sequence															Overall Safety (N=73)	
		THS 2.2 Menthol - mCC (N=22)			mCC - THS 2.2 Menthol (N=22)			THS 2.2 Menthol - NRT gum (N=9)			NRT gum - THS 2.2 Menthol (N=9)			Exposed Not Randomized (N=11)				
		n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events		

No concomitant medication reported

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

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/000000106326/dev/tables/t\_conatc.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.6.9.2 Summary of Concomitant Medication by Preferred Drug Name - Safety Population**

Sequence													
THS 2.2 Menthol - mCC (N=22)		mCC - THS 2.2 Menthol (N=22)		THS 2.2 Menthol - NRT gum (N=9)		NRT gum - THS 2.2 (N=9)		Exposed Not Randomized (N=11)		Overall Safety (N=73)			
Preferred Term	n (%)	Events	n (%)	Events	n (%)	Events	n (%)	Events	n (%)	Events	n (%)	Events	n (%)

No concomitant medication reported

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; THS = Tobacco Heating System.

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

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/000000106326/dev/tables/t\_conmed.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.6.10 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Alanine aminotransferase (IU/L) 5-45	Baseline	n	22	22	9	9	11	73
		Mean (SD)	15.1 (6.00)	16.8 (7.14)	18.1 (7.58)	14.2 (4.92)	16.7 (6.82)	16.1 (6.51)
		Median	14.0	16.0	16.0	13.0	14.0	14.0
		Min, Max	8, 36	6, 39	8, 29	9, 22	9, 29	6, 39
		Low value - n (%)	0	0	0	0	0	0
		Normal value - n (%)	22 (100 %)	22 (100 %)	9 (100 %)	9 (100 %)	11 (100 %)	73 (100 %)
		High value - n (%)	0	0	0	0	0	0
		Abnormal clinically significant - n (%)	0	0	0	0	0	0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Appendix 15.3.6.4

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_labchem.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.6.10 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Alanine aminotransferase (IU/L) 5-45	Day 4/ Discharge	n	22	22	9	9	11	73
		Mean (SD)	17.8 (7.29)	22.0 (10.71)	18.4 (5.44)	16.4 (7.01)	16.8 (5.88)	18.8 (8.21)
		Median	15.5	20.0	18.0	14.0	15.0	17.0
		Min, Max	8, 33	10, 50	11, 28	10, 32	9, 26	8, 50
		Low value - n (%)	0	0	0	0	0	0
		Normal value - n (%)	22 (100 %)	21 ( 95.5%)	9 (100 %)	9 (100 %)	11 (100 %)	72 ( 98.6%)
		High value - n (%)	0	1 ( 4.5%)	0	0	0	1 ( 1.4%)
		Abnormal clinically significant - n (%)	0	0	0	0	0	0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Appendix 15.3.6.4

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_labchem.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.6.10 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Alanine aminotransferase (IU/L) 5-45	Change from n Baseline		22	22	9	9	11	73
		Mean (SD)	2.7 (4.65)	5.3 (5.17)	0.3 (3.58)	2.2 (5.22)	0.1 (1.52)	2.7 (4.76)
		Median	1.5	4.0	1.0	1.0	1.0	2.0
		Min, Max	-4, 13	-3, 17	-7, 5	-4, 15	-3, 2	-7, 17

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Appendix 15.3.6.4

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_labchem.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.6.10 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Aspartate aminotransferase (IU/L) 10-40	Baseline	n	22	22	9	9	11	73
		Mean (SD)	19.3 (2.68)	20.0 (4.08)	20.3 (3.28)	19.6 (2.75)	19.8 (3.49)	19.8 (3.29)
		Median	19.0	19.0	19.0	19.0	20.0	19.0
		Min, Max	14, 26	15, 32	15, 25	15, 24	15, 26	14, 32
		Low value - n (%)	0	0	0	0	0	0
		Normal value - n (%)	22 (100 %)	22 (100 %)	9 (100 %)	9 (100 %)	11 (100 %)	73 (100 %)
		High value - n (%)	0	0	0	0	0	0
		Abnormal clinically significant - n (%)	0	0	0	0	0	0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Appendix 15.3.6.4

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_labchem.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.6.10 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Aspartate aminotransferase (IU/L) 10-40	Day 4/ Discharge	n	22	22	9	9	11	73
		Mean (SD)	19.2 (3.46)	21.4 (5.35)	19.0 (3.75)	18.8 (5.55)	18.5 (3.33)	19.7 (4.44)
		Median	19.0	20.5	19.0	18.0	20.0	19.0
		Min, Max	13, 26	15, 32	14, 25	12, 32	13, 25	12, 32
		Low value - n (%)	0	0	0	0	0	0
		Normal value - n (%)	22 (100 %)	22 (100 %)	9 (100 %)	9 (100 %)	11 (100 %)	73 (100 %)
		High value - n (%)	0	0	0	0	0	0
		Abnormal clinically significant - n (%)	0	0	0	0	0	0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Appendix 15.3.6.4

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_labchem.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.6.10 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Overall Safety (N=73)	
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		Enrolled Not Randomized (N=11)
Aspartate aminotransferase (IU/L) 10-40	Change from n Baseline		22	22	9	9	11	73
		Mean (SD)	-0.1 (3.00)	1.4 (3.13)	-1.3 (3.43)	-0.8 (4.36)	-1.4 (1.97)	-0.1 (3.25)
		Median	-1.0	0.5	0.0	-1.0	-1.0	0.0
		Min, Max	-5, 5	-5, 10	-10, 2	-6, 9	-6, 2	-10, 10

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Appendix 15.3.6.4

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_labchem.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.6.10 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Bilirubin (mg/dL) 0.2-1.2	Baseline	n	22	22	9	9	11	73
		Mean (SD)	0.95 (0.306)	0.91 (0.452)	0.84 (0.275)	0.81 (0.170)	0.83 (0.291)	0.89 (0.336)
		Median	0.90	0.80	0.80	0.90	0.80	0.90
		Min, Max	0.4, 1.8	0.4, 2.1	0.5, 1.4	0.6, 1.0	0.5, 1.4	0.4, 2.1
		Low value - n (%)	0	0	0	0	0	0
		Normal value - n (%)	20 ( 90.9%)	19 ( 86.4%)	8 ( 88.9%)	9 (100 %)	9 ( 81.8%)	65 ( 89.0%)
		High value - n (%)	2 ( 9.1%)	3 ( 13.6%)	1 ( 11.1%)	0	2 ( 18.2%)	8 ( 11.0%)
		Abnormal clinically significant - n (%)	0	1 ( 4.5%)	0	0	0	1 ( 1.4%)

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Appendix 15.3.6.4

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_labchem.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.6.10 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Bilirubin (mg/dL) 0.2-1.2	Day 4/ Discharge	n	22	22	9	9	11	73
		Mean (SD)	0.78 (0.271)	0.76 (0.305)	0.73 (0.174)	0.72 (0.199)	0.81 (0.362)	0.76 (0.275)
		Median	0.70	0.70	0.80	0.70	0.70	0.70
		Min, Max	0.4, 1.4	0.4, 1.5	0.5, 1.0	0.5, 1.1	0.5, 1.7	0.4, 1.7
		Low value - n (%)	0	0	0	0	0	0
		Normal value - n (%)	21 ( 95.5%)	20 ( 90.9%)	9 (100 %)	9 (100 %)	9 ( 81.8%)	68 ( 93.2%)
		High value - n (%)	1 ( 4.5%)	2 ( 9.1%)	0	0	2 ( 18.2%)	5 ( 6.8%)
		Abnormal clinically significant - n (%)	0	0	0	0	0	0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Appendix 15.3.6.4

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_labchem.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.6.10 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Bilirubin (mg/dL) 0.2-1.2	Change from n Baseline		22	22	9	9	11	73
		Mean (SD)	-0.17 (0.279)	-0.15 (0.327)	-0.11 (0.227)	-0.09 (0.117)	-0.02 (0.141)	-0.13 (0.258)
		Median	-0.10	-0.10	0.00	-0.10	0.00	-0.10
		Min, Max	-0.6, 0.5	-1.1, 0.3	-0.6, 0.1	-0.3, 0.1	-0.2, 0.3	-1.1, 0.5

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Appendix 15.3.6.4

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_labchem.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.6.10 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Direct bilirubin (mg/dL) 0-0.3	Baseline	n	22	22	9	9	11	73
		Mean (SD)	0.22 (0.096)	0.20 (0.114)	0.18 (0.084)	0.16 (0.053)	0.16 (0.068)	0.19 (0.094)
		Median	0.20	0.20	0.20	0.20	0.20	0.20
		Min, Max	0.1, 0.4	0.1, 0.5	0.1, 0.3	0.1, 0.2	0.1, 0.3	0.1, 0.5
		Low value - n (%)	0	0	0	0	0	0
		Normal value - n (%)	20 ( 90.9%)	19 ( 86.4%)	9 (100 %)	9 (100 %)	11 (100 %)	68 ( 93.2%)
		High value - n (%)	2 ( 9.1%)	3 ( 13.6%)	0	0	0	5 ( 6.8%)
		Abnormal clinically significant - n (%)	0	0	0	0	0	0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Appendix 15.3.6.4

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_labchem.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.6.10 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Direct bilirubin (mg/dL) 0-0.3	Day 4/ Discharge	n	22	22	9	9	11	73
		Mean (SD)	0.18 (0.076)	0.17 (0.078)	0.14 (0.053)	0.14 (0.053)	0.15 (0.094)	0.16 (0.074)
		Median	0.20	0.20	0.10	0.10	0.10	0.20
		Min, Max	0.1, 0.3	0.1, 0.4	0.1, 0.2	0.1, 0.2	0.1, 0.4	0.1, 0.4
		Low value - n (%)	0	0	0	0	0	0
		Normal value - n (%)	22 (100 %)	21 ( 95.5%)	9 (100 %)	9 (100 %)	10 ( 90.9%)	71 ( 97.3%)
		High value - n (%)	0	1 ( 4.5%)	0	0	1 ( 9.1%)	2 ( 2.7%)
		Abnormal clinically significant - n (%)	0	0	0	0	0	0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Appendix 15.3.6.4

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_labchem.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.6.10 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Direct bilirubin (mg/dL) 0-0.3	Change from n Baseline		22	22	9	9	11	73
		Mean (SD)	-0.04 (0.067)	-0.04 (0.073)	-0.03 (0.050)	-0.01 (0.034)	-0.01 (0.054)	-0.03 (0.062)
		Median	0.00	0.00	0.00	0.00	0.00	0.00
		Min, Max	-0.2, 0.1	-0.2, 0.1	-0.1, 0.0	-0.1, 0.0	-0.1, 0.1	-0.2, 0.1

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Appendix 15.3.6.4

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_labchem.sas

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**Table 15.2.6.10 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Alkaline phosphatase (IU/L) 100-325	Baseline	n	22	22	9	9	11	73
		Mean (SD)	185.2 (57.02)	187.5 (50.53)	195.3 (38.73)	211.1 (81.08)	188.6 (26.58)	190.9 (52.50)
		Median	175.5	187.5	193.0	216.0	186.0	186.0
		Min, Max	113, 337	90, 275	136, 245	103, 365	152, 225	90, 365
		Low value - n (%)	0	1 ( 4.5%)	0	0	0	1 ( 1.4%)
		Normal value - n (%)	21 ( 95.5%)	21 ( 95.5%)	9 (100 %)	8 ( 88.9%)	11 (100 %)	70 ( 95.9%)
		High value - n (%)	1 ( 4.5%)	0	0	1 ( 11.1%)	0	2 ( 2.7%)
		Abnormal clinically significant - n (%)	0	0	0	0	0	0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Appendix 15.3.6.4

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_labchem.sas

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**Table 15.2.6.10 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Alkaline phosphatase (IU/L) 100-325	Day 4/ Discharge	n	22	22	9	9	11	73
		Mean (SD)	179.8 (48.17)	187.5 (53.31)	188.3 (36.49)	208.1 (75.98)	188.1 (28.81)	187.9 (49.82)
		Median	173.5	183.5	198.0	193.0	188.0	185.0
		Min, Max	109, 284	79, 302	134, 244	106, 334	149, 235	79, 334
		Low value - n (%)	0	1 ( 4.5%)	0	0	0	1 ( 1.4%)
		Normal value - n (%)	22 (100 %)	21 ( 95.5%)	9 (100 %)	8 ( 88.9%)	11 (100 %)	71 ( 97.3%)
		High value - n (%)	0	0	0	1 ( 11.1%)	0	1 ( 1.4%)
		Abnormal clinically significant - n (%)	0	0	0	0	0	0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Appendix 15.3.6.4

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_labchem.sas

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**Table 15.2.6.10 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Alkaline phosphatase (IU/L) 100-325	Change from n Baseline		22	22	9	9	11	73
		Mean (SD)	-5.4 (15.73)	0.0 (10.58)	-7.0 (12.46)	-3.0 (17.55)	-0.5 (8.41)	-3.0 (13.17)
		Median	-3.0	-0.5	-10.0	-2.0	-1.0	-2.0
		Min, Max	-53, 16	-18, 27	-24, 12	-31, 21	-15, 11	-53, 27

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Appendix 15.3.6.4

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_labchem.sas

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**Table 15.2.6.10 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Gamma glutamyl transferase (IU/L) <80 (Males)	Baseline	n	11	12	5	4	5	37
		Mean (SD)	17.1 (6.31)	23.9 (13.45)	28.4 (13.47)	21.3 (14.53)	23.6 (20.11)	22.2 (12.79)
		Median	13.0	18.5	24.0	18.5	16.0	19.0
		Min, Max	11, 30	13, 51	13, 48	7, 41	11, 59	7, 59
		Low value - n (%)	0	0	0	0	0	0
		Normal value - n (%)	11 ( 50.0%)	12 ( 54.5%)	5 ( 55.6%)	4 ( 44.4%)	5 ( 45.5%)	37 ( 50.7%)
		High value - n (%)	0	0	0	0	0	0
		Abnormal clinically significant - n (%)	0	0	0	0	0	0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Note: Gamma Glutamyl Transferase and Creatinine are summarized separately by sex.

Appendix 15.3.6.4

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_labchem.sas

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**Table 15.2.6.10 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Gamma glutamyl transferase (IU/L) <80 (Males)	Day 4/ Discharge	n	11	12	5	4	5	37
		Mean (SD)	15.9 (5.53)	22.2 (12.35)	25.0 (11.07)	24.3 (19.14)	23.0 (20.83)	21.0 (12.64)
		Median	14.0	17.0	23.0	19.5	15.0	17.0
		Min, Max	10, 28	12, 47	10, 39	7, 51	11, 60	7, 60
		Low value - n (%)	0	0	0	0	0	0
		Normal value - n (%)	11 ( 50.0%)	12 ( 54.5%)	5 ( 55.6%)	4 ( 44.4%)	5 ( 45.5%)	37 ( 50.7%)
		High value - n (%)	0	0	0	0	0	0
		Abnormal clinically significant - n (%)	0	0	0	0	0	0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Note: Gamma Glutamyl Transferase and Creatinine are summarized separately by sex.

Appendix 15.3.6.4

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**Table 15.2.6.10 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Overall Safety (N=73)	
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		Enrolled Not Randomized (N=11)
Gamma glutamyl transferase (IU/L) <80 (Males)	Change from n Baseline		11	12	5	4	5	37
		Mean (SD)	-1.2 (1.48)	-1.8 (1.87)	-3.4 (3.29)	3.0 (4.77)	-0.6 (1.52)	-1.1 (2.80)
		Median	-1.0	-1.0	-3.0	1.0	0.0	-1.0
		Min, Max	-3, 2	-6, 1	-9, -1	0, 10	-3, 1	-9, 10

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Note: Gamma Glutamyl Transferase and Creatinine are summarized separately by sex.

Appendix 15.3.6.4

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**Table 15.2.6.10 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Gamma glutamyl transferase (IU/L) <30 (Females)	Baseline	n	11	10	4	5	6	36
		Mean (SD)	20.9 (14.16)	11.6 (2.46)	18.5 (13.97)	14.8 (7.95)	18.0 (10.81)	16.7 (10.66)
		Median	15.0	11.5	14.5	15.0	14.5	13.0
		Min, Max	8, 52	8, 16	7, 38	7, 25	9, 39	7, 52
		Low value - n (%)	0	0	0	0	0	0
		Normal value - n (%)	9 ( 40.9%)	10 ( 45.5%)	3 ( 33.3%)	5 ( 55.6%)	5 ( 45.5%)	32 ( 43.8%)
		High value - n (%)	2 ( 9.1%)	0	1 ( 11.1%)	0	1 ( 9.1%)	4 ( 5.5%)
		Abnormal clinically significant - n (%)	0	0	0	0	0	0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Note: Gamma Glutamyl Transferase and Creatinine are summarized separately by sex.

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**Table 15.2.6.10 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Gamma glutamyl transferase (IU/L) <30 (Females)	Day 4/ Discharge	n	11	10	4	5	6	36
		Mean (SD)	19.5 (11.96)	11.5 (3.14)	17.8 (12.50)	14.6 (6.43)	19.0 (10.92)	16.3 (9.51)
		Median	15.0	12.0	13.5	14.0	15.0	13.5
		Min, Max	7, 42	7, 16	8, 36	8, 23	10, 40	7, 42
		Low value - n (%)	0	0	0	0	0	0
		Normal value - n (%)	9 ( 40.9%)	10 ( 45.5%)	3 ( 33.3%)	5 ( 55.6%)	5 ( 45.5%)	32 ( 43.8%)
		High value - n (%)	2 ( 9.1%)	0	1 ( 11.1%)	0	1 ( 9.1%)	4 ( 5.5%)
		Abnormal clinically significant - n (%)	0	0	0	0	0	0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Note: Gamma Glutamyl Transferase and Creatinine are summarized separately by sex.

Appendix 15.3.6.4

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**Table 15.2.6.10 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Overall Safety (N=73)	
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		Enrolled Not Randomized (N=11)
Gamma glutamyl transferase (IU/L) <30 (Females)	Change from n Baseline		11	10	4	5	6	36
		Mean (SD)	-1.4 (4.35)	-0.1 (1.45)	-0.8 (2.76)	-0.2 (1.65)	1.0 (0.64)	-0.4 (2.76)
		Median	0.0	0.0	-0.5	-1.0	1.0	0.0
		Min, Max	-13, 3	-2, 3	-4, 2	-2, 2	0, 2	-13, 3

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Note: Gamma Glutamyl Transferase and Creatinine are summarized separately by sex.

Appendix 15.3.6.4

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**Table 15.2.6.10 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Lactate dehydrogenase (IU/L) 120-240	Baseline	n	22	22	9	9	11	73
		Mean (SD)	133.6 (15.79)	135.1 (19.13)	131.8 (18.73)	134.2 (15.02)	144.2 (27.76)	135.5 (19.11)
		Median	131.5	132.5	129.0	133.0	134.0	133.0
		Min, Max	107, 173	102, 168	105, 157	113, 166	114, 189	102, 189
		Low value - n (%)	3 ( 13.6%)	4 ( 18.2%)	4 ( 44.4%)	1 ( 11.1%)	2 ( 18.2%)	14 ( 19.2%)
		Normal value - n (%)	19 ( 86.4%)	18 ( 81.8%)	5 ( 55.6%)	8 ( 88.9%)	9 ( 81.8%)	59 ( 80.8%)
		High value - n (%)	0	0	0	0	0	0
		Abnormal clinically significant - n (%)	0	0	0	0	0	0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

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**Table 15.2.6.10 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Lactate dehydrogenase (IU/L) 120-240	Day 4/ Discharge	n	22	22	9	9	11	73
		Mean (SD)	122.5 (16.03)	131.2 (20.68)	121.8 (14.59)	125.7 (15.77)	137.8 (24.12)	127.7 (19.08)
		Median	120.5	131.5	118.0	125.0	128.0	124.0
		Min, Max	91, 161	91, 171	107, 145	108, 161	103, 173	91, 173
		Low value - n (%)	9 ( 40.9%)	7 ( 31.8%)	5 ( 55.6%)	4 ( 44.4%)	3 ( 27.3%)	28 ( 38.4%)
		Normal value - n (%)	13 ( 59.1%)	15 ( 68.2%)	4 ( 44.4%)	5 ( 55.6%)	8 ( 72.7%)	45 ( 61.6%)
		High value - n (%)	0	0	0	0	0	0
		Abnormal clinically significant - n (%)	0	0	0	0	0	0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

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**Table 15.2.6.10 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Lactate dehydrogenase (IU/L) 120-240	Change from n Baseline		22	22	9	9	11	73
		Mean (SD)	-11.1 (8.68)	-4.0 (9.29)	-10.0 (11.23)	-8.6 (10.02)	-6.4 (13.37)	-7.8 (10.29)
		Median	-10.0	-3.5	-11.0	-10.0	-8.0	-9.0
		Min, Max	-30, 5	-23, 15	-26, 3	-25, 6	-25, 25	-30, 25

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

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**Table 15.2.6.10 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Glucose (mg/dL) 70-109	Baseline	n	22	22	9	9	11	73
		Mean (SD)	88.7 (6.96)	87.5 (6.49)	88.6 (7.20)	87.6 (5.18)	88.9 (3.79)	88.2 (6.12)
		Median	89.0	87.0	88.0	86.0	89.0	88.0
		Min, Max	77, 108	77, 99	77, 102	81, 97	82, 94	77, 108
		Low value - n (%)	0	0	0	0	0	0
		Normal value - n (%)	22 (100 %)	22 (100 %)	9 (100 %)	9 (100 %)	11 (100 %)	73 (100 %)
		High value - n (%)	0	0	0	0	0	0
		Abnormal clinically significant - n (%)	0	0	0	0	0	0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

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**Table 15.2.6.10 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Glucose (mg/dL) 70-109	Day 4/ Discharge	n	22	22	9	9	11	73
		Mean (SD)	87.9 (5.19)	86.8 (6.54)	88.8 (5.85)	91.7 (5.50)	88.7 (8.29)	88.3 (6.26)
		Median	88.5	87.0	88.0	91.0	89.0	89.0
		Min, Max	79, 96	77, 104	80, 97	83, 102	71, 102	71, 104
		Low value - n (%)	0	0	0	0	0	0
		Normal value - n (%)	22 (100 %)	22 (100 %)	9 (100 %)	9 (100 %)	11 (100 %)	73 (100 %)
		High value - n (%)	0	0	0	0	0	0
		Abnormal clinically significant - n (%)	0	0	0	0	0	0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Appendix 15.3.6.4

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_labchem.sas

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**Table 15.2.6.10 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Glucose (mg/dL) 70-109	Change from n Baseline		22	22	9	9	11	73
		Mean (SD)	-0.8 (7.14)	-0.7 (5.83)	0.2 (6.56)	4.1 (3.34)	-0.2 (6.95)	0.1 (6.32)
		Median	0.0	-2.0	0.0	4.0	2.0	1.0
		Min, Max	-14, 13	-11, 8	-7, 11	0, 11	-11, 8	-14, 13

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Appendix 15.3.6.4

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**Table 15.2.6.10 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Creatinine (mg/dL) 0.47-0.79 (Females)	Baseline	n	11	10	4	5	6	36
		Mean (SD)	0.633 (0.0793)	0.620 (0.0517)	0.618 (0.0550)	0.578 (0.0497)	0.567 (0.0547)	0.609 (0.0642)
		Median	0.610	0.635	0.605	0.570	0.570	0.605
		Min, Max	0.51, 0.76	0.53, 0.70	0.57, 0.69	0.52, 0.65	0.49, 0.63	0.49, 0.76
		Low value - n (%)	0	0	0	0	0	0
		Normal value - n (%)	11 ( 50.0%)	10 ( 45.5%)	4 ( 44.4%)	5 ( 55.6%)	6 ( 54.5%)	36 ( 49.3%)
		High value - n (%)	0	0	0	0	0	0
		Abnormal clinically significant - n (%)	0	0	0	0	0	0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Note: Gamma Glutamyl Transferase and Creatinine are summarized separately by sex.

Appendix 15.3.6.4

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**Table 15.2.6.10 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Creatinine (mg/dL) 0.47-0.79 (Females)	Day 4/ Discharge	n	11	10	4	5	6	36
		Mean (SD)	0.583 (0.0946)	0.566 (0.0451)	0.593 (0.0728)	0.544 (0.0537)	0.510 (0.0639)	0.562 (0.0721)
		Median	0.560	0.555	0.565	0.530	0.525	0.550
		Min, Max	0.43, 0.74	0.48, 0.64	0.54, 0.70	0.47, 0.61	0.40, 0.58	0.40, 0.74
		Low value - n (%)	1 ( 4.5%)	0	0	0	1 ( 9.1%)	2 ( 2.7%)
		Normal value - n (%)	10 ( 45.5%)	10 ( 45.5%)	4 ( 44.4%)	5 ( 55.6%)	5 ( 45.5%)	34 ( 46.6%)
		High value - n (%)	0	0	0	0	0	0
		Abnormal clinically significant - n (%)	0	0	0	0	0	0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Note: Gamma Glutamyl Transferase and Creatinine are summarized separately by sex.

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**Table 15.2.6.10 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Overall Safety (N=73)	
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		Enrolled Not Randomized (N=11)
Creatinine (mg/dL) 0.47-0.79 (Females)	Change from n Baseline		11	10	4	5	6	36
		Mean (SD)	-0.050 (0.0686)	-0.054 (0.0307)	-0.025 (0.0444)	-0.034 (0.0555)	-0.057 (0.0207)	-0.047 (0.0478)
		Median	-0.070	-0.055	-0.010	-0.040	-0.050	-0.050
		Min, Max	-0.16, 0.06	-0.10, -0.01	-0.09, 0.01	-0.08, 0.06	-0.09, -0.03	-0.16, 0.06

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Note: Gamma Glutamyl Transferase and Creatinine are summarized separately by sex.

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**Table 15.2.6.10 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Creatinine (mg/dL) 0.61-1.04 (Males)	Baseline	n	11	12	5	4	5	37
		Mean (SD)	0.835 (0.1035)	0.830 (0.1200)	0.792 (0.1045)	0.770 (0.1120)	0.800 (0.1105)	0.816 (0.1074)
		Median	0.830	0.835	0.850	0.750	0.830	0.830
		Min, Max	0.64, 1.02	0.63, 0.98	0.67, 0.90	0.67, 0.91	0.61, 0.89	0.61, 1.02
		Low value - n (%)	0	0	0	0	0	0
		Normal value - n (%)	11 ( 50.0%)	12 ( 54.5%)	5 ( 55.6%)	4 ( 44.4%)	5 ( 45.5%)	37 ( 50.7%)
		High value - n (%)	0	0	0	0	0	0
		Abnormal clinically significant - n (%)	0	0	0	0	0	0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Note: Gamma Glutamyl Transferase and Creatinine are summarized separately by sex.

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**Table 15.2.6.10 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Creatinine (mg/dL) 0.61-1.04 (Males)	Day 4/ Discharge	n	11	12	5	4	5	37
		Mean (SD)	0.770 (0.1104)	0.779 (0.1259)	0.774 (0.0780)	0.733 (0.1072)	0.744 (0.0953)	0.766 (0.1056)
		Median	0.770	0.770	0.770	0.725	0.770	0.770
		Min, Max	0.56, 0.96	0.60, 0.95	0.70, 0.90	0.61, 0.87	0.58, 0.81	0.56, 0.96
		Low value - n (%)	1 ( 4.5%)	1 ( 4.5%)	0	0	1 ( 9.1%)	3 ( 4.1%)
		Normal value - n (%)	10 ( 45.5%)	11 ( 50.0%)	5 ( 55.6%)	4 ( 44.4%)	4 ( 36.4%)	34 ( 46.6%)
		High value - n (%)	0	0	0	0	0	0
		Abnormal clinically significant - n (%)	0	0	0	0	0	0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Note: Gamma Glutamyl Transferase and Creatinine are summarized separately by sex.

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**Table 15.2.6.10 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Overall Safety (N=73)	
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		Enrolled Not Randomized (N=11)
Creatinine (mg/dL) 0.61-1.04 (Males)	Change from n Baseline		11	12	5	4	5	37
		Mean (SD)	-0.065 (0.0263)	-0.051 (0.0861)	-0.018 (0.0555)	-0.038 (0.0404)	-0.056 (0.0392)	-0.050 (0.0578)
		Median	-0.060	-0.055	0.000	-0.050	-0.080	-0.060
		Min, Max	-0.10, -0.02	-0.20, 0.09	-0.08, 0.05	-0.07, 0.02	-0.09, 0.00	-0.20, 0.09

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Note: Gamma Glutamyl Transferase and Creatinine are summarized separately by sex.

Appendix 15.3.6.4

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**Table 15.2.6.10 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Protein (g/dL) 6.7-8.3	Baseline	n	22	22	9	9	11	73
		Mean (SD)	6.83 (0.445)	6.87 (0.340)	7.14 (0.434)	7.02 (0.441)	6.82 (0.473)	6.90 (0.421)
		Median	6.75	6.90	7.20	7.10	6.80	6.90
		Min, Max	6.2, 7.6	6.2, 7.8	6.5, 7.9	6.2, 7.5	5.9, 7.6	5.9, 7.9
		Low value - n (%)	10 ( 45.5%)	5 ( 22.7%)	2 ( 22.2%)	2 ( 22.2%)	4 ( 36.4%)	23 ( 31.5%)
		Normal value - n (%)	12 ( 54.5%)	17 ( 77.3%)	7 ( 77.8%)	7 ( 77.8%)	7 ( 63.6%)	50 ( 68.5%)
		High value - n (%)	0	0	0	0	0	0
		Abnormal clinically significant - n (%)	0	0	0	0	0	0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

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**Table 15.2.6.10 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Protein (g/dL) 6.7-8.3	Day 4/ Discharge	n	22	22	9	9	11	73
		Mean (SD)	6.79 (0.314)	7.00 (0.419)	7.16 (0.513)	7.02 (0.403)	6.85 (0.391)	6.94 (0.405)
		Median	6.80	7.00	7.00	6.90	6.90	6.90
		Min, Max	6.3, 7.7	6.4, 7.9	6.5, 8.0	6.6, 7.8	6.1, 7.3	6.1, 8.0
		Low value - n (%)	7 ( 31.8%)	6 ( 27.3%)	2 ( 22.2%)	1 ( 11.1%)	2 ( 18.2%)	18 ( 24.7%)
		Normal value - n (%)	15 ( 68.2%)	16 ( 72.7%)	7 ( 77.8%)	8 ( 88.9%)	9 ( 81.8%)	55 ( 75.3%)
		High value - n (%)	0	0	0	0	0	0
		Abnormal clinically significant - n (%)	0	0	0	0	0	0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

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Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_labchem.sas

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**Table 15.2.6.10 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Protein (g/dL) 6.7-8.3	Change from n Baseline		22	22	9	9	11	73
		Mean (SD)	-0.04 (0.307)	0.13 (0.273)	0.01 (0.290)	0.00 (0.343)	0.04 (0.324)	0.03 (0.301)
		Median	0.05	0.10	0.00	0.10	0.20	0.10
		Min, Max	-0.6, 0.4	-0.3, 0.7	-0.4, 0.4	-0.6, 0.5	-0.8, 0.3	-0.8, 0.7

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

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**Table 15.2.6.10 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Blood urea nitrogen (mg/dL) 8-23	Baseline	n	22	22	9	9	11	73
		Mean (SD)	11.5 (2.00)	9.9 (2.48)	9.6 (1.82)	10.2 (3.00)	10.9 (1.71)	10.5 (2.30)
		Median	11.0	10.0	10.0	11.0	11.0	11.0
		Min, Max	9, 15	5, 17	7, 12	6, 14	7, 13	5, 17
		Low value - n (%)	0	2 ( 9.1%)	1 ( 11.1%)	2 ( 22.2%)	1 ( 9.1%)	6 ( 8.2%)
		Normal value - n (%)	22 (100 %)	20 ( 90.9%)	8 ( 88.9%)	7 ( 77.8%)	10 ( 90.9%)	67 ( 91.8%)
		High value - n (%)	0	0	0	0	0	0
		Abnormal clinically significant - n (%)	0	0	0	0	0	0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

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**Table 15.2.6.10 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Blood urea nitrogen (mg/dL) 8-23	Day 4/ Discharge	n	22	22	9	9	11	73
		Mean (SD)	10.4 (2.52)	10.0 (1.99)	8.4 (1.95)	9.2 (1.40)	9.3 (1.96)	9.7 (2.14)
		Median	10.5	10.0	8.0	9.0	9.0	9.0
		Min, Max	6, 16	6, 14	6, 12	7, 11	7, 12	6, 16
		Low value - n (%)	3 ( 13.6%)	2 ( 9.1%)	4 ( 44.4%)	1 ( 11.1%)	2 ( 18.2%)	12 ( 16.4%)
		Normal value - n (%)	19 ( 86.4%)	20 ( 90.9%)	5 ( 55.6%)	8 ( 88.9%)	9 ( 81.8%)	61 ( 83.6%)
		High value - n (%)	0	0	0	0	0	0
		Abnormal clinically significant - n (%)	0	0	0	0	0	0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

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**Table 15.2.6.10 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Overall Safety (N=73)	
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		Enrolled Not Randomized (N=11)
Blood urea nitrogen (mg/dL) 8-23	Change from n Baseline		22	22	9	9	11	73
		Mean (SD)	-1.1 (1.62)	0.1 (2.07)	-1.1 (2.09)	-1.0 (2.13)	-1.6 (1.86)	-0.8 (1.97)
		Median	-1.0	0.0	-1.0	-1.0	-2.0	-1.0
		Min, Max	-4, 2	-4, 4	-4, 2	-4, 1	-4, 1	-4, 4

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Appendix 15.3.6.4

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_labchem.sas

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**Table 15.2.6.10 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Triglycerides (mg/dL) 30-149	Baseline	n	22	22	9	9	11	73
		Mean (SD)	95.5 (36.29)	85.0 (32.37)	137.2 (79.75)	90.2 (61.71)	101.2 (45.92)	97.7 (48.65)
		Median	91.0	77.0	103.0	63.0	79.0	87.0
		Min, Max	37, 193	34, 143	46, 259	36, 202	38, 183	34, 259
		Low value - n (%)	0	0	0	0	0	0
		Normal value - n (%)	21 ( 95.5%)	22 (100 %)	6 ( 66.7%)	7 ( 77.8%)	9 ( 81.8%)	65 ( 89.0%)
		High value - n (%)	1 ( 4.5%)	0	3 ( 33.3%)	2 ( 22.2%)	2 ( 18.2%)	8 ( 11.0%)
		Abnormal clinically significant - n (%)	0	0	0	0	0	0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Appendix 15.3.6.4

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_labchem.sas

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**Table 15.2.6.10 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Triglycerides (mg/dL) 30-149	Day 4/ Discharge	n	22	22	9	9	11	73
		Mean (SD)	120.3 (50.09)	105.5 (42.13)	147.7 (74.39)	108.7 (64.47)	172.5 (70.42)	125.7 (59.84)
		Median	111.5	97.0	128.0	85.0	141.0	110.0
		Min, Max	52, 240	54, 231	67, 276	59, 254	73, 297	52, 297
		Low value - n (%)	0	0	0	0	0	0
		Normal value - n (%)	15 ( 68.2%)	20 ( 90.9%)	6 ( 66.7%)	7 ( 77.8%)	6 ( 54.5%)	54 ( 74.0%)
		High value - n (%)	7 ( 31.8%)	2 ( 9.1%)	3 ( 33.3%)	2 ( 22.2%)	5 ( 45.5%)	19 ( 26.0%)
		Abnormal clinically significant - n (%)	0	0	0	0	0	0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Appendix 15.3.6.4

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_labchem.sas

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**Table 15.2.6.10 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Triglycerides (mg/dL) 30-149	Change from n Baseline	n	22	22	9	9	11	73
		Mean (SD)	24.8 (43.43)	20.5 (29.98)	10.4 (35.71)	18.4 (27.30)	71.4 (49.31)	27.9 (41.62)
		Median	11.5	14.0	24.0	29.0	76.0	23.0
		Min, Max	-45, 106	-29, 92	-49, 60	-21, 52	-15, 157	-49, 157

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

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**Table 15.2.6.10 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Cholesterol (mg/dL) 120-219	Baseline	n	22	22	9	9	11	73
		Mean (SD)	185.8 (28.00)	178.3 (31.22)	187.2 (22.16)	184.9 (38.80)	187.6 (34.29)	183.9 (30.21)
		Median	182.5	172.0	195.0	183.0	183.0	183.0
		Min, Max	126, 259	128, 236	154, 216	122, 258	147, 251	122, 259
		Low value - n (%)	0	0	0	0	0	0
		Normal value - n (%)	20 ( 90.9%)	20 ( 90.9%)	9 (100 %)	8 ( 88.9%)	9 ( 81.8%)	66 ( 90.4%)
		High value - n (%)	2 ( 9.1%)	2 ( 9.1%)	0	1 ( 11.1%)	2 ( 18.2%)	7 ( 9.6%)
		Abnormal clinically significant - n (%)	0	0	0	0	0	0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Appendix 15.3.6.4

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**Table 15.2.6.10 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Cholesterol (mg/dL) 120-219	Day 4/ Discharge	n	22	22	9	9	11	73
		Mean (SD)	172.3 (25.19)	162.3 (28.84)	173.4 (21.55)	175.8 (31.12)	183.1 (33.76)	171.5 (28.19)
		Median	166.5	150.5	175.0	171.0	175.0	168.0
		Min, Max	129, 222	119, 230	133, 201	137, 239	145, 249	119, 249
		Low value - n (%)	0	1 ( 4.5%)	0	0	0	1 ( 1.4%)
		Normal value - n (%)	21 ( 95.5%)	20 ( 90.9%)	9 (100 %)	8 ( 88.9%)	9 ( 81.8%)	67 ( 91.8%)
		High value - n (%)	1 ( 4.5%)	1 ( 4.5%)	0	1 ( 11.1%)	2 ( 18.2%)	5 ( 6.8%)
		Abnormal clinically significant - n (%)	0	0	0	0	0	0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

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**Table 15.2.6.10 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Cholesterol (mg/dL) 120-219	Change from n Baseline	n	22	22	9	9	11	73
		Mean (SD)	-13.5 (13.59)	-16.0 (11.77)	-13.8 (17.84)	-9.1 (22.68)	-4.5 (12.91)	-12.4 (15.01)
		Median	-13.5	-19.0	-8.0	-18.0	-6.0	-13.0
		Min, Max	-42, 8	-41, 9	-41, 12	-43, 32	-32, 16	-43, 32

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

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**Table 15.2.6.10 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Albumin (g/L) 38-53	Baseline	n	22	22	9	9	11	73
		Mean (SD)	44.0 (3.00)	43.9 (2.61)	45.8 (2.54)	45.7 (3.40)	44.7 (4.08)	44.5 (3.08)
		Median	43.0	43.0	46.0	46.0	44.0	44.0
		Min, Max	40, 51	40, 50	43, 50	41, 50	36, 52	36, 52
		Low value - n (%)	0	0	0	0	1 ( 9.1%)	1 ( 1.4%)
		Normal value - n (%)	22 (100 %)	22 (100 %)	9 (100 %)	9 (100 %)	10 ( 90.9%)	72 ( 98.6%)
		High value - n (%)	0	0	0	0	0	0
		Abnormal clinically significant - n (%)	0	0	0	0	0	0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

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**Table 15.2.6.10 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Albumin (g/L) 38-53	Day 4/ Discharge	n	22	22	9	9	11	73
		Mean (SD)	43.5 (2.88)	44.5 (2.99)	45.8 (3.04)	45.6 (2.97)	44.9 (2.55)	44.6 (2.93)
		Median	43.0	44.0	45.0	45.0	45.0	44.0
		Min, Max	39, 50	39, 50	41, 51	42, 51	40, 49	39, 51
		Low value - n (%)	0	0	0	0	0	0
		Normal value - n (%)	22 (100 %)	22 (100 %)	9 (100 %)	9 (100 %)	11 (100 %)	73 (100 %)
		High value - n (%)	0	0	0	0	0	0
		Abnormal clinically significant - n (%)	0	0	0	0	0	0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

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**Table 15.2.6.10 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Albumin (g/L) 38-53	Change from n Baseline		22	22	9	9	11	73
		Mean (SD)	-0.5 (2.35)	0.6 (1.99)	0.0 (2.00)	-0.1 (1.91)	0.2 (2.23)	0.1 (2.13)
		Median	-0.5	1.0	0.0	0.0	0.0	0.0
		Min, Max	-5, 4	-3, 5	-2, 4	-4, 3	-5, 4	-5, 5

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

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**Table 15.2.6.10 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Sodium (mmol/L) 137-147	Baseline	n	22	22	9	9	11	73
		Mean (SD)	140.5 (1.93)	140.6 (1.82)	140.3 (1.50)	140.8 (1.10)	140.6 (1.13)	140.5 (1.62)
		Median	141.0	140.5	140.0	141.0	140.0	141.0
		Min, Max	137, 144	138, 145	137, 142	139, 143	139, 143	137, 145
		Low value - n (%)	0	0	0	0	0	0
		Normal value - n (%)	22 (100 %)	22 (100 %)	9 (100 %)	9 (100 %)	11 (100 %)	73 (100 %)
		High value - n (%)	0	0	0	0	0	0
		Abnormal clinically significant - n (%)	0	0	0	0	0	0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

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**Table 15.2.6.10 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Sodium (mmol/L) 137-147	Day 4/ Discharge	n	22	22	9	9	11	73
		Mean (SD)	140.6 (1.30)	140.7 (1.25)	140.4 (1.51)	141.1 (1.06)	142.1 (2.17)	140.9 (1.51)
		Median	141.0	141.0	141.0	141.0	143.0	141.0
		Min, Max	139, 144	139, 143	138, 142	140, 143	139, 146	138, 146
		Low value - n (%)	0	0	0	0	0	0
		Normal value - n (%)	22 (100 %)	22 (100 %)	9 (100 %)	9 (100 %)	11 (100 %)	73 (100 %)
		High value - n (%)	0	0	0	0	0	0
		Abnormal clinically significant - n (%)	0	0	0	0	0	0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

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**Table 15.2.6.10 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Sodium (mmol/L) 137-147	Change from n Baseline		22	22	9	9	11	73
		Mean (SD)	0.2 (1.79)	0.1 (1.51)	0.1 (1.37)	0.3 (1.23)	1.5 (2.63)	0.4 (1.78)
		Median	1.0	0.0	0.0	0.0	2.0	0.0
		Min, Max	-4, 3	-4, 2	-2, 2	-1, 3	-2, 7	-4, 7

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

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**Table 15.2.6.10 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Potassium (mmol/L) 3.5-5	Baseline	n	22	22	9	9	11	73
		Mean (SD)	4.12 (0.202)	3.93 (0.284)	4.13 (0.230)	4.16 (0.301)	4.05 (0.247)	4.06 (0.261)
		Median	4.10	3.90	4.10	4.20	4.10	4.10
		Min, Max	3.7, 4.5	3.2, 4.5	3.8, 4.6	3.8, 4.6	3.8, 4.6	3.2, 4.6
		Low value - n (%)	0	1 ( 4.5%)	0	0	0	1 ( 1.4%)
		Normal value - n (%)	22 (100 %)	21 ( 95.5%)	9 (100 %)	9 (100 %)	11 (100 %)	72 ( 98.6%)
		High value - n (%)	0	0	0	0	0	0
		Abnormal clinically significant - n (%)	0	0	0	0	0	0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

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**Table 15.2.6.10 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Potassium (mmol/L) 3.5-5	Day 4/ Discharge	n	22	22	9	9	11	73
		Mean (SD)	3.98 (0.180)	4.05 (0.293)	3.98 (0.223)	4.20 (0.260)	4.00 (0.314)	4.03 (0.258)
		Median	4.00	4.00	4.10	4.10	4.00	4.00
		Min, Max	3.7, 4.5	3.5, 4.8	3.6, 4.2	3.9, 4.7	3.6, 4.5	3.5, 4.8
		Low value - n (%)	0	0	0	0	0	0
		Normal value - n (%)	22 (100 %)	22 (100 %)	9 (100 %)	9 (100 %)	11 (100 %)	73 (100 %)
		High value - n (%)	0	0	0	0	0	0
		Abnormal clinically significant - n (%)	0	0	0	0	0	0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Appendix 15.3.6.4

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_labchem.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.6.10 Summary of Clinical Chemistry Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Potassium (mmol/L) 3.5-5	Change from n Baseline		22	22	9	9	11	73
		Mean (SD)	-0.14 (0.214)	0.13 (0.385)	-0.16 (0.321)	0.04 (0.434)	-0.05 (0.234)	-0.02 (0.332)
		Median	-0.10	0.20	-0.20	-0.10	-0.10	0.00
		Min, Max	-0.6, 0.2	-0.5, 1.2	-0.6, 0.3	-0.5, 0.8	-0.5, 0.4	-0.6, 1.2

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Appendix 15.3.6.4

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_labchem.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.6.11 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Erythrocytes (T/L) 3.8-5 (Females)	Baseline	n	11	10	4	5	6	36
		Mean (SD)	4.223 (0.1499)	4.033 (0.3860)	4.435 (0.4206)	4.114 (0.2781)	4.147 (0.3085)	4.166 (0.3116)
		Median	4.190	4.010	4.535	4.150	4.075	4.160
		Min, Max	4.03, 4.46	3.58, 4.68	3.84, 4.83	3.72, 4.42	3.87, 4.68	3.58, 4.83
		Low value - n (%)	0	3 ( 13.6%)	0	1 ( 11.1%)	0	4 ( 5.5%)
		Normal value - n (%)	11 ( 50.0%)	7 ( 31.8%)	4 ( 44.4%)	4 ( 44.4%)	6 ( 54.5%)	32 ( 43.8%)
		High value - n (%)	0	0	0	0	0	0
		Abnormal clinically significant - n (%)	0	0	0	0	0	0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Note: Erythrocytes, Hemoglobin and Hematocrit are summarized separately by sex.

Appendix 15.3.6.5

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_labhem.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.6.11 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Erythrocytes (T/L) 3.8-5 (Females)	Day 4/ Discharge	n	11	10	4	5	6	36
		Mean (SD)	4.082 (0.1328)	3.960 (0.3730)	4.233 (0.4532)	3.980 (0.1337)	4.103 (0.2839)	4.054 (0.2821)
		Median	4.090	4.000	4.420	4.030	4.035	4.070
		Min, Max	3.79, 4.23	3.43, 4.63	3.56, 4.53	3.80, 4.10	3.84, 4.62	3.43, 4.63
		Low value - n (%)	1 ( 4.5%)	4 ( 18.2%)	1 ( 11.1%)	0	0	6 ( 8.2%)
		Normal value - n (%)	10 ( 45.5%)	6 ( 27.3%)	3 ( 33.3%)	5 ( 55.6%)	6 ( 54.5%)	30 ( 41.1%)
		High value - n (%)	0	0	0	0	0	0
		Abnormal clinically significant - n (%)	0	0	0	0	0	0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Note: Erythrocytes, Hemoglobin and Hematocrit are summarized separately by sex.

Appendix 15.3.6.5

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_labhem.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.6.11 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Overall Safety (N=73)	
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		Enrolled Not Randomized (N=11)
Erythrocytes (T/L) 3.8-5 (Females)	Change from n Baseline		11	10	4	5	6	36
		Mean (SD)	-0.141 (0.2034)	-0.073 (0.1890)	-0.203 (0.1145)	-0.134 (0.2275)	-0.043 (0.0525)	-0.112 (0.1759)
		Median	-0.140	-0.035	-0.230	-0.220	-0.055	-0.095
		Min, Max	-0.53, 0.20	-0.33, 0.31	-0.30, -0.05	-0.39, 0.13	-0.10, 0.02	-0.53, 0.31

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Note: Erythrocytes, Hemoglobin and Hematocrit are summarized separately by sex.

Appendix 15.3.6.5

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_labhem.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.6.11 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Erythrocytes (T/L) 4.3-5.7 (Males)	Baseline	n	11	12	5	4	5	37
		Mean (SD)	4.704 (0.3769)	4.739 (0.2702)	4.964 (0.4120)	4.950 (0.2056)	4.840 (0.2050)	4.795 (0.3152)
		Median	4.730	4.750	5.110	4.950	4.790	4.800
		Min, Max	4.21, 5.40	4.26, 5.16	4.49, 5.36	4.70, 5.20	4.61, 5.17	4.21, 5.40
		Low value - n (%)	3 ( 13.6%)	1 ( 4.5%)	0	0	0	4 ( 5.5%)
		Normal value - n (%)	8 ( 36.4%)	11 ( 50.0%)	5 ( 55.6%)	4 ( 44.4%)	5 ( 45.5%)	33 ( 45.2%)
		High value - n (%)	0	0	0	0	0	0
		Abnormal clinically significant - n (%)	0	0	0	0	0	0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Note: Erythrocytes, Hemoglobin and Hematocrit are summarized separately by sex.

Appendix 15.3.6.5

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_labhem.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.6.11 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Erythrocytes (T/L) 4.3-5.7 (Males)	Day 4/ Discharge	n	11	12	5	4	5	37
		Mean (SD)	4.556 (0.3249)	4.639 (0.3253)	4.952 (0.3816)	4.800 (0.3966)	4.756 (0.1235)	4.690 (0.3323)
		Median	4.540	4.745	4.980	4.735	4.780	4.750
		Min, Max	4.07, 5.14	3.89, 5.07	4.36, 5.39	4.45, 5.28	4.56, 4.90	3.89, 5.39
		Low value - n (%)	3 ( 13.6%)	1 ( 4.5%)	0	0	0	4 ( 5.5%)
		Normal value - n (%)	8 ( 36.4%)	11 ( 50.0%)	5 ( 55.6%)	4 ( 44.4%)	5 ( 45.5%)	33 ( 45.2%)
		High value - n (%)	0	0	0	0	0	0
		Abnormal clinically significant - n (%)	0	0	0	0	0	0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Note: Erythrocytes, Hemoglobin and Hematocrit are summarized separately by sex.

Appendix 15.3.6.5

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_labhem.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.6.11 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Overall Safety (N=73)	
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		Enrolled Not Randomized (N=11)
Erythrocytes (T/L) 4.3-5.7 (Males)	Change from n Baseline		11	12	5	4	5	37
		Mean (SD)	-0.147 (0.1625)	-0.100 (0.1241)	-0.012 (0.2053)	-0.150 (0.2432)	-0.084 (0.1881)	-0.105 (0.1662)
		Median	-0.200	-0.065	-0.130	-0.105	-0.050	-0.090
		Min, Max	-0.37, 0.12	-0.37, 0.08	-0.16, 0.33	-0.47, 0.08	-0.39, 0.11	-0.47, 0.33

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Note: Erythrocytes, Hemoglobin and Hematocrit are summarized separately by sex.

Appendix 15.3.6.5

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_labhem.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.6.11 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Hemoglobin (g/dL) 11.5-15 (Females)	Baseline	n	11	10	4	5	6	36
		Mean (SD)	13.15 (0.619)	12.91 (0.861)	13.43 (0.940)	12.78 (1.260)	13.38 (0.585)	13.10 (0.811)
		Median	13.00	12.75	13.60	12.90	13.45	13.10
		Min, Max	12.1, 14.0	11.8, 14.4	12.2, 14.3	10.8, 14.3	12.7, 14.0	10.8, 14.4
		Low value - n (%)	0	0	0	1 ( 11.1%)	0	1 ( 1.4%)
		Normal value - n (%)	11 ( 50.0%)	10 ( 45.5%)	4 ( 44.4%)	4 ( 44.4%)	6 ( 54.5%)	35 ( 47.9%)
		High value - n (%)	0	0	0	0	0	0
		Abnormal clinically significant - n (%)	0	0	0	0	0	0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Note: Erythrocytes, Hemoglobin and Hematocrit are summarized separately by sex.

Appendix 15.3.6.5

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_labhem.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.6.11 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Hemoglobin (g/dL) 11.5-15 (Females)	Day 4/ Discharge	n	11	10	4	5	6	36
		Mean (SD)	12.80 (0.548)	12.74 (0.773)	12.88 (1.261)	12.36 (0.802)	13.33 (0.551)	12.82 (0.755)
		Median	12.90	12.90	13.00	12.20	13.55	12.90
		Min, Max	11.6, 13.4	11.7, 14.3	11.3, 14.2	11.2, 13.3	12.6, 13.8	11.2, 14.3
		Low value - n (%)	0	0	1 ( 11.1%)	1 ( 11.1%)	0	2 ( 2.7%)
		Normal value - n (%)	11 ( 50.0%)	10 ( 45.5%)	3 ( 33.3%)	4 ( 44.4%)	6 ( 54.5%)	34 ( 46.6%)
		High value - n (%)	0	0	0	0	0	0
		Abnormal clinically significant - n (%)	0	0	0	0	0	0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Note: Erythrocytes, Hemoglobin and Hematocrit are summarized separately by sex.

Appendix 15.3.6.5

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_labhem.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.6.11 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Overall Safety (N=73)	
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		Enrolled Not Randomized (N=11)
Hemoglobin (g/dL) 11.5-15 (Females)	Change from n Baseline		11	10	4	5	6	36
		Mean (SD)	-0.35 (0.684)	-0.17 (0.626)	-0.55 (0.342)	-0.42 (0.730)	-0.05 (0.198)	-0.28 (0.580)
		Median	-0.40	-0.05	-0.60	-0.60	-0.10	-0.20
		Min, Max	-1.7, 0.8	-1.0, 1.2	-0.9, -0.1	-1.4, 0.4	-0.2, 0.3	-1.7, 1.2

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Note: Erythrocytes, Hemoglobin and Hematocrit are summarized separately by sex.

Appendix 15.3.6.5

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_labhem.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.6.11 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Hemoglobin (g/dL) 13.5-17.5 (Males)	Baseline	n	11	12	5	4	5	37
		Mean (SD)	15.06 (1.192)	15.30 (0.951)	15.32 (0.868)	14.90 (0.688)	15.64 (0.372)	15.24 (0.925)
		Median	14.50	15.00	15.70	14.65	15.70	15.30
		Min, Max	13.6, 17.5	13.7, 17.3	14.1, 16.3	14.4, 15.9	15.3, 16.2	13.6, 17.5
		Low value - n (%)	0	0	0	0	0	0
		Normal value - n (%)	11 ( 50.0%)	12 ( 54.5%)	5 ( 55.6%)	4 ( 44.4%)	5 ( 45.5%)	37 ( 50.7%)
		High value - n (%)	0	0	0	0	0	0
		Abnormal clinically significant - n (%)	0	0	0	0	0	0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Note: Erythrocytes, Hemoglobin and Hematocrit are summarized separately by sex.

Appendix 15.3.6.5

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_labhem.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.6.11 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Hemoglobin (g/dL) 13.5-17.5 (Males)	Day 4/ Discharge	n	11	12	5	4	5	37
		Mean (SD)	14.66 (1.002)	14.89 (0.964)	15.22 (0.650)	14.43 (1.103)	15.40 (0.255)	14.89 (0.900)
		Median	14.70	14.85	15.40	14.20	15.50	15.00
		Min, Max	13.1, 16.8	13.6, 16.9	14.2, 15.8	13.4, 15.9	15.0, 15.6	13.1, 16.9
		Low value - n (%)	1 ( 4.5%)	0	0	1 ( 11.1%)	0	2 ( 2.7%)
		Normal value - n (%)	10 ( 45.5%)	12 ( 54.5%)	5 ( 55.6%)	3 ( 33.3%)	5 ( 45.5%)	35 ( 47.9%)
		High value - n (%)	0	0	0	0	0	0
		Abnormal clinically significant - n (%)	0	0	0	0	0	0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Note: Erythrocytes, Hemoglobin and Hematocrit are summarized separately by sex.

Appendix 15.3.6.5

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_labhem.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.6.11 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Hemoglobin (g/dL) 13.5-17.5 (Males)	Change from n Baseline		11	12	5	4	5	37
		Mean (SD)	-0.40 (0.470)	-0.41 (0.451)	-0.10 (0.605)	-0.48 (0.728)	-0.24 (0.434)	-0.35 (0.493)
		Median	-0.50	-0.30	-0.30	-0.35	-0.20	-0.30
		Min, Max	-1.0, 0.4	-1.3, 0.2	-0.6, 0.9	-1.4, 0.2	-0.9, 0.3	-1.4, 0.9

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Note: Erythrocytes, Hemoglobin and Hematocrit are summarized separately by sex.

Appendix 15.3.6.5

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_labhem.sas

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**Table 15.2.6.11 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Hematocrit (%) 34.8-45 (Females)	Baseline	n	11	10	4	5	6	36
		Mean (SD)	38.68 (1.732)	37.99 (2.350)	39.50 (2.632)	37.94 (3.324)	39.38 (1.620)	38.59 (2.209)
		Median	38.50	37.65	40.40	38.40	39.45	38.60
		Min, Max	35.6, 41.3	34.9, 41.8	35.7, 41.5	32.6, 41.8	37.5, 41.2	32.6, 41.8
		Low value - n (%)	0	0	0	1 ( 11.1%)	0	1 ( 1.4%)
		Normal value - n (%)	11 ( 50.0%)	10 ( 45.5%)	4 ( 44.4%)	4 ( 44.4%)	6 ( 54.5%)	35 ( 47.9%)
		High value - n (%)	0	0	0	0	0	0
		Abnormal clinically significant - n (%)	0	0	0	0	0	0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Note: Erythrocytes, Hemoglobin and Hematocrit are summarized separately by sex.

Appendix 15.3.6.5

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_labhem.sas

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**Table 15.2.6.11 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Hematocrit (%) 34.8-45 (Females)	Day 4/ Discharge	n	11	10	4	5	6	36
		Mean (SD)	37.44 (1.383)	37.26 (2.570)	37.73 (3.125)	36.74 (2.632)	39.20 (1.275)	37.62 (2.166)
		Median	37.50	37.75	38.35	36.10	39.15	37.80
		Min, Max	34.6, 39.3	33.3, 42.2	33.5, 40.7	33.2, 40.1	37.3, 40.9	33.2, 42.2
		Low value - n (%)	1 ( 4.5%)	2 ( 9.1%)	1 ( 11.1%)	1 ( 11.1%)	0	5 ( 6.8%)
		Normal value - n (%)	10 ( 45.5%)	8 ( 36.4%)	3 ( 33.3%)	4 ( 44.4%)	6 ( 54.5%)	31 ( 42.5%)
		High value - n (%)	0	0	0	0	0	0
		Abnormal clinically significant - n (%)	0	0	0	0	0	0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Note: Erythrocytes, Hemoglobin and Hematocrit are summarized separately by sex.

Appendix 15.3.6.5

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**Table 15.2.6.11 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Overall Safety (N=73)	
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		Enrolled Not Randomized (N=11)
Hematocrit (%) 34.8-45 (Females)	Change from n Baseline		11	10	4	5	6	36
		Mean (SD)	-1.25 (1.780)	-0.73 (1.762)	-1.78 (0.714)	-1.20 (2.125)	-0.18 (0.602)	-0.98 (1.595)
		Median	-1.00	-0.55	-1.95	-2.20	-0.25	-0.85
		Min, Max	-4.6, 1.8	-3.3, 2.8	-2.4, -0.8	-3.4, 1.5	-0.9, 0.7	-4.6, 2.8

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Note: Erythrocytes, Hemoglobin and Hematocrit are summarized separately by sex.

Appendix 15.3.6.5

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_labhem.sas

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**Table 15.2.6.11 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Hematocrit (%) 39.7-52.4 (Males)	Baseline	n	11	12	5	4	5	37
		Mean (SD)	44.85 (3.327)	45.32 (2.504)	45.44 (2.870)	44.78 (1.069)	45.84 (1.218)	45.21 (2.509)
		Median	44.20	44.70	45.80	44.90	45.50	44.70
		Min, Max	40.3, 51.3	41.5, 50.6	41.2, 48.3	43.5, 45.8	44.4, 47.5	40.3, 51.3
		Low value - n (%)	0	0	0	0	0	0
		Normal value - n (%)	11 ( 50.0%)	12 ( 54.5%)	5 ( 55.6%)	4 ( 44.4%)	5 ( 45.5%)	37 ( 50.7%)
		High value - n (%)	0	0	0	0	0	0
		Abnormal clinically significant - n (%)	0	0	0	0	0	0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Note: Erythrocytes, Hemoglobin and Hematocrit are summarized separately by sex.

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**Table 15.2.6.11 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Hematocrit (%) 39.7-52.4 (Males)	Day 4/ Discharge	n	11	12	5	4	5	37
		Mean (SD)	43.10 (2.624)	43.99 (3.022)	44.70 (1.722)	43.20 (2.399)	44.90 (1.840)	43.86 (2.524)
		Median	43.10	43.70	45.50	42.90	43.70	43.80
		Min, Max	38.5, 48.5	39.3, 50.7	42.1, 46.1	40.9, 46.1	43.5, 47.5	38.5, 50.7
		Low value - n (%)	1 ( 4.5%)	1 ( 4.5%)	0	0	0	2 ( 2.7%)
		Normal value - n (%)	10 ( 45.5%)	11 ( 50.0%)	5 ( 55.6%)	4 ( 44.4%)	5 ( 45.5%)	35 ( 47.9%)
		High value - n (%)	0	0	0	0	0	0
		Abnormal clinically significant - n (%)	0	0	0	0	0	0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Note: Erythrocytes, Hemoglobin and Hematocrit are summarized separately by sex.

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**Table 15.2.6.11 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Overall Safety (N=73)	
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		Enrolled Not Randomized (N=11)
Hematocrit (%) 39.7-52.4 (Males)	Change from n Baseline		11	12	5	4	5	37
		Mean (SD)	-1.75 (1.556)	-1.33 (1.250)	-0.74 (2.016)	-1.58 (2.722)	-0.94 (1.986)	-1.35 (1.669)
		Median	-2.50	-0.90	-1.70	-1.05	-0.80	-1.00
		Min, Max	-3.3, 0.4	-4.1, 0.2	-2.2, 2.6	-4.9, 0.7	-4.0, 0.9	-4.9, 2.6

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Note: Erythrocytes, Hemoglobin and Hematocrit are summarized separately by sex.

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**Table 15.2.6.11 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Leukocytes (G/L) 3.3-9	Baseline	n	22	22	9	9	11	73
		Mean (SD)	6.78 (1.233)	6.50 (1.608)	6.26 (1.103)	6.22 (1.660)	5.92 (1.575)	6.43 (1.439)
		Median	6.70	6.05	5.70	5.60	5.50	6.10
		Min, Max	3.9, 8.6	3.4, 9.8	4.5, 7.9	4.6, 10.0	4.3, 9.0	3.4, 10.0
		Low value - n (%)	0	0	0	0	0	0
		Normal value - n (%)	22 (100 %)	20 ( 90.9%)	9 (100 %)	8 ( 88.9%)	11 (100 %)	70 ( 95.9%)
		High value - n (%)	0	2 ( 9.1%)	0	1 ( 11.1%)	0	3 ( 4.1%)
		Abnormal clinically significant - n (%)	0	0	0	0	0	0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

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**Table 15.2.6.11 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Leukocytes (G/L) 3.3-9	Day 4/ Discharge	n	22	22	9	9	11	73
		Mean (SD)	5.96 (1.204)	5.57 (1.277)	5.64 (1.138)	5.50 (0.966)	5.79 (1.302)	5.72 (1.189)
		Median	5.70	5.85	5.70	5.20	5.40	5.70
		Min, Max	4.3, 9.2	3.5, 7.8	3.8, 7.5	4.1, 6.9	4.2, 7.9	3.5, 9.2
		Low value - n (%)	0	0	0	0	0	0
		Normal value - n (%)	21 ( 95.5%)	22 (100 %)	9 (100 %)	9 (100 %)	11 (100 %)	72 ( 98.6%)
		High value - n (%)	1 ( 4.5%)	0	0	0	0	1 ( 1.4%)
		Abnormal clinically significant - n (%)	0	0	0	0	0	0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

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**Table 15.2.6.11 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Overall Safety (N=73)	
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		Enrolled Not Randomized (N=11)
Leukocytes (G/L) 3.3-9	Change from n Baseline		22	22	9	9	11	73
		Mean (SD)	-0.81 (0.952)	-0.92 (0.811)	-0.61 (1.153)	-0.72 (1.406)	-0.13 (1.372)	-0.71 (1.075)
		Median	-0.90	-0.80	-0.70	-0.90	-0.20	-0.80
		Min, Max	-2.4, 1.0	-2.4, 0.4	-2.0, 1.8	-3.1, 2.0	-2.5, 2.4	-3.1, 2.4

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Appendix 15.3.6.5

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**Table 15.2.6.11 Summary of Hematology Parameters - Safety Population**

Parameter (units)	Reference range	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
				THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Neutrophils (G/L)		Baseline	n	22	22	9	9	11	73
			Mean (SD)	3.64 (1.091)	3.43 (1.254)	3.36 (0.916)	3.47 (1.316)	2.94 (0.986)	3.42 (1.130)
			Median	3.55	3.40	3.20	3.20	2.70	3.40
			Min, Max	2.0, 6.1	1.5, 6.5	1.9, 5.0	2.0, 6.2	1.8, 5.1	1.5, 6.5

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Appendix 15.3.6.5

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_labhem.sas

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**Table 15.2.6.11 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Neutrophils (G/L)	Day 4/ Discharge	n	22	22	9	9	11	73
		Mean (SD)	3.60 (1.146)	3.22 (0.734)	3.48 (1.053)	3.38 (0.767)	3.58 (1.396)	3.44 (1.013)
		Median	3.50	3.15	3.40	3.30	3.50	3.30
		Min, Max	2.1, 6.9	1.9, 4.8	2.2, 5.4	2.2, 4.3	1.8, 5.9	1.8, 6.9

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

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**Table 15.2.6.11 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Overall Safety (N=73)	
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		Enrolled Not Randomized (N=11)
Neutrophils (G/L)	Change from n Baseline		22	22	9	9	11	73
		Mean (SD)	-0.04 (0.685)	-0.21 (0.796)	0.12 (1.035)	-0.09 (1.013)	0.65 (1.194)	0.03 (0.912)
		Median	-0.15	0.00	0.20	0.00	0.50	0.00
		Min, Max	-1.0, 1.4	-2.1, 0.9	-1.0, 2.5	-1.9, 1.6	-1.2, 3.4	-2.1, 3.4

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Appendix 15.3.6.5

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**Table 15.2.6.11 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Neutrophils/ leukocytes (%) 40-75	Baseline	n	22	22	9	9	11	73
		Mean (SD)	53.13 (8.398)	51.75 (9.004)	52.82 (7.971)	54.91 (9.764)	49.26 (7.219)	52.32 (8.468)
		Median	53.35	52.60	52.60	57.00	52.00	53.00
		Min, Max	40.3, 72.0	36.9, 71.0	41.3, 65.5	37.0, 68.0	32.7, 56.8	32.7, 72.0
		Low value - n (%)	0	3 ( 13.6%)	0	1 ( 11.1%)	1 ( 9.1%)	5 ( 6.8%)
		Normal value - n (%)	22 (100 %)	19 ( 86.4%)	9 (100 %)	8 ( 88.9%)	10 ( 90.9%)	68 ( 93.2%)
		High value - n (%)	0	0	0	0	0	0
		Abnormal clinically significant - n (%)	0	0	0	0	0	0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

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**Table 15.2.6.11 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Neutrophils/ leukocytes (%) 40-75	Day 4/ Discharge	n	22	22	9	9	11	73
		Mean (SD)	59.35 (9.053)	58.14 (4.970)	60.12 (6.960)	60.91 (6.364)	59.95 (12.496)	59.36 (7.972)
		Median	60.55	57.30	57.10	59.50	63.50	59.00
		Min, Max	42.4, 75.0	50.6, 68.1	50.7, 71.7	53.0, 71.5	37.8, 81.3	37.8, 81.3
		Low value - n (%)	0	0	0	0	1 ( 9.1%)	1 ( 1.4%)
		Normal value - n (%)	22 (100 %)	22 (100 %)	9 (100 %)	9 (100 %)	9 ( 81.8%)	71 ( 97.3%)
		High value - n (%)	0	0	0	0	1 ( 9.1%)	1 ( 1.4%)
		Abnormal clinically significant - n (%)	0	0	0	0	0	0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Appendix 15.3.6.5

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_labhem.sas

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**Table 15.2.6.11 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Overall Safety (N=73)	
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		Enrolled Not Randomized (N=11)
Neutrophils/ leukocytes (%) 40-75	Change from n Baseline		22	22	9	9	11	73
		Mean (SD)	6.22 (6.544)	6.39 (7.423)	7.30 (6.404)	6.00 (6.030)	10.68 (7.960)	7.05 (6.962)
		Median	5.65	6.20	7.70	5.80	11.50	6.90
		Min, Max	-8.7, 16.4	-4.8, 18.6	-1.5, 20.7	-2.5, 18.0	-1.1, 28.8	-8.7, 28.8

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Appendix 15.3.6.5

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**Table 15.2.6.11 Summary of Hematology Parameters - Safety Population**

Parameter (units)	Reference range	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
				THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Lymphocytes (G/L)		Baseline	n	22	22	9	9	11	73
			Mean (SD)	2.37 (0.597)	2.36 (0.581)	2.16 (0.451)	2.20 (0.513)	2.28 (0.705)	2.31 (0.574)
			Median	2.35	2.30	2.00	2.10	1.90	2.30
			Min, Max	1.4, 4.4	1.2, 3.5	1.5, 3.0	1.5, 2.9	1.5, 3.8	1.2, 4.4

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

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**Table 15.2.6.11 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Lymphocytes (G/L)	Day 4/ Discharge	n	22	22	9	9	11	73
		Mean (SD)	1.76 (0.441)	1.77 (0.500)	1.62 (0.304)	1.66 (0.256)	1.65 (0.451)	1.72 (0.424)
		Median	1.65	1.85	1.60	1.70	1.50	1.70
		Min, Max	1.2, 2.6	0.9, 2.8	1.3, 2.1	1.3, 2.0	1.1, 2.6	0.9, 2.8

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

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**Table 15.2.6.11 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Lymphocytes (G/L)	Change from n Baseline		22	22	9	9	11	73
		Mean (SD)	-0.61 (0.537)	-0.59 (0.364)	-0.53 (0.391)	-0.54 (0.442)	-0.64 (0.439)	-0.59 (0.435)
		Median	-0.65	-0.55	-0.60	-0.40	-0.70	-0.60
		Min, Max	-2.0, 0.5	-1.1, 0.2	-1.0, 0.2	-1.2, 0.2	-1.3, 0.2	-2.0, 0.5

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

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**Table 15.2.6.11 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Lymphocytes/ leukocytes (%) 18-49	Baseline	n	22	22	9	9	11	73
		Mean (SD)	35.51 (7.638)	37.32 (8.115)	34.66 (6.585)	36.22 (8.367)	38.76 (7.717)	36.53 (7.669)
		Median	36.85	35.65	34.90	33.00	39.00	36.20
		Min, Max	16.5, 51.0	24.0, 49.4	23.0, 44.1	29.2, 53.5	30.1, 56.7	16.5, 56.7
		Low value - n (%)	1 ( 4.5%)	0	0	0	0	1 ( 1.4%)
		Normal value - n (%)	20 ( 90.9%)	20 ( 90.9%)	9 (100 %)	8 ( 88.9%)	10 ( 90.9%)	67 ( 91.8%)
		High value - n (%)	1 ( 4.5%)	2 ( 9.1%)	0	1 ( 11.1%)	1 ( 9.1%)	5 ( 6.8%)
		Abnormal clinically significant - n (%)	0	0	0	0	0	0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

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**Table 15.2.6.11 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Lymphocytes/ leukocytes (%) 18-49	Day 4/ Discharge	n	22	22	9	9	11	73
		Mean (SD)	30.03 (7.419)	31.67 (4.487)	29.69 (6.749)	30.40 (4.821)	29.33 (8.854)	30.42 (6.421)
		Median	28.60	32.95	29.30	30.50	27.00	29.50
		Min, Max	17.3, 41.5	24.5, 38.2	17.3, 39.7	23.0, 37.9	15.4, 44.6	15.4, 44.6
		Low value - n (%)	1 ( 4.5%)	0	1 ( 11.1%)	0	1 ( 9.1%)	3 ( 4.1%)
		Normal value - n (%)	21 ( 95.5%)	22 (100 %)	8 ( 88.9%)	9 (100 %)	10 ( 90.9%)	70 ( 95.9%)
		High value - n (%)	0	0	0	0	0	0
		Abnormal clinically significant - n (%)	0	0	0	0	0	0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

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**Table 15.2.6.11 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Lymphocytes/ leukocytes (%) 18-49	Change from n Baseline		22	22	9	9	11	73
	Mean (SD)		-5.48 (5.758)	-5.65 (6.704)	-4.97 (6.846)	-5.82 (5.425)	-9.44 (6.502)	-6.10 (6.259)
	Median		-4.30	-5.65	-4.40	-6.10	-8.90	-5.70
	Min, Max		-16.3, 4.5	-17.4, 5.3	-17.6, 6.0	-15.6, 2.2	-23.7, -1.8	-23.7, 6.0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

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**Table 15.2.6.11 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Monocytes (G/L)	Baseline	n	22	22	9	9	11	73
		Mean (SD)	0.451 (0.1220)	0.432 (0.1452)	0.442 (0.0937)	0.349 (0.1491)	0.405 (0.0782)	0.425 (0.1258)
		Median	0.435	0.405	0.440	0.300	0.390	0.400
		Min, Max	0.26, 0.66	0.21, 0.69	0.31, 0.58	0.13, 0.56	0.30, 0.52	0.13, 0.69

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

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**Table 15.2.6.11 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Monocytes (G/L)	Day 4/ Discharge	n	22	22	9	9	11	73
		Mean (SD)	0.394 (0.1155)	0.353 (0.1363)	0.331 (0.1093)	0.328 (0.1042)	0.355 (0.1347)	0.360 (0.1223)
		Median	0.385	0.330	0.310	0.370	0.380	0.350
		Min, Max	0.19, 0.66	0.13, 0.70	0.20, 0.56	0.15, 0.48	0.15, 0.52	0.13, 0.70

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

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**Table 15.2.6.11 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Overall Safety (N=73)	
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		Enrolled Not Randomized (N=11)
Monocytes (G/L)	Change from n Baseline		22	22	9	9	11	73
		Mean (SD)	-0.057 (0.1278)	-0.079 (0.1355)	-0.111 (0.0964)	-0.021 (0.1435)	-0.050 (0.1543)	-0.065 (0.1318)
		Median	-0.090	-0.085	-0.130	-0.050	-0.080	-0.090
		Min, Max	-0.26, 0.20	-0.31, 0.13	-0.24, 0.08	-0.18, 0.20	-0.36, 0.20	-0.36, 0.20

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

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**Table 15.2.6.11 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Monocytes/ leukocytes (%) 2-10	Baseline	n	22	22	9	9	11	73
		Mean (SD)	6.73 (1.635)	6.69 (1.624)	7.16 (1.376)	5.62 (1.980)	7.00 (1.032)	6.67 (1.590)
		Median	6.80	6.95	7.50	5.60	7.10	6.90
		Min, Max	3.0, 9.6	3.0, 8.9	4.6, 8.6	2.0, 8.3	5.4, 8.4	2.0, 9.6
		Low value - n (%)	0	0	0	0	0	0
		Normal value - n (%)	22 (100 %)	22 (100 %)	9 (100 %)	9 (100 %)	11 (100 %)	73 (100 %)
		High value - n (%)	0	0	0	0	0	0
		Abnormal clinically significant - n (%)	0	0	0	0	0	0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

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**Table 15.2.6.11 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Monocytes/ leukocytes (%) 2-10	Day 4/ Discharge	n	22	22	9	9	11	73
		Mean (SD)	6.72 (1.954)	6.30 (1.797)	5.84 (1.192)	5.98 (1.807)	6.48 (2.994)	6.36 (1.979)
		Median	6.65	6.05	5.40	5.50	6.70	5.90
		Min, Max	3.2, 11.0	3.5, 11.3	4.8, 8.2	3.1, 9.0	2.0, 11.1	2.0, 11.3
		Low value - n (%)	0	0	0	0	0	0
		Normal value - n (%)	21 ( 95.5%)	21 ( 95.5%)	9 (100 %)	9 (100 %)	9 ( 81.8%)	69 ( 94.5%)
		High value - n (%)	1 ( 4.5%)	1 ( 4.5%)	0	0	2 ( 18.2%)	4 ( 5.5%)
		Abnormal clinically significant - n (%)	0	0	0	0	0	0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

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**Table 15.2.6.11 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Overall Safety (N=73)	
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		Enrolled Not Randomized (N=11)
Monocytes/ leukocytes (%) 2-10	Change from n Baseline		22	22	9	9	11	73
		Mean (SD)	-0.01 (1.739)	-0.39 (1.901)	-1.31 (1.059)	0.36 (1.659)	-0.52 (3.337)	-0.32 (2.032)
		Median	-0.30	-0.25	-1.10	0.10	-0.40	-0.40
		Min, Max	-2.8, 5.0	-3.6, 3.1	-2.9, 0.4	-2.4, 3.5	-6.2, 5.2	-6.2, 5.2

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

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**Table 15.2.6.11 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Eosinophils (G/L)	Baseline	n	22	22	9	9	11	73
		Mean (SD)	0.272 (0.2120)	0.240 (0.1572)	0.301 (0.1263)	0.183 (0.1183)	0.265 (0.1513)	0.254 (0.1670)
		Median	0.205	0.210	0.290	0.180	0.230	0.220
		Min, Max	0.12, 1.06	0.02, 0.69	0.15, 0.54	0.00, 0.38	0.06, 0.48	0.00, 1.06

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Appendix 15.3.6.5

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_labhem.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.6.11 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Eosinophils (G/L)	Day 4/ Discharge	n	22	22	9	9	11	73
		Mean (SD)	0.199 (0.1866)	0.195 (0.1388)	0.223 (0.1365)	0.131 (0.0869)	0.212 (0.0979)	0.195 (0.1438)
		Median	0.145	0.165	0.180	0.140	0.220	0.170
		Min, Max	0.00, 0.92	0.00, 0.58	0.08, 0.46	0.00, 0.26	0.01, 0.34	0.00, 0.92

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Appendix 15.3.6.5

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_labhem.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.6.11 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Overall Safety (N=73)	
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		Enrolled Not Randomized (N=11)
Eosinophils (G/L)	Change from n Baseline		22	22	9	9	11	73
		Mean (SD)	-0.073 (0.0690)	-0.044 (0.0725)	-0.078 (0.0390)	-0.052 (0.0367)	-0.053 (0.1047)	-0.059 (0.0703)
		Median	-0.080	-0.035	-0.080	-0.050	-0.050	-0.060
		Min, Max	-0.22, 0.05	-0.14, 0.08	-0.15, -0.01	-0.12, 0.00	-0.22, 0.07	-0.22, 0.08

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Appendix 15.3.6.5

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_labhem.sas

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**Table 15.2.6.11 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Eosinophils/ leukocytes (%) 0-8	Baseline	n	22	22	9	9	11	73
		Mean (SD)	4.13 (3.230)	3.79 (2.477)	4.91 (2.099)	2.96 (1.634)	4.46 (2.529)	4.03 (2.607)
		Median	3.40	3.75	5.00	2.80	4.40	3.70
		Min, Max	1.4, 16.5	0.5, 11.7	2.4, 7.9	0.0, 5.7	1.0, 9.6	0.0, 16.5
		Low value - n (%)	0	0	0	0	0	0
		Normal value - n (%)	20 ( 90.9%)	21 ( 95.5%)	9 (100 %)	9 (100 %)	10 ( 90.9%)	69 ( 94.5%)
		High value - n (%)	2 ( 9.1%)	1 ( 4.5%)	0	0	1 ( 9.1%)	4 ( 5.5%)
		Abnormal clinically significant - n (%)	0	0	0	0	0	0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

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**Table 15.2.6.11 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Eosinophils/ leukocytes (%) 0-8	Day 4/ Discharge	n	22	22	9	9	11	73
		Mean (SD)	3.49 (3.252)	3.42 (2.169)	4.07 (2.655)	2.32 (1.405)	3.91 (2.170)	3.46 (2.521)
		Median	2.80	2.85	3.20	2.10	4.00	2.90
		Min, Max	0.0, 16.1	0.0, 9.4	1.6, 8.5	0.0, 4.3	0.1, 8.0	0.0, 16.1
		Low value - n (%)	0	0	0	0	0	0
		Normal value - n (%)	20 ( 90.9%)	21 ( 95.5%)	7 ( 77.8%)	9 (100 %)	11 (100 %)	68 ( 93.2%)
		High value - n (%)	2 ( 9.1%)	1 ( 4.5%)	2 ( 22.2%)	0	0	5 ( 6.8%)
		Abnormal clinically significant - n (%)	0	0	0	0	0	0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

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**Table 15.2.6.11 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Overall Safety (N=73)	
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		Enrolled Not Randomized (N=11)
Eosinophils/ leukocytes (%) 0-8	Change from n Baseline		22	22	9	9	11	73
		Mean (SD)	-0.64 (0.950)	-0.37 (1.431)	-0.84 (1.011)	-0.63 (0.775)	-0.55 (1.655)	-0.57 (1.202)
		Median	-0.65	-0.30	-1.00	-0.40	-1.10	-0.60
		Min, Max	-2.4, 1.1	-2.3, 3.3	-1.9, 0.8	-1.9, 0.2	-3.2, 2.2	-3.2, 3.3

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

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**Table 15.2.6.11 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Basophils (G/L)	Baseline	n	22	22	9	9	11	73
		Mean (SD)	0.0360 (0.02612)	0.0302 (0.02756)	0.0293 (0.02302)	0.0190 (0.01509)	0.0301 (0.02941)	0.0304 (0.02551)
		Median	0.0320	0.0205	0.0230	0.0260	0.0270	0.0260
		Min, Max	0.000, 0.086	0.000, 0.091	0.000, 0.077	0.000, 0.040	0.000, 0.107	0.000, 0.107

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

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**Table 15.2.6.11 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Basophils (G/L)	Day 4/ Discharge	n	22	22	9	9	11	73
		Mean (SD)	0.0252 (0.01743)	0.0250 (0.01799)	0.0159 (0.01082)	0.0226 (0.02046)	0.0196 (0.02383)	0.0228 (0.01823)
		Median	0.0200	0.0210	0.0160	0.0210	0.0110	0.0190
		Min, Max	0.010, 0.092	0.000, 0.091	0.000, 0.034	0.000, 0.066	0.000, 0.079	0.000, 0.092

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

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**Table 15.2.6.11 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Overall Safety (N=73)	
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		Enrolled Not Randomized (N=11)
Basophils (G/L)	Change from n Baseline		22	22	9	9	11	73
		Mean (SD)	-0.0107 (0.02305)	-0.0051 (0.02856)	-0.0134 (0.02988)	0.0036 (0.02721)	-0.0105 (0.02315)	-0.0076 (0.02598)
		Median	-0.0075	-0.0020	-0.0090	0.0000	-0.0170	-0.0070
		Min, Max	-0.068, 0.031	-0.064, 0.045	-0.077, 0.034	-0.021, 0.066	-0.049, 0.028	-0.077, 0.066

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

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**Table 15.2.6.11 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Basophils/ leukocytes (%) 0-2	Baseline	n	22	22	9	9	11	73
		Mean (SD)	0.50 (0.318)	0.45 (0.361)	0.46 (0.313)	0.29 (0.221)	0.51 (0.473)	0.45 (0.346)
		Median	0.50	0.35	0.50	0.40	0.40	0.40
		Min, Max	0.0, 1.0	0.0, 1.0	0.0, 1.0	0.0, 0.5	0.0, 1.7	0.0, 1.7
		Low value - n (%)	0	0	0	0	0	0
		Normal value - n (%)	22 (100 %)	22 (100 %)	9 (100 %)	9 (100 %)	11 (100 %)	73 (100 %)
		High value - n (%)	0	0	0	0	0	0
		Abnormal clinically significant - n (%)	0	0	0	0	0	0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

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**Table 15.2.6.11 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Basophils/ leukocytes (%) 0-2	Day 4/ Discharge	n	22	22	9	9	11	73
		Mean (SD)	0.40 (0.192)	0.47 (0.381)	0.28 (0.179)	0.39 (0.318)	0.34 (0.367)	0.40 (0.301)
		Median	0.40	0.40	0.30	0.40	0.20	0.40
		Min, Max	0.2, 1.0	0.0, 1.9	0.0, 0.6	0.0, 1.0	0.0, 1.0	0.0, 1.9
		Low value - n (%)	0	0	0	0	0	0
		Normal value - n (%)	22 (100 %)	22 (100 %)	9 (100 %)	9 (100 %)	11 (100 %)	73 (100 %)
		High value - n (%)	0	0	0	0	0	0
		Abnormal clinically significant - n (%)	0	0	0	0	0	0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

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**Table 15.2.6.11 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Basophils/ leukocytes (%) 0-2	Change from n Baseline		22	22	9	9	11	73
		Mean (SD)	-0.10 (0.308)	0.02 (0.430)	-0.18 (0.436)	0.10 (0.416)	-0.17 (0.476)	-0.06 (0.403)
		Median	-0.05	0.00	-0.10	0.00	-0.30	0.00
		Min, Max	-0.8, 0.4	-0.6, 1.0	-1.0, 0.6	-0.4, 1.0	-0.9, 0.7	-1.0, 1.0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

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**Table 15.2.6.11 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Platelets (G/L) 140-340	Baseline	n	22	22	9	9	11	73
		Mean (SD)	216.7 (46.47)	235.5 (46.40)	215.3 (35.56)	233.9 (28.34)	230.5 (40.15)	226.4 (42.32)
		Median	208.0	228.5	222.0	231.0	230.0	227.0
		Min, Max	146, 338	171, 348	144, 264	198, 272	176, 318	144, 348
		Low value - n (%)	0	0	0	0	0	0
		Normal value - n (%)	22 (100 %)	21 ( 95.5%)	9 (100 %)	9 (100 %)	11 (100 %)	72 ( 98.6%)
		High value - n (%)	0	1 ( 4.5%)	0	0	0	1 ( 1.4%)
		Abnormal clinically significant - n (%)	0	0	0	0	0	0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

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**Table 15.2.6.11 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Platelets (G/L) 140-340	Day 4/ Discharge	n	21	22	9	9	11	72
		Mean (SD)	214.0 (42.69)	231.6 (42.63)	226.3 (41.19)	224.4 (37.73)	228.7 (40.12)	224.5 (41.00)
		Median	209.0	221.5	234.0	235.0	228.0	221.5
		Min, Max	151, 341	172, 341	153, 292	166, 281	172, 298	151, 341
		Low value - n (%)	0	0	0	0	0	0
		Normal value - n (%)	20 ( 90.9%)	21 ( 95.5%)	9 (100 %)	9 (100 %)	11 (100 %)	70 ( 95.9%)
		High value - n (%)	1 ( 4.5%)	1 ( 4.5%)	0	0	0	2 ( 2.7%)
		Abnormal clinically significant - n (%)	0	0	0	0	0	0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

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**Table 15.2.6.11 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Platelets (G/L) 140-340	Change from n Baseline		21	22	9	9	11	72
		Mean (SD)	-3.0 (15.44)	-3.9 (18.52)	11.0 (15.90)	-9.4 (17.91)	-1.8 (14.23)	-2.2 (17.08)
		Median	-6.0	-2.0	9.0	-6.0	-2.0	-2.5
		Min, Max	-23, 35	-42, 33	-13, 31	-33, 23	-21, 27	-42, 35

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

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**Table 15.2.6.11 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Ery. mean corpuscular hemoglobin (pg) 28-34	Baseline	n	22	22	9	9	11	73
		Mean (SD)	31.60 (1.024)	32.23 (1.529)	30.69 (1.616)	30.64 (1.698)	32.33 (0.980)	31.67 (1.465)
		Median	31.60	32.10	30.80	30.90	32.80	31.80
		Min, Max	29.6, 33.5	29.9, 35.0	27.3, 33.0	27.7, 33.0	29.9, 33.2	27.3, 35.0
		Low value - n (%)	0	0	1 ( 11.1%)	1 ( 11.1%)	0	2 ( 2.7%)
		Normal value - n (%)	22 (100 %)	18 ( 81.8%)	8 ( 88.9%)	8 ( 88.9%)	11 (100 %)	67 ( 91.8%)
		High value - n (%)	0	4 ( 18.2%)	0	0	0	4 ( 5.5%)
		Abnormal clinically significant - n (%)	0	0	0	0	0	0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Appendix 15.3.6.5

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_labhem.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.6.11 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Ery. mean corpuscular hemoglobin (pg) 28-34	Day 4/ Discharge	n	22	22	9	9	11	73
		Mean (SD)	31.78 (1.051)	32.22 (1.521)	30.67 (1.503)	30.62 (1.514)	32.47 (1.037)	31.74 (1.451)
		Median	31.80	32.00	30.90	30.70	32.60	31.80
		Min, Max	29.4, 33.7	30.0, 35.0	27.6, 32.6	27.7, 32.4	29.9, 33.9	27.6, 35.0
		Low value - n (%)	0	0	1 ( 11.1%)	1 ( 11.1%)	0	2 ( 2.7%)
		Normal value - n (%)	22 (100 %)	18 ( 81.8%)	8 ( 88.9%)	8 ( 88.9%)	11 (100 %)	67 ( 91.8%)
		High value - n (%)	0	4 ( 18.2%)	0	0	0	4 ( 5.5%)
		Abnormal clinically significant - n (%)	0	0	0	0	0	0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Appendix 15.3.6.5

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_labhem.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.6.11 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Overall Safety (N=73)	
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		Enrolled Not Randomized (N=11)
Ery. mean corpuscular hemoglobin (pg) 28-34	Change from n Baseline		22	22	9	9	11	73
		Mean (SD)	0.18 (0.212)	-0.01 (0.343)	-0.02 (0.234)	-0.02 (0.339)	0.15 (0.391)	0.07 (0.310)
		Median	0.20	0.00	-0.10	0.00	0.00	0.10
		Min, Max	-0.2, 0.6	-0.6, 0.9	-0.4, 0.3	-0.6, 0.5	-0.3, 0.8	-0.6, 0.9

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Appendix 15.3.6.5

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_labhem.sas

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**Table 15.2.6.11 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Ery. mean corpuscular hgb concentration (g/dL) 30.2-35.1	Baseline	n	22	22	9	9	11	73
		Mean (SD)	33.78 (0.551)	33.85 (0.462)	33.86 (0.655)	33.47 (0.793)	34.04 (0.543)	33.81 (0.577)
		Median	33.85	33.85	34.20	33.40	34.10	33.90
		Min, Max	32.6, 34.9	32.9, 34.8	32.5, 34.5	32.3, 34.9	32.8, 34.7	32.3, 34.9
		Low value - n (%)	0	0	0	0	0	0
		Normal value - n (%)	22 (100 %)	22 (100 %)	9 (100 %)	9 (100 %)	11 (100 %)	73 (100 %)
		High value - n (%)	0	0	0	0	0	0
		Abnormal clinically significant - n (%)	0	0	0	0	0	0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Appendix 15.3.6.5

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_labhem.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.6.11 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Ery. mean corpuscular hgb concentration (g/dL) 30.2-35.1	Day 4/ Discharge	n	22	22	9	9	11	73
		Mean (SD)	34.10 (0.452)	34.02 (0.456)	34.06 (0.528)	33.53 (0.537)	34.15 (0.842)	34.01 (0.562)
		Median	34.10	33.95	34.20	33.60	34.30	34.00
		Min, Max	33.4, 35.2	33.3, 35.1	33.4, 34.9	32.8, 34.5	32.8, 35.5	32.8, 35.5
		Low value - n (%)	0	0	0	0	0	0
		Normal value - n (%)	21 ( 95.5%)	22 (100 %)	9 (100 %)	9 (100 %)	9 ( 81.8%)	70 ( 95.9%)
		High value - n (%)	1 ( 4.5%)	0	0	0	2 ( 18.2%)	3 ( 4.1%)
		Abnormal clinically significant - n (%)	0	0	0	0	0	0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Appendix 15.3.6.5

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_labhem.sas

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**Table 15.2.6.11 Summary of Hematology Parameters - Safety Population**

Parameter (units)	Reference range	Study Day	Statistic	Sequence				Overall Safety (N=73)	
				THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		Enrolled Not Randomized (N=11)
Ery. mean corpuscular hgb concentration (g/dL)	30.2-35.1	Change from n Baseline		22	22	9	9	11	73
		Mean (SD)		0.31 (0.411)	0.17 (0.532)	0.20 (0.368)	0.07 (0.520)	0.12 (0.587)	0.20 (0.482)
		Median		0.45	0.25	0.20	0.40	0.00	0.30
		Min, Max		-0.5, 1.1	-0.9, 1.2	-0.5, 0.9	-0.7, 0.6	-0.7, 1.1	-0.9, 1.2

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Appendix 15.3.6.5

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_labhem.sas

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**Table 15.2.6.11 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Ery. mean corpuscular volume (fL) 85-102	Baseline	n	22	22	9	9	11	73
		Mean (SD)	93.5 (3.38)	95.2 (4.21)	90.6 (4.86)	91.6 (4.01)	94.9 (3.02)	93.6 (4.11)
		Median	93.0	95.0	90.0	93.0	96.0	93.0
		Min, Max	87, 103	89, 102	82, 98	84, 97	88, 98	82, 103
		Low value - n (%)	0	0	1 ( 11.1%)	1 ( 11.1%)	0	2 ( 2.7%)
		Normal value - n (%)	21 ( 95.5%)	22 (100 %)	8 ( 88.9%)	8 ( 88.9%)	11 (100 %)	70 ( 95.9%)
		High value - n (%)	1 ( 4.5%)	0	0	0	0	1 ( 1.4%)
		Abnormal clinically significant - n (%)	0	0	0	0	0	0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

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Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_labhem.sas

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**Table 15.2.6.11 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Ery. mean corpuscular volume (fL) 85-102	Day 4/ Discharge	n	22	22	9	9	11	73
		Mean (SD)	93.2 (2.76)	94.8 (4.25)	90.1 (4.49)	91.3 (4.31)	95.2 (3.29)	93.4 (4.03)
		Median	93.0	95.0	90.0	92.0	96.0	93.0
		Min, Max	88, 100	88, 101	83, 97	84, 98	89, 99	83, 101
		Low value - n (%)	0	0	2 ( 22.2%)	1 ( 11.1%)	0	3 ( 4.1%)
		Normal value - n (%)	22 (100 %)	22 (100 %)	7 ( 77.8%)	8 ( 88.9%)	11 (100 %)	70 ( 95.9%)
		High value - n (%)	0	0	0	0	0	0
		Abnormal clinically significant - n (%)	0	0	0	0	0	0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Appendix 15.3.6.5

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_labhem.sas

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**Table 15.2.6.11 Summary of Hematology Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Overall Safety (N=73)	
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		Enrolled Not Randomized (N=11)
Ery. mean corpuscular volume (fL) 85-102	Change from n Baseline		22	22	9	9	11	73
		Mean (SD)	-0.2 (1.11)	-0.4 (1.01)	-0.4 (1.02)	-0.2 (1.21)	0.3 (0.79)	-0.2 (1.04)
		Median	0.0	-1.0	-1.0	0.0	0.0	0.0
		Min, Max	-3, 2	-2, 2	-2, 1	-2, 2	-1, 1	-3, 2

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Appendix 15.3.6.5

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_labhem.sas

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**Table 15.2.6.12 Summary of Urinalysis Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
pH 5-8	Baseline	n	22	22	9	9	11	73
		Mean (SD)	6.00 (0.463)	6.14 (0.658)	5.89 (0.334)	6.22 (0.364)	6.23 (0.262)	6.09 (0.489)
		Median	6.00	6.00	6.00	6.00	6.00	6.00
		Min, Max	5.5, 7.0	5.0, 8.0	5.5, 6.5	6.0, 7.0	6.0, 6.5	5.0, 8.0
		Low value - n (%)	0	0	0	0	0	0
		Normal value - n (%)	22 (100 %)	22 (100 %)	9 (100 %)	9 (100 %)	11 (100 %)	73 (100 %)
		High value - n (%)	0	0	0	0	0	0
		Abnormal clinically significant - n (%)	0	0	0	0	0	0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Appendix 15.3.6.6

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_laburin.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.6.12 Summary of Urinalysis Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
pH 5-8	Day 4/ Discharge	n	22	22	9	9	11	73
		Mean (SD)	6.82 (0.525)	6.80 (0.527)	6.78 (0.507)	6.89 (0.651)	6.82 (0.717)	6.82 (0.556)
		Median	6.75	6.75	6.50	7.00	7.00	7.00
		Min, Max	6.0, 7.5	6.0, 8.0	6.0, 7.5	6.0, 7.5	6.0, 7.5	6.0, 8.0
		Low value - n (%)	0	0	0	0	0	0
		Normal value - n (%)	22 (100 %)	22 (100 %)	9 (100 %)	9 (100 %)	11 (100 %)	73 (100 %)
		High value - n (%)	0	0	0	0	0	0
		Abnormal clinically significant - n (%)	0	0	0	0	0	0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Appendix 15.3.6.6

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_laburin.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.6.12 Summary of Urinalysis Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
pH 5-8	Change from n Baseline		22	22	9	9	11	73
		Mean (SD)	0.82 (0.749)	0.66 (0.606)	0.89 (0.547)	0.67 (0.500)	0.59 (0.626)	0.73 (0.630)
		Median	0.50	0.50	1.00	0.50	0.50	0.50
		Min, Max	-0.5, 2.0	0.0, 1.5	0.0, 1.5	0.0, 1.5	0.0, 1.5	-0.5, 2.0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Appendix 15.3.6.6

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_laburin.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.6.12 Summary of Urinalysis Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Specific gravity 1.002-1.03	Baseline	n	22	22	9	9	11	73
		Mean (SD)	1.0178 (0.00652)	1.0175 (0.00831)	1.0180 (0.00679)	1.0193 (0.00711)	1.0135 (0.00652)	1.0173 (0.00722)
		Median	1.0195	1.0190	1.0170	1.0210	1.0130	1.0180
		Min, Max	1.006, 1.029	1.003, 1.029	1.009, 1.026	1.006, 1.027	1.003, 1.025	1.003, 1.029
		Low value - n (%)	0	0	0	0	0	0
		Normal value - n (%)	22 (100 %)	22 (100 %)	9 (100 %)	9 (100 %)	11 (100 %)	73 (100 %)
		High value - n (%)	0	0	0	0	0	0
		Abnormal clinically significant - n (%)	0	0	0	0	0	0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Appendix 15.3.6.6

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Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.6.12 Summary of Urinalysis Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Specific gravity 1.002-1.03	Day 4/ Discharge	n	22	22	9	9	11	73
		Mean (SD)	1.0095 (0.00646)	1.0102 (0.00627)	1.0094 (0.00615)	1.0063 (0.00350)	1.0114 (0.00861)	1.0096 (0.00644)
		Median	1.0070	1.0080	1.0080	1.0050	1.0080	1.0070
		Min, Max	1.002, 1.023	1.001, 1.022	1.003, 1.023	1.002, 1.012	1.002, 1.030	1.001, 1.030
		Low value - n (%)	0	1 ( 4.5%)	0	0	0	1 ( 1.4%)
		Normal value - n (%)	22 (100 %)	21 ( 95.5%)	9 (100 %)	9 (100 %)	11 (100 %)	72 ( 98.6%)
		High value - n (%)	0	0	0	0	0	0
		Abnormal clinically significant - n (%)	0	0	0	0	0	0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

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**Table 15.2.6.12 Summary of Urinalysis Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Overall Safety (N=73)	
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		Enrolled Not Randomized (N=11)
Specific gravity 1.002-1.03	Change from n Baseline		22	22	9	9	11	73
		Mean (SD)	-0.0084 (0.00791)	-0.0073 (0.00725)	-0.0086 (0.00535)	-0.0130 (0.00862)	-0.0021 (0.00718)	-0.0077 (0.00780)
		Median	-0.0065	-0.0060	-0.0060	-0.0120	-0.0010	-0.0060
		Min, Max	-0.025, 0.003	-0.022, 0.005	-0.019, - 0.003	-0.022, 0.004	-0.018, 0.011	-0.025, 0.011

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Appendix 15.3.6.6

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_laburin.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.6.12 Summary of Urinalysis Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Bilirubin (urine) NEGATIVE	Baseline	Negative - n (%)	22 (100 %)	22 (100 %)	9 (100 %)	9 (100 %)	11 (100 %)	73 (100 %)
		Positive - n (%)	0	0	0	0	0	0
		Normal value - n (%)	22 (100 %)	22 (100 %)	9 (100 %)	9 (100 %)	11 (100 %)	73 (100 %)
		Abnormal value - n (%)	0	0	0	0	0	0
		Abnormal clinically significant - n (%)	0	0	0	0	0	0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Appendix 15.3.6.6

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_laburin.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.6.12 Summary of Urinalysis Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Bilirubin (urine) NEGATIVE	Day 4/ Discharge	Negative - n (%)	22 (100 %)	22 (100 %)	9 (100 %)	9 (100 %)	11 (100 %)	73 (100 %)
		Positive - n (%)	0	0	0	0	0	0
		Normal value - n (%)	22 (100 %)	22 (100 %)	9 (100 %)	9 (100 %)	11 (100 %)	73 (100 %)
		Abnormal value - n (%)	0	0	0	0	0	0
		Abnormal clinically significant - n (%)	0	0	0	0	0	0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Appendix 15.3.6.6

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_laburin.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.6.12 Summary of Urinalysis Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Glucose (urine) NEGATIVE	Baseline	Negative - n (%)	22 (100 %)	22 (100 %)	9 (100 %)	9 (100 %)	11 (100 %)	73 (100 %)
		Positive - n (%)	0	0	0	0	0	0
		Normal value - n (%)	22 (100 %)	22 (100 %)	9 (100 %)	9 (100 %)	11 (100 %)	73 (100 %)
		Abnormal value - n (%)	0	0	0	0	0	0
		Abnormal clinically significant - n (%)	0	0	0	0	0	0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Appendix 15.3.6.6

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_laburin.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.6.12 Summary of Urinalysis Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Glucose (urine) NEGATIVE	Day 4/ Discharge	Negative - n (%)	22 (100 %)	22 (100 %)	9 (100 %)	9 (100 %)	11 (100 %)	73 (100 %)
		Positive - n (%)	0	0	0	0	0	0
		Normal value - n (%)	22 (100 %)	22 (100 %)	9 (100 %)	9 (100 %)	11 (100 %)	73 (100 %)
		Abnormal value - n (%)	0	0	0	0	0	0
		Abnormal clinically significant - n (%)	0	0	0	0	0	0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Appendix 15.3.6.6

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_laburin.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.6.12 Summary of Urinalysis Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Nitrite NEGATIVE	Baseline	Negative - n (%)	21 ( 95.5%)	22 (100 %)	9 (100 %)	9 (100 %)	11 (100 %)	72 ( 98.6%)
		Positive - n (%)	1 ( 4.5%)	0	0	0	0	1 ( 1.4%)
		Normal value - n (%)	21 ( 95.5%)	22 (100 %)	9 (100 %)	9 (100 %)	11 (100 %)	72 ( 98.6%)
		Abnormal value - n (%)	1 ( 4.5%)	0	0	0	0	1 ( 1.4%)
		Abnormal clinically significant - n (%)	0	0	0	0	0	0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Appendix 15.3.6.6

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_laburin.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.6.12 Summary of Urinalysis Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Nitrite NEGATIVE	Day 4/ Discharge	Negative - n (%)	22 (100 %)	22 (100 %)	9 (100 %)	9 (100 %)	11 (100 %)	73 (100 %)
		Positive - n (%)	0	0	0	0	0	0
		Normal value - n (%)	22 (100 %)	22 (100 %)	9 (100 %)	9 (100 %)	11 (100 %)	73 (100 %)
		Abnormal value - n (%)	0	0	0	0	0	0
		Abnormal clinically significant - n (%)	0	0	0	0	0	0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Appendix 15.3.6.6

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_laburin.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.6.12 Summary of Urinalysis Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Protein (urine) NEGATIVE	Baseline	Negative - n (%)	22 (100 %)	22 (100 %)	9 (100 %)	8 ( 88.9%)	11 (100 %)	72 ( 98.6%)
		Trace - n (%)	0	0	0	1 ( 11.1%)	0	1 ( 1.4%)
		Positive - n (%)	0	0	0	0	0	0
		Normal value - n (%)	22 (100 %)	22 (100 %)	9 (100 %)	8 ( 88.9%)	11 (100 %)	72 ( 98.6%)
		Abnormal value - n (%)	0	0	0	1 ( 11.1%)	0	1 ( 1.4%)
		Abnormal clinically significant - n (%)	0	0	0	0	0	0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Appendix 15.3.6.6

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_laburin.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.6.12 Summary of Urinalysis Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Protein (urine) NEGATIVE	Day 4/ Discharge	Negative - n (%)	22 (100 %)	22 (100 %)	9 (100 %)	9 (100 %)	10 ( 90.9%)	72 ( 98.6%)
		Trace - n (%)	0	0	0	0	1 ( 9.1%)	1 ( 1.4%)
		Positive - n (%)	0	0	0	0	0	0
		Normal value - n (%)	22 (100 %)	22 (100 %)	9 (100 %)	9 (100 %)	10 ( 90.9%)	72 ( 98.6%)
		Abnormal value - n (%)	0	0	0	0	1 ( 9.1%)	1 ( 1.4%)
		Abnormal clinically significant - n (%)	0	0	0	0	0	0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Appendix 15.3.6.6

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_laburin.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.6.12 Summary of Urinalysis Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Occult blood NEGATIVE	Baseline	Negative - n (%)	17 ( 77.3%)	20 ( 90.9%)	6 ( 66.7%)	6 ( 66.7%)	10 ( 90.9%)	59 ( 80.8%)
		Trace - n (%)	1 ( 4.5%)	0	0	0	0	1 ( 1.4%)
		1+ - n (%)	0	0	0	1 ( 11.1%)	1 ( 9.1%)	2 ( 2.7%)
		2+ - n (%)	1 ( 4.5%)	0	1 ( 11.1%)	1 ( 11.1%)	0	3 ( 4.1%)
		3+ - n (%)	3 ( 13.6%)	2 ( 9.1%)	2 ( 22.2%)	1 ( 11.1%)	0	8 ( 11.0%)
		Positive - n (%)	0	0	0	0	0	0
		Normal value - n (%)	17 ( 77.3%)	20 ( 90.9%)	6 ( 66.7%)	6 ( 66.7%)	10 ( 90.9%)	59 ( 80.8%)
		Abnormal value - n (%)	5 ( 22.7%)	2 ( 9.1%)	3 ( 33.3%)	3 ( 33.3%)	1 ( 9.1%)	14 ( 19.2%)
		Abnormal clinically significant - n (%)	0	0	0	0	0	0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Appendix 15.3.6.6

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_laburin.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.6.12 Summary of Urinalysis Parameters - Safety Population**

Parameter (units) Reference range	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Occult blood NEGATIVE	Day 4/ Discharge	Negative - n (%)	19 ( 86.4%)	20 ( 90.9%)	8 ( 88.9%)	9 (100 %)	11 (100 %)	67 ( 91.8%)
		Trace - n (%)	1 ( 4.5%)	1 ( 4.5%)	1 ( 11.1%)	0	0	3 ( 4.1%)
		1+ - n (%)	0	1 ( 4.5%)	0	0	0	1 ( 1.4%)
		2+ - n (%)	1 ( 4.5%)	0	0	0	0	1 ( 1.4%)
		3+ - n (%)	1 ( 4.5%)	0	0	0	0	1 ( 1.4%)
		Positive - n (%)	0	0	0	0	0	0
		Normal value - n (%)	19 ( 86.4%)	20 ( 90.9%)	8 ( 88.9%)	9 (100 %)	11 (100 %)	67 ( 91.8%)
		Abnormal value - n (%)	3 ( 13.6%)	2 ( 9.1%)	1 ( 11.1%)	0	0	6 ( 8.2%)
		Abnormal clinically significant - n (%)	0	0	0	0	0	0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Appendix 15.3.6.6

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_laburin.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.6.13 Summary of Supine Vital Signs - Safety Population**

Parameter (units)	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Supine Systolic Blood Pressure (mmHg)	Baseline	n	22	22	9	9	11	73
		Mean (SD)	104.7 (9.18)	109.0 (10.30)	110.9 (9.54)	109.0 (11.77)	106.4 (6.86)	107.5 (9.62)
		Median	101.5	108.5	112.0	111.0	107.0	107.0
		Min, Max	92, 127	92, 127	94, 124	92, 129	95, 114	92, 129
	Day 0	n	22	22	9	9	5	67
		Mean (SD)	104.6 (8.48)	108.3 (9.77)	108.4 (8.67)	110.2 (7.25)	106.2 (8.65)	107.2 (8.80)
		Median	103.5	107.5	107.0	111.0	105.0	106.0
		Min, Max	90, 119	91, 129	99, 127	100, 120	94, 115	90, 129

Note: mCC = menthol Conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Appendix 15.3.6.7

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_vit.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.6.13 Summary of Supine Vital Signs - Safety Population**

Parameter (units)	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Supine Systolic Blood Pressure (mmHg)	Change from baseline on Day 0	n	22	22	9	9	5	67
		Mean (SD)	-0.1 (5.23)	-0.7 (6.59)	-2.4 (9.94)	1.2 (8.78)	-1.2 (4.50)	-0.5 (6.79)
		Median	-0.5	-1.0	0.0	2.0	1.0	0.0
		Min, Max	-9, 11	-15, 10	-23, 8	-18, 11	-9, 2	-23, 11
	Day 1	n	22	22	9	9	0	62
		Mean (SD)	104.4 (7.78)	108.5 (9.19)	109.0 (6.31)	113.2 (9.72)		107.8 (8.74)
		Median	103.5	108.0	110.0	117.0		107.0
		Min, Max	92, 120	93, 128	96, 117	99, 125		92, 128

Note: mCC = menthol Conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Appendix 15.3.6.7

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_vit.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.6.13 Summary of Supine Vital Signs - Safety Population**

Parameter (units)	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Supine Systolic Blood Pressure (mmHg)	Change from baseline on Day 1	n	22	22	9	9	0	62
		Mean (SD)	-0.3 (7.19)	-0.5 (7.30)	-1.9 (6.85)	4.2 (6.84)		0.1 (7.19)
		Median	-0.5	-0.5	2.0	7.0		1.0
		Min, Max	-14, 15	-17, 14	-16, 5	-12, 11		-17, 15
	Day 2	n	21	22	9	9	0	61
		Mean (SD)	103.7 (6.53)	106.1 (8.15)	106.6 (9.80)	106.4 (9.08)		105.4 (7.93)
		Median	105.0	104.5	104.0	105.0		105.0
		Min, Max	94, 116	95, 119	94, 121	93, 118		93, 121

Note: mCC = menthol Conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Appendix 15.3.6.7

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_vit.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.6.13 Summary of Supine Vital Signs - Safety Population**

Parameter (units)	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Supine Systolic Blood Pressure (mmHg)	Change from baseline on Day 2	n	21	22	9	9	0	61
		Mean (SD)	-1.2 (7.30)	-2.9 (8.28)	-4.3 (5.94)	-2.6 (6.73)		-2.5 (7.33)
		Median	0.0	-2.0	-5.0	-1.0		-2.0
		Min, Max	-17, 12	-16, 15	-11, 8	-12, 10		-17, 15
	Day 3	n	21	22	9	9	0	61
		Mean (SD)	106.7 (9.67)	109.3 (9.79)	112.0 (8.78)	110.7 (8.36)		109.0 (9.38)
		Median	105.0	111.0	112.0	108.0		111.0
		Min, Max	93, 126	93, 130	95, 126	98, 121		93, 130

Note: mCC = menthol Conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Appendix 15.3.6.7

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_vit.sas

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**Table 15.2.6.13 Summary of Supine Vital Signs - Safety Population**

Parameter (units)	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Supine Systolic Blood Pressure (mmHg)	Change from baseline on Day 3	n	21	22	9	9	0	61
		Mean (SD)	1.8 (7.00)	0.3 (8.45)	1.1 (7.61)	1.7 (5.57)		1.1 (7.32)
		Median	1.0	1.0	1.0	2.0		1.0
		Min, Max	-9, 19	-16, 13	-12, 13	-10, 8		-16, 19
	Day 4/ Discharge	n	22	22	9	9	11	73
		Mean (SD)	108.0 (8.54)	111.4 (8.73)	113.0 (10.94)	112.9 (14.14)	107.2 (8.16)	110.1 (9.66)
		Median	108.0	111.0	114.0	109.0	107.0	109.0
		Min, Max	82, 124	96, 130	99, 130	93, 129	94, 125	82, 130

Note: mCC = menthol Conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Appendix 15.3.6.7

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_vit.sas

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**Table 15.2.6.13 Summary of Supine Vital Signs - Safety Population**

Parameter (units)	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Supine Systolic Blood Pressure (mmHg)	Change from baseline on Day 4/ Discharge	n	22	22	9	9	11	73
		Mean (SD)	3.3 (8.32)	2.4 (9.60)	2.1 (9.85)	3.9 (8.63)	0.8 (5.46)	2.6 (8.44)
		Median	4.5	4.5	1.0	9.0	0.0	3.0
		Min, Max	-12, 16	-16, 18	-8, 20	-12, 13	-7, 12	-16, 20
Supine Diastolic Blood Pressure (mmHg)	Baseline	n	22	22	9	9	11	73
		Mean (SD)	57.4 (5.18)	57.9 (6.66)	57.0 (5.66)	55.1 (5.04)	56.9 (5.98)	57.1 (5.73)
		Median	56.5	56.5	58.0	55.0	56.0	57.0
		Min, Max	49, 70	48, 73	45, 63	46, 62	48, 66	45, 73

Note: mCC = menthol Conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Appendix 15.3.6.7

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_vit.sas

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**Table 15.2.6.13 Summary of Supine Vital Signs - Safety Population**

Parameter (units)	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Supine Diastolic Blood Pressure (mmHg)	Day 0	n	22	22	9	9	5	67
		Mean (SD)	54.4 (4.50)	55.9 (7.70)	55.6 (7.46)	58.1 (8.01)	56.4 (5.13)	55.7 (6.55)
		Median	55.0	53.0	53.0	58.0	54.0	54.0
		Min, Max	47, 64	46, 80	46, 71	49, 71	52, 65	46, 80
	Change from baseline on Day 0	n	22	22	9	9	5	67
		Mean (SD)	-3.0 (4.95)	-2.0 (7.24)	-1.4 (8.74)	3.0 (6.75)	2.8 (3.43)	-1.2 (6.70)
		Median	-3.5	-2.0	-1.0	3.0	3.0	-1.0
		Min, Max	-11, 7	-13, 10	-16, 12	-7, 12	-2, 6	-16, 12

Note: mCC = menthol Conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Appendix 15.3.6.7

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**Table 15.2.6.13 Summary of Supine Vital Signs - Safety Population**

Parameter (units)	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Supine Diastolic Blood Pressure (mmHg)	Day 1	n	22	22	9	9	0	62
		Mean (SD)	56.2 (4.50)	56.6 (6.53)	54.4 (6.47)	59.4 (8.85)		56.6 (6.27)
		Median	56.5	56.0	52.0	58.0		56.0
		Min, Max	48, 63	46, 73	48, 68	49, 76		46, 76
	Change from baseline on Day 1	n	22	22	9	9	0	62
		Mean (SD)	-1.2 (6.39)	-1.2 (5.23)	-2.6 (7.02)	4.3 (7.42)		-0.6 (6.45)
		Median	-1.0	-1.0	-2.0	5.0		-1.0
		Min, Max	-13, 10	-10, 9	-14, 9	-6, 14		-14, 14

Note: mCC = menthol Conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

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**Table 15.2.6.13 Summary of Supine Vital Signs - Safety Population**

Parameter (units)	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Supine Diastolic Blood Pressure (mmHg)	Day 2	n	21	22	9	9	0	61
		Mean (SD)	53.7 (3.23)	54.8 (5.81)	55.3 (5.90)	55.2 (7.97)		54.6 (5.37)
		Median	54.0	53.0	53.0	52.0		53.0
		Min, Max	48, 60	47, 69	47, 66	45, 68		45, 69
	Change from baseline on Day 2	n	21	22	9	9	0	61
		Mean (SD)	-3.5 (5.38)	-3.1 (5.89)	-1.7 (6.54)	0.1 (7.05)		-2.5 (5.98)
		Median	-4.0	-4.0	-1.0	-1.0		-2.0
		Min, Max	-16, 6	-15, 8	-10, 7	-10, 10		-16, 10

Note: mCC = menthol Conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

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**Table 15.2.6.13 Summary of Supine Vital Signs - Safety Population**

Parameter (units)	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Supine Diastolic Blood Pressure (mmHg)	Day 3	n	21	22	9	9	0	61
		Mean (SD)	56.6 (5.49)	57.5 (7.48)	56.6 (7.18)	56.4 (7.20)		56.9 (6.61)
		Median	57.0	56.5	55.0	57.0		57.0
		Min, Max	48, 66	47, 78	46, 66	46, 71		46, 78
	Change from baseline on Day 3	n	21	22	9	9	0	61
		Mean (SD)	-0.6 (5.08)	-0.3 (6.12)	-0.4 (5.51)	1.3 (6.66)		-0.2 (5.66)
		Median	0.0	1.0	0.0	3.0		0.0
		Min, Max	-13, 11	-12, 12	-11, 8	-9, 9		-13, 12

Note: mCC = menthol Conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

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**Table 15.2.6.13 Summary of Supine Vital Signs - Safety Population**

Parameter (units)	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Supine Diastolic Blood Pressure (mmHg)	Day 4/ Discharge	n	22	22	9	9	11	73
		Mean (SD)	59.6 (6.07)	59.2 (7.36)	60.8 (5.68)	60.1 (7.19)	58.7 (5.41)	59.5 (6.35)
		Median	61.0	59.5	62.0	57.0	59.0	60.0
		Min, Max	44, 67	48, 74	50, 68	53, 76	50, 67	44, 76
	Change from baseline on Day 4/ Discharge	n	22	22	9	9	11	73
		Mean (SD)	2.2 (7.22)	1.3 (6.49)	3.8 (4.77)	5.0 (6.60)	1.8 (5.66)	2.4 (6.39)
		Median	4.0	1.5	5.0	4.0	3.0	3.0
		Min, Max	-11, 11	-10, 17	-4, 10	-5, 14	-12, 8	-12, 17

Note: mCC = menthol Conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

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**Table 15.2.6.13 Summary of Supine Vital Signs - Safety Population**

Parameter (units)	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Supine Pulse Rate (beats/min)	Baseline	n	22	22	9	9	11	73
		Mean (SD)	62.5 (7.15)	61.9 (8.77)	60.8 (5.92)	64.6 (7.56)	68.0 (11.66)	63.2 (8.46)
		Median	62.5	61.0	61.0	64.0	66.0	62.0
		Min, Max	46, 77	43, 82	51, 72	54, 77	54, 88	43, 88
	Day 0	n	22	22	9	9	5	67
		Mean (SD)	61.2 (7.89)	61.8 (6.79)	58.6 (5.58)	61.4 (7.25)	55.8 (2.95)	60.7 (6.94)
		Median	63.0	61.0	57.0	63.0	55.0	61.0
		Min, Max	45, 74	45, 77	51, 71	49, 72	52, 60	45, 77

Note: mCC = menthol Conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

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**Table 15.2.6.13 Summary of Supine Vital Signs - Safety Population**

Parameter (units)	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Supine Pulse Rate (beats/min)	Change from baseline on Day 0	n	22	22	9	9	5	67
		Mean (SD)	-1.3 (6.64)	-0.1 (4.94)	-2.2 (4.82)	-3.1 (7.51)	-4.6 (5.23)	-1.5 (5.91)
		Median	-1.0	0.0	-1.0	-4.0	-3.0	-1.0
		Min, Max	-15, 11	-9, 11	-11, 4	-15, 7	-13, 0	-15, 11
	Day 1	n	22	22	9	9	0	62
		Mean (SD)	59.9 (7.18)	58.9 (7.65)	56.0 (4.22)	66.3 (6.43)		59.9 (7.37)
		Median	60.5	58.5	55.0	66.0		60.0
		Min, Max	48, 71	45, 72	49, 62	58, 77		45, 77

Note: mCC = menthol Conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

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**Table 15.2.6.13 Summary of Supine Vital Signs - Safety Population**

Parameter (units)	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Supine Pulse Rate (beats/min)	Change from baseline on Day 1	n	22	22	9	9	0	62
		Mean (SD)	-2.6 (6.17)	-3.0 (5.85)	-4.8 (6.98)	1.8 (7.68)		-2.4 (6.53)
		Median	-1.5	-2.5	-6.0	0.0		-2.0
		Min, Max	-16, 8	-15, 6	-15, 6	-9, 15		-16, 15
	Day 2	n	21	22	9	9	0	61
		Mean (SD)	60.4 (7.20)	59.0 (7.35)	58.3 (3.54)	62.4 (8.12)		59.9 (6.96)
		Median	62.0	57.5	59.0	63.0		59.0
		Min, Max	47, 71	49, 76	54, 64	52, 76		47, 76

Note: mCC = menthol Conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

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**Table 15.2.6.13 Summary of Supine Vital Signs - Safety Population**

Parameter (units)	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Supine Pulse Rate (beats/min)	Change from baseline on Day 2	n	21	22	9	9	0	61
		Mean (SD)	-2.6 (5.49)	-2.9 (6.66)	-2.4 (5.25)	-2.1 (7.85)		-2.6 (6.13)
		Median	-2.0	-3.5	-2.0	-3.0		-3.0
		Min, Max	-11, 12	-17, 7	-11, 4	-12, 11		-17, 12
	Day 3	n	21	22	9	9	0	61
		Mean (SD)	59.2 (7.12)	59.5 (7.12)	56.0 (4.31)	60.4 (4.37)		59.0 (6.44)
		Median	60.0	58.5	57.0	60.0		59.0
		Min, Max	47, 70	49, 73	50, 64	53, 67		47, 73

Note: mCC = menthol Conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

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**Table 15.2.6.13 Summary of Supine Vital Signs - Safety Population**

Parameter (units)	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Supine Pulse Rate (beats/min)	Change from baseline on Day 3	n	21	22	9	9	0	61
		Mean (SD)	-3.8 (6.23)	-2.5 (6.73)	-4.8 (6.62)	-4.1 (5.69)		-3.5 (6.30)
		Median	-5.0	-2.5	-7.0	-4.0		-5.0
		Min, Max	-14, 11	-16, 11	-12, 6	-14, 7		-16, 11
	Day 4/ Discharge	n	22	22	9	9	11	73
		Mean (SD)	55.6 (5.70)	55.8 (7.83)	54.6 (7.25)	58.2 (9.08)	63.7 (8.36)	57.1 (7.81)
		Median	57.0	54.0	51.0	58.0	60.0	57.0
		Min, Max	44, 65	42, 72	48, 66	45, 73	52, 77	42, 77

Note: mCC = menthol Conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

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**Table 15.2.6.13 Summary of Supine Vital Signs - Safety Population**

Parameter (units)	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Supine Pulse Rate (beats/min)	Change from baseline on Day 4/ Discharge	n	22	22	9	9	11	73
		Mean (SD)	-6.9 (7.68)	-6.1 (6.06)	-6.2 (10.00)	-6.3 (8.74)	-4.3 (7.70)	-6.1 (7.52)
		Median	-6.0	-6.5	-11.0	-9.0	-4.0	-7.0
		Min, Max	-20, 6	-17, 12	-14, 15	-19, 6	-17, 6	-20, 15
Supine Respiratory Rate (breaths/ min)	Baseline	n	22	22	9	9	11	73
		Mean (SD)	15.5 (2.25)	16.5 (3.10)	15.6 (2.41)	16.4 (2.41)	16.0 (3.10)	16.0 (2.67)
		Median	16.0	16.0	16.0	16.0	16.0	16.0
		Min, Max	12, 20	12, 20	12, 20	12, 20	12, 20	12, 20

Note: mCC = menthol Conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Appendix 15.3.6.7

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_vit.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.6.13 Summary of Supine Vital Signs - Safety Population**

Parameter (units)	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Supine Respiratory Rate (breaths/ min)	Day 0	n	22	22	9	9	5	67
		Mean (SD)	16.4 (3.48)	15.8 (2.31)	16.4 (2.41)	16.0 (2.00)	13.6 (3.58)	15.9 (2.83)
		Median	16.0	16.0	16.0	16.0	12.0	16.0
		Min, Max	12, 20	12, 20	12, 20	12, 20	12, 20	12, 20
	Change from baseline on Day 0	n	22	22	9	9	5	67
		Mean (SD)	0.9 (3.26)	-0.7 (2.66)	0.9 (3.34)	-0.4 (1.34)	-1.6 (2.20)	0.0 (2.88)
		Median	0.0	0.0	0.0	0.0	0.0	0.0
		Min, Max	-4, 4	-4, 4	-4, 4	-4, 0	-4, 0	-4, 4

Note: mCC = menthol Conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Appendix 15.3.6.7

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_vit.sas

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**Table 15.2.6.13 Summary of Supine Vital Signs - Safety Population**

Parameter (units)	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Supine Respiratory Rate (breaths/ min)	Day 1	n	22	22	9	9	0	62
		Mean (SD)	16.2 (2.62)	16.5 (2.85)	16.4 (3.13)	15.6 (2.41)		16.3 (2.70)
		Median	16.0	16.0	16.0	16.0		16.0
		Min, Max	12, 20	12, 20	12, 20	12, 20		12, 20
	Change from baseline on Day 1	n	22	22	9	9	0	62
		Mean (SD)	0.7 (3.19)	0.0 (3.03)	0.9 (4.38)	-0.9 (2.67)		0.3 (3.23)
		Median	0.0	0.0	0.0	0.0		0.0
		Min, Max	-4, 4	-8, 4	-4, 8	-4, 4		-8, 8

Note: mCC = menthol Conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Appendix 15.3.6.7

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_vit.sas

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**Table 15.2.6.13 Summary of Supine Vital Signs - Safety Population**

Parameter (units)	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Supine Respiratory Rate (breaths/ min)	Day 2	n	21	22	9	9	0	61
		Mean (SD)	15.6 (2.16)	16.0 (2.77)	16.0 (2.00)	15.1 (1.77)		15.7 (2.30)
		Median	16.0	16.0	16.0	16.0		16.0
		Min, Max	12, 20	12, 20	12, 20	12, 16		12, 20
	Change from baseline on Day 2	n	21	22	9	9	0	61
		Mean (SD)	0.0 (2.83)	-0.5 (3.34)	0.4 (3.13)	-1.3 (2.83)		-0.3 (3.04)
		Median	0.0	0.0	0.0	0.0		0.0
		Min, Max	-4, 4	-8, 4	-4, 8	-4, 4		-8, 8

Note: mCC = menthol Conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Appendix 15.3.6.7

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_vit.sas

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**Table 15.2.6.13 Summary of Supine Vital Signs - Safety Population**

Parameter (units)	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Supine Respiratory Rate (breaths/ min)	Day 3	n	21	22	9	9	0	61
		Mean (SD)	15.8 (2.68)	16.0 (2.47)	15.6 (1.34)	14.2 (2.11)		15.6 (2.39)
		Median	16.0	16.0	16.0	16.0		16.0
		Min, Max	12, 20	12, 20	12, 16	12, 16		12, 20
	Change from baseline on Day 3	n	21	22	9	9	0	61
		Mean (SD)	0.2 (3.22)	-0.5 (3.34)	0.0 (2.00)	-2.2 (2.11)		-0.5 (3.02)
		Median	0.0	0.0	0.0	-4.0		0.0
		Min, Max	-4, 4	-4, 8	-4, 4	-4, 0		-4, 8

Note: mCC = menthol Conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Appendix 15.3.6.7

Path: /cvm/projects/prj/development/000000106326/dev/tables/t\_vit.sas

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**Table 15.2.6.13 Summary of Supine Vital Signs - Safety Population**

Parameter (units)	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Supine Respiratory Rate (breaths/ min)	Day 4/ Discharge	n	22	22	9	9	11	73
		Mean (SD)	16.0 (2.14)	15.1 (2.45)	15.6 (3.13)	16.0 (2.00)	15.6 (2.81)	15.6 (2.42)
		Median	16.0	16.0	16.0	16.0	16.0	16.0
		Min, Max	12, 20	12, 20	12, 20	12, 20	12, 20	12, 20
	Change from baseline on Day 4/ Discharge	n	22	22	9	9	11	73
		Mean (SD)	0.5 (2.85)	-1.5 (2.33)	0.0 (4.00)	-0.4 (1.34)	-0.4 (2.16)	-0.4 (2.69)
		Median	0.0	0.0	0.0	0.0	0.0	0.0
		Min, Max	-4, 4	-4, 4	-4, 8	-4, 0	-4, 4	-4, 8

Note: mCC = menthol Conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Appendix 15.3.6.7

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_vit.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.6.14 Summary of ECG Measurements - Safety Population**

Parameter (units)	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Heart rate (beats/min)	Baseline	n	22	22	9	9	11	73
		Mean (SD)	62.6 (5.65)	63.0 (8.41)	59.8 (8.85)	63.8 (7.89)	62.0 (8.62)	62.4 (7.56)
		Median	61.0	64.0	56.0	61.0	62.0	61.0
		Min, Max	53, 74	47, 79	49, 74	55, 79	50, 76	47, 79
	Day 1	n	22	22	9	9	0	62
		Mean (SD)	59.1 (8.47)	58.1 (7.66)	55.8 (5.45)	64.3 (8.25)		59.0 (7.99)
		Median	60.0	57.0	55.0	68.0		58.0
		Min, Max	42, 74	44, 74	49, 64	54, 75		42, 75

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Appendix 15.3.6.9

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_ecg.sas

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**Table 15.2.6.14 Summary of ECG Measurements - Safety Population**

Parameter (units)	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Heart rate (beats/min)	Change from Baseline on Day 1	n	22	22	9	9	0	62
		Mean (SD)	-3.5 (7.36)	-4.9 (9.28)	-4.0 (6.52)	0.6 (6.72)		-3.5 (7.93)
		Median	-4.5	-3.0	-6.0	0.0		-3.0
		Min, Max	-15, 14	-22, 8	-13, 7	-10, 12		-22, 14
	Day 3	n	21	22	9	9	0	61
		Mean (SD)	59.3 (8.04)	57.3 (6.92)	58.0 (4.95)	59.6 (6.43)		58.4 (6.93)
		Median	59.0	57.0	56.0	59.0		58.0
		Min, Max	45, 74	46, 71	52, 67	50, 69		45, 74

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Appendix 15.3.6.9

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_ecg.sas

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**Table 15.2.6.14 Summary of ECG Measurements - Safety Population**

Parameter (units)	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Heart rate (beats/min)	Change from Baseline on Day 3	n	21	22	9	9	0	61
		Mean (SD)	-3.8 (6.94)	-5.6 (8.09)	-1.8 (8.11)	-4.2 (7.76)		-4.2 (7.58)
		Median	-1.0	-2.0	0.0	-7.0		-2.0
		Min, Max	-16, 8	-24, 5	-18, 9	-13, 10		-24, 10
PR duration (msec)	Baseline	n	22	22	9	9	11	73
		Mean (SD)	152.5 (22.98)	153.7 (17.10)	156.9 (28.01)	148.0 (24.98)	160.7 (18.51)	154.1 (21.37)
		Median	148.0	153.0	148.0	142.0	162.0	154.0
		Min, Max	120, 194	126, 192	124, 202	118, 184	136, 198	118, 202

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Appendix 15.3.6.9

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**Table 15.2.6.14 Summary of ECG Measurements - Safety Population**

Parameter (units)	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
PR duration (msec)	Day 1	n	22	22	9	9	0	62
		Mean (SD)	158.3 (20.77)	158.9 (18.45)	162.0 (26.76)	148.7 (24.96)		157.6 (21.37)
		Median	156.0	157.0	164.0	162.0		156.0
		Min, Max	126, 212	128, 206	130, 204	112, 184		112, 212
	Change from Baseline on Day 1	n	22	22	9	9	0	62
		Mean (SD)	5.7 (18.45)	5.2 (11.94)	5.1 (6.65)	0.7 (11.36)		4.7 (13.85)
		Median	3.0	6.0	6.0	4.0		4.0
		Min, Max	-38, 56	-22, 28	-4, 18	-20, 20		-38, 56

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Appendix 15.3.6.9

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**Table 15.2.6.14 Summary of ECG Measurements - Safety Population**

Parameter (units)	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
PR duration (msec)	Day 3	n	21	22	9	9	0	61
		Mean (SD)	153.9 (23.03)	157.5 (18.40)	162.9 (20.55)	153.1 (24.64)		156.4 (21.05)
		Median	150.0	155.0	164.0	164.0		156.0
		Min, Max	122, 204	130, 202	136, 206	118, 188		118, 206
	Change from Baseline on Day 3	n	21	22	9	9	0	61
		Mean (SD)	1.8 (19.53)	3.8 (11.33)	6.0 (15.75)	5.1 (10.06)		3.6 (14.87)
		Median	0.0	3.0	8.0	6.0		6.0
		Min, Max	-46, 60	-16, 20	-32, 22	-14, 22		-46, 60

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Appendix 15.3.6.9

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**Table 15.2.6.14 Summary of ECG Measurements - Safety Population**

Parameter (units)	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
QT duration (msec)	Baseline	n	22	22	9	9	11	73
		Mean (SD)	402.9 (18.82)	402.7 (32.22)	410.9 (28.99)	398.7 (21.80)	407.5 (27.75)	404.0 (25.91)
		Median	398.0	392.0	410.0	404.0	410.0	400.0
		Min, Max	380, 446	358, 478	362, 460	352, 426	372, 456	352, 478
	Day 1	n	22	22	9	9	0	62
		Mean (SD)	408.2 (28.45)	409.9 (29.40)	407.3 (19.73)	388.7 (17.98)		405.8 (26.86)
		Median	406.0	404.0	404.0	388.0		399.0
		Min, Max	360, 492	368, 478	384, 436	366, 420		360, 492

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

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**Table 15.2.6.14 Summary of ECG Measurements - Safety Population**

Parameter (units)	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
QT duration (msec)	Change from Baseline on Day 1	n	22	22	9	9	0	62
		Mean (SD)	5.3 (24.75)	7.2 (32.18)	-3.6 (19.44)	-10.0 (23.90)		2.5 (27.04)
		Median	5.0	11.0	-2.0	-8.0		0.0
		Min, Max	-52, 52	-52, 86	-28, 26	-48, 28		-52, 86
	Day 3	n	21	22	9	9	0	61
		Mean (SD)	404.3 (23.31)	408.7 (24.38)	403.6 (17.49)	401.1 (23.52)		405.3 (22.61)
		Median	410.0	408.0	406.0	402.0		406.0
		Min, Max	358, 446	368, 456	376, 432	362, 436		358, 456

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

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**Table 15.2.6.14 Summary of ECG Measurements - Safety Population**

Parameter (units)	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
QT duration (msec)	Change from Baseline on Day 3	n	21	22	9	9	0	61
		Mean (SD)	2.6 (22.46)	6.0 (23.15)	-7.3 (16.59)	2.4 (28.34)		2.3 (22.77)
		Median	0.0	1.0	-10.0	16.0		0.0
		Min, Max	-36, 42	-38, 58	-28, 20	-44, 30		-44, 58
QTcB - Bazett's correction formula (msec)	Baseline	n	22	22	9	9	11	73
		Mean (SD)	412.0 (16.14)	411.2 (18.88)	408.7 (16.85)	410.6 (18.31)	413.2 (14.94)	411.4 (16.77)
		Median	410.0	411.5	408.0	407.0	416.0	411.0
		Min, Max	381, 442	369, 444	382, 432	389, 448	379, 432	369, 448

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

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**Table 15.2.6.14 Summary of ECG Measurements - Safety Population**

Parameter (units)	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
QTcB - Bazett's correction formula (msec)	Day 1	n	22	22	9	9	0	62
		Mean (SD)	403.9 (20.45)	402.1 (19.62)	393.2 (9.55)	402.6 (19.12)		401.5 (18.68)
		Median	402.0	400.0	393.0	408.0		401.0
		Min, Max	366, 437	375, 448	377, 406	371, 423		366, 448
	Change from Baseline on Day 1	n	22	22	9	9	0	62
		Mean (SD)	-8.1 (12.90)	-9.1 (20.35)	-15.4 (14.08)	-8.0 (14.96)		-9.5 (16.17)
		Median	-7.5	-9.0	-15.0	-9.0		-8.5
		Min, Max	-45, 10	-44, 30	-34, 1	-37, 8		-45, 30

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Appendix 15.3.6.9

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_ecg.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.6.14 Summary of ECG Measurements - Safety Population**

Parameter (units)	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
QTcB - Bazett's correction formula (msec)	Day 3	n	21	22	9	9	0	61
		Mean (SD)	401.2 (19.93)	398.9 (18.62)	398.1 (20.61)	399.6 (21.64)		399.7 (19.35)
		Median	400.0	404.0	403.0	399.0		400.0
		Min, Max	367, 428	371, 435	361, 424	360, 435		360, 435
		Change from Baseline on Day 3						
		n	21	22	9	9	0	61
		Mean (SD)	-11.2 (13.35)	-12.4 (18.76)	-10.6 (21.90)	-11.0 (16.12)		-11.5 (16.78)
		Median	-8.0	-7.0	-17.0	-9.0		-9.0
		Min, Max	-43, 9	-56, 10	-45, 30	-34, 20		-56, 30

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Appendix 15.3.6.9

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_ecg.sas

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**Table 15.2.6.14 Summary of ECG Measurements - Safety Population**

Parameter (units)	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
QTcF - Fridericia's correction formula (msec)	Baseline	n	22	22	9	9	11	73
		Mean (SD)	408.0 (14.50)	407.3 (20.32)	408.5 (16.88)	405.7 (16.11)	410.2 (15.79)	407.9 (16.69)
		Median	408.1	406.3	408.1	404.0	403.0	408.0
		Min, Max	383, 438	364, 452	385, 436	386, 436	388, 438	364, 452
	Day 1	n	22	22	9	9	0	62
		Mean (SD)	404.2 (17.99)	403.8 (19.40)	396.7 (10.03)	396.7 (14.68)		401.9 (17.14)
		Median	406.6	401.4	397.2	395.6		401.5
		Min, Max	364, 437	372, 446	381, 410	375, 413		364, 446

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Appendix 15.3.6.9

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_ecg.sas

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**Table 15.2.6.14 Summary of ECG Measurements - Safety Population**

Parameter (units)	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
QTcF - Fridericia's correction formula (msec)	Change from Baseline on Day 1	n	22	22	9	9	0	62
		Mean (SD)	-3.8 (12.50)	-3.5 (20.09)	-11.8 (12.23)	-9.0 (16.30)		-5.6 (16.04)
		Median	-1.5	-1.5	-11.4	-8.5		-4.2
		Min, Max	-32, 16	-42, 49	-32, 9	-42, 14		-42, 49
	Day 3	n	21	22	9	9	0	61
		Mean (SD)	401.2 (16.95)	401.3 (17.51)	398.6 (18.22)	399.2 (19.71)		400.6 (17.33)
		Median	404.2	401.2	396.8	404.4		404.0
		Min, Max	370, 430	372, 438	365, 421	360, 429		360, 438

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Appendix 15.3.6.9

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_ecg.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.6.14 Summary of ECG Measurements - Safety Population**

Parameter (units)	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
QTcF - Fridericia's correction formula (msec)	Change from Baseline on Day 3	n	21	22	9	9	0	61
		Mean (SD)	-6.7 (12.42)	-6.0 (15.96)	-9.8 (16.81)	-6.5 (17.85)		-6.9 (14.91)
		Median	-5.5	-2.4	-15.9	-2.6		-5.5
		Min, Max	-33, 19	-49, 18	-30, 22	-34, 19		-49, 22
QRS duration (msec)	Baseline	n	22	22	9	9	11	73
		Mean (SD)	92.1 (10.06)	93.5 (8.29)	96.2 (11.03)	92.0 (9.06)	93.6 (12.74)	93.2 (9.82)
		Median	93.0	93.0	94.0	90.0	94.0	94.0
		Min, Max	74, 114	82, 110	82, 120	80, 104	80, 126	74, 126

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Appendix 15.3.6.9

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_ecg.sas

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**Table 15.2.6.14 Summary of ECG Measurements - Safety Population**

Parameter (units)	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
QRS duration (msec)	Day 1	n	22	22	9	9	0	62
		Mean (SD)	92.5 (10.63)	96.9 (9.72)	99.6 (12.20)	93.1 (9.12)		95.2 (10.43)
		Median	91.0	92.0	96.0	92.0		92.0
		Min, Max	74, 116	84, 114	88, 128	80, 104		74, 128
	Change from Baseline on Day 1	n	22	22	9	9	0	62
		Mean (SD)	0.5 (3.86)	3.5 (4.87)	3.3 (4.48)	1.1 (1.77)		2.0 (4.28)
		Median	0.0	2.0	2.0	2.0		2.0
		Min, Max	-6, 8	-4, 16	-2, 12	-2, 4		-6, 16

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Appendix 15.3.6.9

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**Table 15.2.6.14 Summary of ECG Measurements - Safety Population**

Parameter (units)	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
QRS duration (msec)	Day 3	n	21	22	9	9	0	61
		Mean (SD)	92.9 (9.86)	96.9 (9.95)	97.8 (10.84)	93.3 (10.91)		95.1 (10.15)
		Median	94.0	94.0	96.0	90.0		94.0
		Min, Max	76, 114	84, 116	86, 122	82, 108		76, 122
	Change from Baseline on Day 3	n	21	22	9	9	0	61
		Mean (SD)	1.0 (3.01)	3.5 (4.67)	1.6 (3.29)	1.3 (3.47)		2.0 (3.87)
		Median	0.0	2.0	2.0	0.0		2.0
		Min, Max	-4, 8	-2, 14	-4, 8	-4, 6		-4, 14

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Appendix 15.3.6.9

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**Table 15.2.6.14 Summary of ECG Measurements - Safety Population**

Parameter (units)	Study Day	Statistic	Sequence										Overall Safety (N=73)	
			THS 2.2 Menthol - mCC (N=22)		mCC - THS 2.2 Menthol (N=22)		THS 2.2 Menthol- NRT gum (N=9)		NRT gum - THS 2.2 Menthol (N=9)		Enrolled Not Randomized (N=11)			
Interpretation	Baseline	Normal - n (%)	22	(100 %)	22	(100 %)	9	(100 %)	9	(100 %)	11	(100 %)	73	(100 %)
		Abnormal non- clinically significant - n (%)	0		0		0		0		0		0	
		Abnormal clinically significant - n (%)	0		0		0		0		0		0	
	A Single Use Day 1	Normal - n (%)	22	(100 %)	22	(100 %)	9	(100 %)	9	(100 %)	0		62	( 84.9%)
		Abnormal non- clinically significant - n (%)	0		0		0		0		0		0	
		Abnormal clinically significant - n (%)	0		0		0		0		0		0	

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

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Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.6.14 Summary of ECG Measurements - Safety Population**

Parameter (units)	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol- NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Interpretation 3	Single Use Day	Normal - n (%)	21 (95.5%)	22 (100 %)	9 (100 %)	9 (100 %)	0	61 (83.6%)
		Abnormal non- clinically significant - n (%)	0	0	0	0	0	0
		Abnormal clinically significant - n (%)	0	0	0	0	0	0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Appendix 15.3.6.9

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_ecg.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.6.15 Summary of Spirometry Measurements - Safety Population**

Parameter (units)	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Best measured FVC value (with bronchodilator) (L)	Baseline	n	22	22	9	9	11	73
		Mean (SD)	4.175 (0.7271)	4.271 (0.7581)	4.390 (1.0946)	4.087 (0.8458)	3.677 (0.7948)	4.144 (0.8177)
		Median	4.145	4.290	4.180	3.760	3.550	4.150
		Min, Max	2.82, 5.49	3.10, 5.53	3.06, 6.05	3.00, 5.66	2.74, 4.91	2.74, 6.05
Percent of predicted FVC value (with bronchodilator) (%)	Baseline	n	22	22	9	9	11	73
		Mean (SD)	107.82 (12.809)	106.86 (13.432)	106.67 (6.083)	107.00 (13.172)	98.82 (10.039)	105.93 (12.131)
		Median	104.50	104.00	106.00	103.00	98.00	103.00
		Min, Max	88.0, 136.0	82.0, 131.0	97.0, 116.0	93.0, 123.0	84.0, 115.0	82.0, 136.0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

**Table 15.2.6.15 Summary of Spirometry Measurements - Safety Population**

Parameter (units)	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Best measured FEV1 value (with bronchodilator) (L)	Baseline	n	22	22	9	9	11	73
		Mean (SD)	3.439 (0.6289)	3.559 (0.6393)	3.832 (0.9549)	3.519 (0.8233)	3.173 (0.7611)	3.493 (0.7239)
		Median	3.435	3.745	3.490	3.100	3.210	3.380
		Min, Max	2.41, 4.55	2.40, 4.91	2.79, 5.25	2.48, 4.76	2.24, 4.47	2.24, 5.25
Percent of predicted FEV1 value (with bronchodilator) (%)	Baseline	n	22	22	9	9	11	73
		Mean (SD)	102.45 (10.765)	101.41 (11.057)	106.22 (11.915)	104.89 (11.208)	97.55 (9.533)	102.16 (10.875)
		Median	100.00	99.50	108.00	109.00	96.00	101.00
		Min, Max	85.0, 126.0	82.0, 124.0	81.0, 126.0	89.0, 118.0	83.0, 118.0	81.0, 126.0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

**Table 15.2.6.15 Summary of Spirometry Measurements - Safety Population**

Parameter (units)	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Calculated ratio between FEV1/FVC (with bronchodilator) (RATIO)	Baseline	n	22	22	9	9	11	73
		Mean (SD)	0.824 (0.0565)	0.836 (0.0719)	0.877 (0.0775)	0.859 (0.0516)	0.861 (0.0733)	0.844 (0.0670)
		Median	0.810	0.840	0.870	0.840	0.880	0.840
		Min, Max	0.73, 0.95	0.72, 0.99	0.72, 0.96	0.79, 0.95	0.74, 0.97	0.72, 0.99
Best measured FVC value (L)	Baseline	n	22	22	9	9	11	73
		Mean (SD)	4.077 (0.7008)	4.270 (0.7019)	4.378 (1.1155)	4.024 (0.9487)	3.626 (0.7873)	4.098 (0.8154)
		Median	4.035	4.530	3.960	3.840	3.620	3.980
		Min, Max	2.90, 5.49	3.14, 5.51	3.10, 6.19	2.91, 5.73	2.67, 4.82	2.67, 6.19

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Appendix 15.3.6.8

Path: /cvm/projects/prj/development/000000106326/dev/tables/t\_spiro.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.6.15 Summary of Spirometry Measurements - Safety Population**

Parameter (units)	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Best measured FVC value (L)	Day 4/ Discharge	n	22	22	9	9	11	73
		Mean (SD)	4.067 (0.6950)	4.292 (0.7153)	4.377 (1.0543)	4.066 (0.8071)	3.585 (0.7613)	4.100 (0.7927)
		Median	4.070	4.420	4.200	3.800	3.670	4.050
		Min, Max	2.71, 5.18	3.18, 5.63	3.12, 6.12	3.05, 5.38	2.69, 4.68	2.69, 6.12
	Change from Baseline	n	22	22	9	9	11	73
		Mean (SD)	-0.010 (0.1318)	0.022 (0.1304)	-0.001 (0.1446)	0.041 (0.2080)	-0.042 (0.1689)	0.002 (0.1476)
		Median	0.005	0.040	-0.060	0.030	-0.070	0.020
		Min, Max	-0.31, 0.20	-0.28, 0.25	-0.18, 0.24	-0.35, 0.43	-0.33, 0.25	-0.35, 0.43

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Appendix 15.3.6.8

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_spiro.sas

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**Table 15.2.6.15 Summary of Spirometry Measurements - Safety Population**

Parameter (units)	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Percent of predicted FVC value (%)	Baseline	n	22	22	9	9	11	73
		Mean (SD)	105.83 (12.991)	107.33 (12.585)	106.69 (7.706)	105.63 (15.659)	97.96 (10.184)	105.18 (12.389)
		Median	102.85	106.55	106.80	107.70	98.90	104.40
		Min, Max	85.1, 133.6	88.3, 135.8	92.0, 115.8	84.9, 124.5	83.2, 117.1	83.2, 135.8
	Day 4/ Discharge	n	22	22	9	9	11	73
		Mean (SD)	105.82 (15.173)	107.99 (13.774)	106.83 (5.384)	106.99 (12.157)	96.92 (10.377)	105.40 (13.067)
		Median	102.55	106.90	106.40	106.20	95.30	105.00
		Min, Max	79.4, 135.9	85.5, 136.2	97.6, 113.7	87.8, 123.3	84.3, 119.6	79.4, 136.2

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Appendix 15.3.6.8

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_spiro.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.6.15 Summary of Spirometry Measurements - Safety Population**

Parameter (units)	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Percent of predicted FVC value (%)	Change from Baseline	n	22	22	9	9	11	73
		Mean (SD)	-0.01 (3.433)	0.66 (3.001)	0.14 (3.601)	1.36 (5.280)	-1.05 (4.316)	0.22 (3.698)
		Median	0.15	1.10	-1.30	1.00	-2.50	0.70
		Min, Max	-6.9, 6.6	-5.6, 5.7	-3.7, 6.6	-7.6, 12.0	-7.5, 5.0	-7.6, 12.0
Best measured FEV1 value (L)	Baseline	n	22	22	9	9	11	73
		Mean (SD)	3.290 (0.5885)	3.560 (0.6190)	3.596 (0.7964)	3.413 (0.8427)	3.045 (0.7373)	3.387 (0.6877)
		Median	3.275	3.755	3.490	2.950	3.050	3.300
		Min, Max	2.29, 4.19	2.65, 4.82	2.54, 4.46	2.46, 4.64	2.10, 4.26	2.10, 4.82

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Appendix 15.3.6.8

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_spiro.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.6.15 Summary of Spirometry Measurements - Safety Population**

Parameter (units)	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Best measured FEV1 value (L)	Day 4/ Discharge	n	22	22	9	9	11	73
		Mean (SD)	3.334 (0.6021)	3.588 (0.5903)	3.771 (0.9378)	3.499 (0.7972)	2.972 (0.7254)	3.430 (0.7117)
		Median	3.295	3.735	3.390	3.020	2.630	3.390
		Min, Max	2.39, 4.32	2.56, 4.82	2.63, 5.32	2.61, 4.66	2.12, 4.18	2.12, 5.32
	Change from Baseline	n	22	22	9	9	11	73
		Mean (SD)	0.044 (0.1327)	0.029 (0.1342)	0.176 (0.2977)	0.086 (0.0887)	-0.074 (0.2250)	0.043 (0.1809)
		Median	0.075	0.035	0.090	0.100	-0.010	0.060
		Min, Max	-0.27, 0.27	-0.22, 0.23	-0.10, 0.91	-0.05, 0.18	-0.66, 0.15	-0.66, 0.91

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Appendix 15.3.6.8

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_spiro.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.6.15 Summary of Spirometry Measurements - Safety Population**

Parameter (units)	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Percent of predicted FEV1 value (%)	Baseline	n	22	22	9	9	11	73
		Mean (SD)	98.47 (10.599)	102.00 (10.210)	100.74 (14.367)	101.77 (9.577)	94.00 (10.048)	99.55 (10.863)
		Median	97.75	101.30	101.10	103.70	92.70	99.30
		Min, Max	81.4, 117.5	84.3, 121.4	77.0, 126.9	88.8, 114.8	79.4, 117.3	77.0, 126.9
	Day 4/ Discharge	n	22	22	9	9	11	73
		Mean (SD)	99.88 (11.555)	103.04 (11.198)	104.79 (11.500)	104.70 (9.998)	91.53 (5.602)	100.77 (11.158)
		Median	97.20	103.10	102.70	108.30	93.00	98.20
		Min, Max	82.8, 124.5	86.0, 128.5	84.0, 123.2	93.8, 120.8	79.6, 98.6	79.6, 128.5

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Appendix 15.3.6.8

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_spiro.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.6.15 Summary of Spirometry Measurements - Safety Population**

Parameter (units)	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Percent of predicted FEV1 value (%)	Change from Baseline	n	22	22	9	9	11	73
		Mean (SD)	1.41 (4.259)	1.04 (3.807)	4.04 (6.372)	2.93 (3.191)	-2.47 (8.107)	1.23 (5.269)
		Median	2.00	1.15	3.40	3.00	-0.20	1.80
		Min, Max	-8.8, 8.6	-4.9, 7.7	-3.7, 18.7	-1.3, 7.2	-23.8, 6.3	-23.8, 18.7
Calculated ratio between FEV1/FVC (RATIO)	Baseline	n	22	22	9	9	11	73
		Mean (SD)	0.812 (0.0680)	0.838 (0.0699)	0.863 (0.0799)	0.829 (0.0582)	0.862 (0.0623)	0.836 (0.0691)
		Median	0.800	0.835	0.860	0.820	0.890	0.830
		Min, Max	0.69, 1.00	0.72, 0.95	0.70, 0.95	0.75, 0.91	0.76, 0.95	0.69, 1.00

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Appendix 15.3.6.8

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Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.6.15 Summary of Spirometry Measurements - Safety Population**

Parameter (units)	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Calculated ratio between FEV1/FVC <sup>1</sup> (RATIO)	Baseline	n	22	22	9	9	11	73
		Mean (SD)	0.808 (0.0580)	0.834 (0.0480)	0.829 (0.0839)	0.849 (0.0729)	0.836 (0.0453)	0.828 (0.0593)
		Median	0.800	0.825	0.820	0.850	0.840	0.820
		Min, Max	0.71, 0.95	0.74, 0.94	0.71, 0.94	0.73, 0.96	0.75, 0.88	0.71, 0.96
	Day 4/ Discharge	n	22	22	9	9	11	73
		Mean (SD)	0.820 (0.0606)	0.839 (0.0567)	0.863 (0.0747)	0.857 (0.0560)	0.827 (0.0614)	0.837 (0.0611)
		Median	0.815	0.825	0.870	0.840	0.830	0.830
		Min, Max	0.70, 0.99	0.74, 0.98	0.74, 0.96	0.78, 0.96	0.68, 0.89	0.68, 0.99

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Note: 1: Ratio is derived in analysis datasets.

**Table 15.2.6.15 Summary of Spirometry Measurements - Safety Population**

Parameter (units)	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Calculated ratio between FEV1/FVC <sup>1</sup> (RATIO)	Change from Baseline	n	22	22	9	9	11	73
		Mean (SD)	0.013 (0.0337)	0.005 (0.0276)	0.034 (0.0525)	0.008 (0.0515)	-0.009 (0.0635)	0.009 (0.0431)
		Median	0.015	0.005	0.020	0.010	0.010	0.010
		Min, Max	-0.06, 0.07	-0.05, 0.06	-0.02, 0.16	-0.12, 0.05	-0.19, 0.04	-0.19, 0.16

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Note: 1: Ratio is derived in analysis datasets.

Appendix 15.3.6.8

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**Table 15.2.6.15 Summary of Spirometry Measurements - Safety Population**

Parameter (units)	Study Day	Statistic	Sequence					Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)	Enrolled Not Randomized (N=11)	
Interpretation (with bronchodilator)	Baseline	Normal - n (%)	22 (100 %)	22 (100 %)	9 (100 %)	9 (100 %)	11 (100 %)	73 (100 %)
		Abnormal non- clinically significant - n (%)	0	0	0	0	0	0
		Abnormal clinically significant - n (%)	0	0	0	0	0	0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Appendix 15.3.6.8

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Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.6.15 Summary of Spirometry Measurements - Safety Population**

Parameter (units)	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Interpretation	Baseline	Normal - n (%)	22 (100 %)	22 (100 %)	9 (100 %)	9 (100 %)	11 (100 %)	73 (100 %)
		Abnormal non- clinically significant - n (%)	0	0	0	0	0	0
		Abnormal clinically significant - n (%)	0	0	0	0	0	0
	Day 4/ Discharge	Normal - n (%)	22 (100 %)	22 (100 %)	9 (100 %)	9 (100 %)	11 (100 %)	73 (100 %)
		Abnormal non- clinically significant - n (%)	0	0	0	0	0	0
		Abnormal clinically significant - n (%)	0	0	0	0	0	0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Appendix 15.3.6.8

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_spiro.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.6.16 Summary of Weight and BMI Measurements - Safety Population**

Parameter (units)	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Weight (kg)	Baseline	n	22	22	9	9	11	73
		Mean (SD)	61.85 (6.450)	62.08 (9.310)	66.11 (11.702)	59.07 (9.221)	57.50 (8.329)	61.44 (8.814)
		Median	60.95	63.35	67.50	55.10	53.80	60.80
		Min, Max	49.4, 77.5	46.0, 78.7	51.9, 82.2	48.2, 75.0	47.4, 75.5	46.0, 82.2
	Day 4/ Discharge	n	22	22	9	9	11	73
		Mean (SD)	61.31 (6.388)	61.50 (9.041)	65.71 (11.474)	58.46 (8.734)	57.21 (8.125)	60.94 (8.596)
		Median	60.75	62.50	68.60	54.30	53.80	60.60
		Min, Max	49.5, 76.5	45.9, 77.6	51.9, 81.8	48.3, 73.3	47.4, 75.0	45.9, 81.8

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Note: 1: BMI is derived in analysis datasets.

Appendix 15.3.6.10

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_bmi.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.6.16 Summary of Weight and BMI Measurements - Safety Population**

Parameter (units)	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
BMI <sup>1</sup> (kg/m <sup>2</sup> )	Baseline	n	22	22	9	9	11	73
		Mean (SD)	22.21 (1.411)	22.46 (2.968)	23.84 (2.582)	21.88 (2.117)	21.77 (2.326)	22.38 (2.347)
		Median	22.15	21.50	23.40	21.90	21.20	22.20
		Min, Max	19.3, 24.5	18.7, 30.4	19.1, 28.4	18.8, 26.0	19.5, 26.7	18.7, 30.4
	Day 4/ Discharge	n	22	22	9	9	11	73
		Mean (SD)	22.02 (1.376)	22.25 (2.921)	23.71 (2.535)	21.66 (1.966)	21.68 (2.319)	22.20 (2.304)
		Median	21.95	21.50	23.20	21.60	21.10	22.00
		Min, Max	19.3, 24.2	18.6, 29.9	19.1, 28.3	18.7, 25.4	19.3, 26.6	18.6, 29.9

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Note: 1: BMI is derived in analysis datasets.

Appendix 15.3.6.10

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_bmi.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.6.16 Summary of Weight and BMI Measurements - Safety Population**

Parameter (units)	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
BMI (kg/m <sup>2</sup> )	Baseline	Underweight - n (%)	0	0	0	0	0	0
		Normal weight - n (%)	22 (100 %)	18 (81.8%)	7 (77.8%)	8 (88.9%)	10 (90.9%)	65 (89.0%)
		Overweight - n (%)	0	3 (13.6%)	2 (22.2%)	1 (11.1%)	1 (9.1%)	7 (9.6%)
		Obese - n (%)	0	1 (4.5%)	0	0	0	1 (1.4%)
	Day 4/ Discharge	Underweight - n (%)	0	0	0	0	0	0
		Normal weight - n (%)	22 (100 %)	18 (81.8%)	7 (77.8%)	8 (88.9%)	10 (90.9%)	65 (89.0%)
		Overweight - n (%)	0	4 (18.2%)	2 (22.2%)	1 (11.1%)	1 (9.1%)	8 (11.0%)
		Obese - n (%)	0	0	0	0	0	0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Note: 1: BMI is derived in analysis datasets.

Appendix 15.3.6.10

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_bmi.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.6.16 Summary of Weight and BMI Measurements - Safety Population**

Parameter (units)	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Height (m)	Screening	n	22	22	9	9	11	73
		Mean (SD)	1.668 (0.0773)	1.661 (0.0715)	1.661 (0.1129)	1.640 (0.0764)	1.624 (0.0870)	1.655 (0.0812)
		Median	1.655	1.675	1.670	1.650	1.590	1.650
		Min, Max	1.54, 1.80	1.53, 1.76	1.50, 1.86	1.54, 1.76	1.50, 1.76	1.50, 1.86

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Note: 1: BMI is derived in analysis datasets.

Appendix 15.3.6.10

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_bmi.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.6.17 Summary of Physical Examination of Body Systems - Safety Population**

Body System	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)		Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)			
General appearance	Baseline	n	22	22	9	9	11		73
		Normal - n (%)	22 (100 %)	22 (100 %)	9 (100 %)	9 (100 %)	11 (100 %)		73 (100 %)
		Abnormal non-clinically significant - n (%)	0	0	0	0	0		0
		Abnormal clinically significant - n (%)	0	0	0	0	0		0
	Day 4/ Discharge	n	22	22	9	9	11		73
		Normal - n (%)	22 (100 %)	22 (100 %)	9 (100 %)	9 (100 %)	11 (100 %)		73 (100 %)
		Abnormal non-clinically significant - n (%)	0	0	0	0	0		0
		Abnormal clinically significant - n (%)	0	0	0	0	0		0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Appendix 15.3.6.10

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_phyex.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.6.17 Summary of Physical Examination of Body Systems - Safety Population**

Body System	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)		Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)			
Head, eyes, ears, nose, throat	Baseline	n	22	22	9	9	11		73
		Normal - n (%)	22 (100 %)	22 (100 %)	9 (100 %)	9 (100 %)	11 (100 %)		73 (100 %)
		Abnormal non-clinically significant - n (%)	0	0	0	0	0		0
		Abnormal clinically significant - n (%)	0	0	0	0	0		0
	Day 4/ Discharge	n	22	22	9	9	11		73
		Normal - n (%)	22 (100 %)	22 (100 %)	9 (100 %)	9 (100 %)	11 (100 %)		73 (100 %)
		Abnormal non-clinically significant - n (%)	0	0	0	0	0		0
		Abnormal clinically significant - n (%)	0	0	0	0	0		0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Appendix 15.3.6.10

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_phyex.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.6.17 Summary of Physical Examination of Body Systems - Safety Population**

Body System	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Thyroid gland	Baseline	n	22	22	9	9	11	73
		Normal - n (%)	22 (100 %)	22 (100 %)	9 (100 %)	9 (100 %)	11 (100 %)	73 (100 %)
		Abnormal non-clinically significant - n (%)	0	0	0	0	0	0
		Abnormal clinically significant - n (%)	0	0	0	0	0	0
	Day 4/ Discharge	n	22	22	9	9	11	73
		Normal - n (%)	22 (100 %)	22 (100 %)	9 (100 %)	9 (100 %)	11 (100 %)	73 (100 %)
		Abnormal non-clinically significant - n (%)	0	0	0	0	0	0
		Abnormal clinically significant - n (%)	0	0	0	0	0	0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Appendix 15.3.6.10

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_phyex.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.6.17 Summary of Physical Examination of Body Systems - Safety Population**

Body System	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Heart	Baseline	n	22	22	9	9	11	73
		Normal - n (%)	22 (100 %)	22 (100 %)	9 (100 %)	9 (100 %)	11 (100 %)	73 (100 %)
		Abnormal non-clinically significant - n (%)	0	0	0	0	0	0
		Abnormal clinically significant - n (%)	0	0	0	0	0	0
	Day 4/ Discharge	n	22	22	9	9	11	73
		Normal - n (%)	22 (100 %)	22 (100 %)	9 (100 %)	9 (100 %)	11 (100 %)	73 (100 %)
		Abnormal non-clinically significant - n (%)	0	0	0	0	0	0
		Abnormal clinically significant - n (%)	0	0	0	0	0	0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Appendix 15.3.6.10

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_phyex.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.6.17 Summary of Physical Examination of Body Systems - Safety Population**

Body System	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Chest	Baseline	n	22	22	9	9	11	73
		Normal - n (%)	22 (100 %)	22 (100 %)	9 (100 %)	9 (100 %)	11 (100 %)	73 (100 %)
		Abnormal non-clinically significant - n (%)	0	0	0	0	0	0
		Abnormal clinically significant - n (%)	0	0	0	0	0	0
	Day 4/ Discharge	n	22	22	9	9	11	73
		Normal - n (%)	22 (100 %)	22 (100 %)	9 (100 %)	9 (100 %)	11 (100 %)	73 (100 %)
		Abnormal non-clinically significant - n (%)	0	0	0	0	0	0
		Abnormal clinically significant - n (%)	0	0	0	0	0	0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Appendix 15.3.6.10

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_phyex.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.6.17 Summary of Physical Examination of Body Systems - Safety Population**

Body System	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Lungs	Baseline	n	22	22	9	9	11	73
		Normal - n (%)	22 (100 %)	22 (100 %)	9 (100 %)	9 (100 %)	11 (100 %)	73 (100 %)
		Abnormal non-clinically significant - n (%)	0	0	0	0	0	0
		Abnormal clinically significant - n (%)	0	0	0	0	0	0
	Day 4/ Discharge	n	22	22	9	9	11	73
		Normal - n (%)	22 (100 %)	22 (100 %)	9 (100 %)	9 (100 %)	11 (100 %)	73 (100 %)
		Abnormal non-clinically significant - n (%)	0	0	0	0	0	0
		Abnormal clinically significant - n (%)	0	0	0	0	0	0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Appendix 15.3.6.10

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_phyex.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.6.17 Summary of Physical Examination of Body Systems - Safety Population**

Body System	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Gastrointestinal	Baseline	n	22	22	9	9	11	73
		Normal - n (%)	22 (100 %)	22 (100 %)	9 (100 %)	9 (100 %)	11 (100 %)	73 (100 %)
		Abnormal non-clinically significant - n (%)	0	0	0	0	0	0
		Abnormal clinically significant - n (%)	0	0	0	0	0	0
	Day 4/ Discharge	n	22	22	9	9	11	73
		Normal - n (%)	22 (100 %)	22 (100 %)	9 (100 %)	9 (100 %)	11 (100 %)	73 (100 %)
		Abnormal non-clinically significant - n (%)	0	0	0	0	0	0
		Abnormal clinically significant - n (%)	0	0	0	0	0	0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Appendix 15.3.6.10

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_phyex.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.6.17 Summary of Physical Examination of Body Systems - Safety Population**

Body System	Study Day	Statistic	Sequence				Enrolled Not		Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)	Randomized (N=11)		
Cardiovascular system	Baseline	n	22	22	9	9	11		73
		Normal - n (%)	22 (100 %)	22 (100 %)	9 (100 %)	9 (100 %)	11 (100 %)		73 (100 %)
		Abnormal non-clinically significant - n (%)	0	0	0	0	0		0
		Abnormal clinically significant - n (%)	0	0	0	0	0		0
	Day 4/ Discharge	n	22	22	9	9	11		73
		Normal - n (%)	22 (100 %)	22 (100 %)	9 (100 %)	9 (100 %)	11 (100 %)		73 (100 %)
		Abnormal non-clinically significant - n (%)	0	0	0	0	0		0
		Abnormal clinically significant - n (%)	0	0	0	0	0		0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Appendix 15.3.6.10

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_phyex.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.6.17 Summary of Physical Examination of Body Systems - Safety Population**

Body System	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Neurologic	Baseline	n	22	22	9	9	11	73
		Normal - n (%)	22 (100 %)	22 (100 %)	9 (100 %)	9 (100 %)	11 (100 %)	73 (100 %)
		Abnormal non-clinically significant - n (%)	0	0	0	0	0	0
		Abnormal clinically significant - n (%)	0	0	0	0	0	0
	Day 4/ Discharge	n	22	22	9	9	11	73
		Normal - n (%)	22 (100 %)	22 (100 %)	9 (100 %)	9 (100 %)	11 (100 %)	73 (100 %)
		Abnormal non-clinically significant - n (%)	0	0	0	0	0	0
		Abnormal clinically significant - n (%)	0	0	0	0	0	0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Appendix 15.3.6.10

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_phyex.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.6.17 Summary of Physical Examination of Body Systems - Safety Population**

Body System	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Skin	Baseline	n	22	22	9	9	11	73
		Normal - n (%)	22 (100 %)	22 (100 %)	9 (100 %)	9 (100 %)	11 (100 %)	73 (100 %)
		Abnormal non-clinically significant - n (%)	0	0	0	0	0	0
		Abnormal clinically significant - n (%)	0	0	0	0	0	0
	Day 4/ Discharge	n	22	22	9	9	11	73
		Normal - n (%)	22 (100 %)	22 (100 %)	9 (100 %)	9 (100 %)	11 (100 %)	73 (100 %)
		Abnormal non-clinically significant - n (%)	0	0	0	0	0	0
		Abnormal clinically significant - n (%)	0	0	0	0	0	0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Appendix 15.3.6.10

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_phyex.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.6.17 Summary of Physical Examination of Body Systems - Safety Population**

Body System	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Back	Baseline	n	22	22	9	9	11	73
		Normal - n (%)	22 (100 %)	22 (100 %)	9 (100 %)	9 (100 %)	11 (100 %)	73 (100 %)
		Abnormal non-clinically significant - n (%)	0	0	0	0	0	0
		Abnormal clinically significant - n (%)	0	0	0	0	0	0
	Day 4/ Discharge	n	22	22	9	9	11	73
		Normal - n (%)	22 (100 %)	22 (100 %)	9 (100 %)	9 (100 %)	11 (100 %)	73 (100 %)
		Abnormal non-clinically significant - n (%)	0	0	0	0	0	0
		Abnormal clinically significant - n (%)	0	0	0	0	0	0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Appendix 15.3.6.10

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_phyex.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.6.17 Summary of Physical Examination of Body Systems - Safety Population**

Body System	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Musculoskeletal	Baseline	n	22	22	9	9	11	73
		Normal - n (%)	22 (100 %)	22 (100 %)	9 (100 %)	9 (100 %)	11 (100 %)	73 (100 %)
		Abnormal non-clinically significant - n (%)	0	0	0	0	0	0
		Abnormal clinically significant - n (%)	0	0	0	0	0	0
	Day 4/ Discharge	n	22	22	9	9	11	73
		Normal - n (%)	22 (100 %)	22 (100 %)	9 (100 %)	9 (100 %)	11 (100 %)	73 (100 %)
		Abnormal non-clinically significant - n (%)	0	0	0	0	0	0
		Abnormal clinically significant - n (%)	0	0	0	0	0	0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Appendix 15.3.6.10

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_phyex.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.6.17 Summary of Physical Examination of Body Systems - Safety Population**

Body System	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Abdomen	Baseline	n	22	22	9	9	11	73
		Normal - n (%)	22 (100 %)	22 (100 %)	9 (100 %)	9 (100 %)	11 (100 %)	73 (100 %)
		Abnormal non-clinically significant - n (%)	0	0	0	0	0	0
		Abnormal clinically significant - n (%)	0	0	0	0	0	0
	Day 4/ Discharge	n	22	22	9	9	11	73
		Normal - n (%)	22 (100 %)	22 (100 %)	9 (100 %)	9 (100 %)	11 (100 %)	73 (100 %)
		Abnormal non-clinically significant - n (%)	0	0	0	0	0	0
		Abnormal clinically significant - n (%)	0	0	0	0	0	0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Appendix 15.3.6.10

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_phyex.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.6.17 Summary of Physical Examination of Body Systems - Safety Population**

Body System	Study Day	Statistic	Sequence				Enrolled Not Randomized (N=11)	Overall Safety (N=73)
			THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)		
Dentition	Baseline	n	22	22	9	9	11	73
		Normal - n (%)	22 (100 %)	22 (100 %)	9 (100 %)	9 (100 %)	11 (100 %)	73 (100 %)
		Abnormal non-clinically significant - n (%)	0	0	0	0	0	0
		Abnormal clinically significant - n (%)	0	0	0	0	0	0
	Day 4/ Discharge	n	22	22	9	9	11	73
		Normal - n (%)	22 (100 %)	22 (100 %)	9 (100 %)	9 (100 %)	11 (100 %)	73 (100 %)
		Abnormal non-clinically significant - n (%)	0	0	0	0	0	0
		Abnormal clinically significant - n (%)	0	0	0	0	0	0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: Baseline is the last available time point prior to the product test (THS 2.2 Menthol or NRT gum) at Admission (Day -1).

Appendix 15.3.6.10

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_phyex.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.6.18 Summary of Cough Assessments - Safety Population**

Variable	Sequence											
	THS 2.2 Menthol - mCC (N=22)			mCC - THS 2.2 Menthol (N=22)			THS 2.2 Menthol - NRT gum (N=9)			NRT gum - THS 2.2 Menthol (N=9)		
	n	(%)	Events	n	(%)	Events	n	(%)	Events	n	(%)	Events
Has subject experienced cough in the study period assessed?												
Yes	14	( 63.6%)	38	9	( 40.9%)	29	7	( 77.8%)	24	5	( 55.6%)	16
No	8	( 36.4%)	40	13	( 59.1%)	65	2	( 22.2%)	10	4	( 44.4%)	20
Intensity of cough												
Very mild	9	( 40.9%)	23	3	( 13.6%)	7	3	( 33.3%)	7	1	( 11.1%)	3
Mild	3	( 13.6%)	4	5	( 22.7%)	7	3	( 33.3%)	7	3	( 33.3%)	4
Moderate	1	( 4.5%)	1	1	( 4.5%)	1	1	( 11.1%)	1	1	( 11.1%)	1
Severe	1	( 4.5%)	1	0		0	0		0	0		0
Very severe	0		0	0		0	0		0	0		0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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 4 (to assess Day -1 to Day 3).

Note: If subject has answered question more than once then the most severe intensity is presented.

Appendix 15.3.6.13

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_cough.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.6.18 Summary of Cough Assessments - Safety Population**

Variable	Sequence										Overall Safety (N=73)  n (%) Events		
	THS 2.2 Menthol - mCC (N=22)		mCC - THS 2.2 Menthol (N=22)		THS 2.2 Menthol - NRT gum (N=9)		NRT gum - THS 2.2 Menthol (N=9)		Enrolled Not Randomized (N=11)				
	n	(%) Events	n	(%) Events	n	(%) Events	n	(%) Events	n	(%) Events			
Frequency of cough													
Rarely	8	( 36.4%)	21	5 ( 22.7%)	13	3 ( 33.3%)	9	1 ( 11.1%)	3	1 ( 9.1%)	1	18 ( 24.7%)	47
Sometimes	4	( 18.2%)	6	4 ( 18.2%)	13	4 ( 44.4%)	6	4 ( 44.4%)	8	0		16 ( 21.9%)	33
Fairly often	2	( 9.1%)	2	0		0		0		0		2 ( 2.7%)	2
Often	0			0		0		0		0		0	
Almost always	0			0		0		0		0		0	
Amount of sputum													
No sputum	7	( 31.8%)	19	5 ( 22.7%)	14	3 ( 33.3%)	7	2 ( 22.2%)	6	1 ( 9.1%)	1	18 ( 24.7%)	47
A moderate amount of sputum	7	( 31.8%)	12	4 ( 18.2%)	6	4 ( 44.4%)	12	3 ( 33.3%)	9	0		18 ( 24.7%)	39
A large amount of sputum	0			0		0		0		0		0	
A very large amount of sputum	0			0		0		0		0		0	

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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 4 (to assess Day -1 to Day 3).

Note: If subject has answered question more than once then the most severe intensity is presented.

Appendix 15.3.6.13

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_cough.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.6.18.1 Summary of Cough Assessments by Study Day - Safety Population**

Study Day/ Variable	Statistic	THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)	Enrolled Not Randomized (N=11)	Overall Safety (N=73)
<b>Day -1</b>							
Has subject experienced cough in last 24 hours							
Yes	n (%)	13 ( 59.1%)	6 ( 27.3%)	6 ( 66.7%)	4 ( 44.4%)	1 ( 9.1%)	30 ( 41.1%)
No	n (%)	9 ( 40.9%)	16 ( 72.7%)	3 ( 33.3%)	5 ( 55.6%)	9 ( 81.8%)	42 ( 57.5%)
VAS	n	13	6	6	4	1	30
	Mean (SD)	15.8 (17.54)	10.7 (13.52)	10.2 (9.63)	24.0 (25.71)	NC	14.2 (16.45)
	Median	9.0	6.0	7.5	19.0	NC	8.5
	Min, Max	0, 50	0, 36	0, 25	0, 58	NC	0, 58

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: The assessments performed at Day 0 to Day 4 are used to evaluate cough at Day -1 to Day 3, respectively.

Note: NC = Not calculated.

Appendix 15.3.6.13

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_cough2.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.6.18.1 Summary of Cough Assessments by Study Day - Safety Population**

Study Day/ Variable	Statistic	THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)	Enrolled Not Randomized (N=11)	Overall Safety (N=73)
Intensity of cough							
Very mild	n (%)	11 ( 50.0%)	4 ( 18.2%)	3 ( 33.3%)	1 ( 11.1%)	1 ( 9.1%)	20 ( 27.4%)
Mild	n (%)	1 ( 4.5%)	2 ( 9.1%)	3 ( 33.3%)	3 ( 33.3%)	0	9 ( 12.3%)
Moderate	n (%)	1 ( 4.5%)	0	0	0	0	1 ( 1.4%)
Severe	n (%)	0	0	0	0	0	0
Very severe	n (%)	0	0	0	0	0	0
Frequency of cough							
Rarely	n (%)	8 ( 36.4%)	3 ( 13.6%)	4 ( 44.4%)	2 ( 22.2%)	1 ( 9.1%)	18 ( 24.7%)
Sometimes	n (%)	4 ( 18.2%)	3 ( 13.6%)	2 ( 22.2%)	2 ( 22.2%)	0	11 ( 15.1%)
Fairly often	n (%)	1 ( 4.5%)	0	0	0	0	1 ( 1.4%)
Often	n (%)	0	0	0	0	0	0
Almost always	n (%)	0	0	0	0	0	0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: The assessments performed at Day 0 to Day 4 are used to evaluate cough at Day -1 to Day 3, respectively.

Appendix 15.3.6.13

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_cough2.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.6.18.1 Summary of Cough Assessments by Study Day - Safety Population**

Study Day/ Variable	Statistic	THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)	Enrolled Not Randomized (N=11)	Overall Safety (N=73)
Amount of sputum produced							
No sputum	n (%)	9 ( 40.9%)	5 ( 22.7%)	3 ( 33.3%)	1 ( 11.1%)	1 ( 9.1%)	19 ( 26.0%)
A moderate amount of sputum	n (%)	4 ( 18.2%)	1 ( 4.5%)	3 ( 33.3%)	3 ( 33.3%)	0	11 ( 15.1%)
A larger amount of sputum	n (%)	0	0	0	0	0	0
A very large amount of sputum	n (%)	0	0	0	0	0	0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: The assessments performed at Day 0 to Day 4 are used to evaluate cough at Day -1 to Day 3, respectively.

Appendix 15.3.6.13

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_cough2.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.6.18.1 Summary of Cough Assessments by Study Day - Safety Population**

Study Day/ Variable	Statistic	THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)	Enrolled Not Randomized (N=11)	Overall Safety (N=73)
<b>Day 0</b>							
Has subject experienced cough in last 24 hours							
Yes	n (%)	9 ( 40.9%)	8 ( 36.4%)	5 ( 55.6%)	4 ( 44.4%)	0	26 ( 35.6%)
No	n (%)	13 ( 59.1%)	14 ( 63.6%)	4 ( 44.4%)	5 ( 55.6%)	0	36 ( 49.3%)
VAS	n	9	8	5	4		26
	Mean (SD)	22.6 (23.98)	11.1 (11.21)	4.0 (4.53)	23.3 (24.68)		15.6 (18.86)
	Median	22.0	7.0	2.0	19.0		8.0
	Min, Max	0, 69	0, 31	0, 11	0, 55		0, 69

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: The assessments performed at Day 0 to Day 4 are used to evaluate cough at Day -1 to Day 3, respectively.

Appendix 15.3.6.13

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_cough2.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.6.18.1 Summary of Cough Assessments by Study Day - Safety Population**

Study Day/ Variable	Statistic	THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)	Enrolled Not Randomized (N=11)	Overall Safety (N=73)
Intensity of cough							
Very mild	n (%)	6 ( 27.3%)	4 ( 18.2%)	2 ( 22.2%)	3 ( 33.3%)		15 ( 20.5%)
Mild	n (%)	2 ( 9.1%)	4 ( 18.2%)	3 ( 33.3%)	1 ( 11.1%)		10 ( 13.7%)
Moderate	n (%)	0	0	0	0		0
Severe	n (%)	1 ( 4.5%)	0	0	0		1 ( 1.4%)
Very severe	n (%)	0	0	0	0		0
Frequency of cough							
Rarely	n (%)	6 ( 27.3%)	4 ( 18.2%)	4 ( 44.4%)	1 ( 11.1%)		15 ( 20.5%)
Sometimes	n (%)	2 ( 9.1%)	4 ( 18.2%)	1 ( 11.1%)	3 ( 33.3%)		10 ( 13.7%)
Fairly often	n (%)	1 ( 4.5%)	0	0	0		1 ( 1.4%)
Often	n (%)	0	0	0	0		0
Almost always	n (%)	0	0	0	0		0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: The assessments performed at Day 0 to Day 4 are used to evaluate cough at Day -1 to Day 3, respectively.

Appendix 15.3.6.13

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_cough2.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.6.18.1 Summary of Cough Assessments by Study Day - Safety Population**

Study Day/ Variable	Statistic	THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)	Enrolled Not Randomized (N=11)	Overall Safety (N=73)
Amount of sputum produced							
No sputum	n (%)	5 ( 22.7%)	5 ( 22.7%)	3 ( 33.3%)	1 ( 11.1%)		14 ( 19.2%)
A moderate amount of sputum	n (%)	4 ( 18.2%)	3 ( 13.6%)	2 ( 22.2%)	3 ( 33.3%)		12 ( 16.4%)
A larger amount of sputum	n (%)	0	0	0	0		0
A very large amount of sputum	n (%)	0	0	0	0		0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: The assessments performed at Day 0 to Day 4 are used to evaluate cough at Day -1 to Day 3, respectively.

Appendix 15.3.6.13

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_cough2.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.6.18.1 Summary of Cough Assessments by Study Day - Safety Population**

Study Day/ Variable	Statistic	THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)	Enrolled Not Randomized (N=11)	Overall Safety (N=73)
<b>Day 1</b>							
Has subject experienced cough in last 24 hours							
Yes	n (%)	6 ( 27.3%)	3 ( 13.6%)	5 ( 55.6%)	4 ( 44.4%)	0	18 ( 24.7%)
No	n (%)	15 ( 68.2%)	19 ( 86.4%)	4 ( 44.4%)	5 ( 55.6%)	0	43 ( 58.9%)
VAS	n	6	3	5	4		18
	Mean (SD)	14.2 (15.69)	NC	7.8 (7.53)	14.3 (11.88)		11.7 (11.57)
	Median	9.5	NC	4.0	14.0		10.0
	Min, Max	0, 41	NC	1, 19	0, 29		0, 41

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: The assessments performed at Day 0 to Day 4 are used to evaluate cough at Day -1 to Day 3, respectively.

Note: NC = Not calculated.

Appendix 15.3.6.13

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_cough2.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.6.18.1 Summary of Cough Assessments by Study Day - Safety Population**

Study Day/ Variable	Statistic	THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)	Enrolled Not Randomized (N=11)	Overall Safety (N=73)
Intensity of cough							
Very mild	n (%)	4 ( 18.2%)	1 ( 4.5%)	2 ( 22.2%)	3 ( 33.3%)		10 ( 13.7%)
Mild	n (%)	2 ( 9.1%)	1 ( 4.5%)	3 ( 33.3%)	0		6 ( 8.2%)
Moderate	n (%)	0	1 ( 4.5%)	0	1 ( 11.1%)		2 ( 2.7%)
Severe	n (%)	0	0	0	0		0
Very severe	n (%)	0	0	0	0		0
Frequency of cough							
Rarely	n (%)	4 ( 18.2%)	1 ( 4.5%)	4 ( 44.4%)	2 ( 22.2%)		11 ( 15.1%)
Sometimes	n (%)	2 ( 9.1%)	2 ( 9.1%)	1 ( 11.1%)	2 ( 22.2%)		7 ( 9.6%)
Fairly often	n (%)	0	0	0	0		0
Often	n (%)	0	0	0	0		0
Almost always	n (%)	0	0	0	0		0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: The assessments performed at Day 0 to Day 4 are used to evaluate cough at Day -1 to Day 3, respectively.

Appendix 15.3.6.13

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_cough2.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.6.18.1 Summary of Cough Assessments by Study Day - Safety Population**

Study Day/ Variable	Statistic	THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)	Enrolled Not Randomized (N=11)	Overall Safety (N=73)
Amount of sputum produced							
No sputum	n (%)	3 ( 13.6%)	3 ( 13.6%)	2 ( 22.2%)	2 ( 22.2%)		10 ( 13.7%)
A moderate amount of sputum	n (%)	3 ( 13.6%)	0	3 ( 33.3%)	2 ( 22.2%)		8 ( 11.0%)
A larger amount of sputum	n (%)	0	0	0	0		0
A very large amount of sputum	n (%)	0	0	0	0		0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: The assessments performed at Day 0 to Day 4 are used to evaluate cough at Day -1 to Day 3, respectively.

Appendix 15.3.6.13

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_cough2.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.6.18.1 Summary of Cough Assessments by Study Day - Safety Population**

Study Day/ Variable	Statistic	THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)	Enrolled Not Randomized (N=11)	Overall Safety (N=73)
<b>Day 2</b>							
Has subject experienced cough in last 24 hours							
Yes	n (%)	4 ( 18.2%)	5 ( 22.7%)	3 ( 33.3%)	1 ( 11.1%)	0	13 ( 17.8%)
No	n (%)	17 ( 77.3%)	17 ( 77.3%)	6 ( 66.7%)	8 ( 88.9%)	0	48 ( 65.8%)
VAS	n	4	5	3	1		13
	Mean (SD)	4.3 (5.00)	10.0 (8.00)	NC	NC		8.0 (6.97)
	Median	3.0	8.0	NC	NC		6.0
	Min, Max	0, 11	0, 20	NC	NC		0, 20

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: The assessments performed at Day 0 to Day 4 are used to evaluate cough at Day -1 to Day 3, respectively.

Note: NC = Not calculated.

Appendix 15.3.6.13

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_cough2.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.6.18.1 Summary of Cough Assessments by Study Day - Safety Population**

Study Day/ Variable	Statistic	THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)	Enrolled Not Randomized (N=11)	Overall Safety (N=73)
Intensity of cough							
Very mild	n (%)	4 ( 18.2%)	4 ( 18.2%)	3 ( 33.3%)	1 ( 11.1%)		12 ( 16.4%)
Mild	n (%)	0	1 ( 4.5%)	0	0		1 ( 1.4%)
Moderate	n (%)	0	0	0	0		0
Severe	n (%)	0	0	0	0		0
Very severe	n (%)	0	0	0	0		0
Frequency of cough							
Rarely	n (%)	4 ( 18.2%)	3 ( 13.6%)	2 ( 22.2%)	0		9 ( 12.3%)
Sometimes	n (%)	0	2 ( 9.1%)	1 ( 11.1%)	1 ( 11.1%)		4 ( 5.5%)
Fairly often	n (%)	0	0	0	0		0
Often	n (%)	0	0	0	0		0
Almost always	n (%)	0	0	0	0		0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: The assessments performed at Day 0 to Day 4 are used to evaluate cough at Day -1 to Day 3, respectively.

Appendix 15.3.6.13

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_cough2.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.6.18.1 Summary of Cough Assessments by Study Day - Safety Population**

Study Day/ Variable	Statistic	THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)	Enrolled Not Randomized (N=11)	Overall Safety (N=73)
Amount of sputum produced							
No sputum	n (%)	3 ( 13.6%)	4 ( 18.2%)	2 ( 22.2%)	1 ( 11.1%)		10 ( 13.7%)
A moderate amount of sputum	n (%)	1 ( 4.5%)	1 ( 4.5%)	1 ( 11.1%)	0		3 ( 4.1%)
A larger amount of sputum	n (%)	0	0	0	0		0
A very large amount of sputum	n (%)	0	0	0	0		0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: The assessments performed at Day 0 to Day 4 are used to evaluate cough at Day -1 to Day 3, respectively.

Appendix 15.3.6.13

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_cough2.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.6.18.1 Summary of Cough Assessments by Study Day - Safety Population**

Study Day/ Variable	Statistic	THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)	Enrolled Not Randomized (N=11)	Overall Safety (N=73)
<b>Day 3</b>							
Has subject experienced cough in last 24 hours							
Yes	n (%)	6 ( 27.3%)	7 ( 31.8%)	5 ( 55.6%)	3 ( 33.3%)	0	21 ( 28.8%)
No	n (%)	15 ( 68.2%)	15 ( 68.2%)	4 ( 44.4%)	6 ( 66.7%)	0	40 ( 54.8%)
VAS	n	6	7	5	3		21
	Mean (SD)	8.5 (10.14)	5.7 (4.93)	10.2 (17.87)	NC		7.4 (10.08)
	Median	5.5	6.0	4.0	NC		5.0
	Min, Max	1, 28	0, 14	0, 42	NC		0, 42

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: The assessments performed at Day 0 to Day 4 are used to evaluate cough at Day -1 to Day 3, respectively.

Note: NC = Not calculated.

Appendix 15.3.6.13

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_cough2.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.6.18.1 Summary of Cough Assessments by Study Day - Safety Population**

Study Day/ Variable	Statistic	THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)	Enrolled Not Randomized (N=11)	Overall Safety (N=73)
Intensity of cough							
Very mild	n (%)	6 ( 27.3%)	6 ( 27.3%)	4 ( 44.4%)	3 ( 33.3%)		19 ( 26.0%)
Mild	n (%)	0	1 ( 4.5%)	0	0		1 ( 1.4%)
Moderate	n (%)	0	0	1 ( 11.1%)	0		1 ( 1.4%)
Severe	n (%)	0	0	0	0		0
Very severe	n (%)	0	0	0	0		0
Frequency of cough							
Rarely	n (%)	5 ( 22.7%)	5 ( 22.7%)	4 ( 44.4%)	3 ( 33.3%)		17 ( 23.3%)
Sometimes	n (%)	1 ( 4.5%)	2 ( 9.1%)	1 ( 11.1%)	0		4 ( 5.5%)
Fairly often	n (%)	0	0	0	0		0
Often	n (%)	0	0	0	0		0
Almost always	n (%)	0	0	0	0		0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: The assessments performed at Day 0 to Day 4 are used to evaluate cough at Day -1 to Day 3, respectively.

Appendix 15.3.6.13

Path: /cvn/projects/prj/development/000000106326/dev/tables/t\_cough2.sas

Program Run: 27OCT14 cvn\_kbooth Program Status: FINAL

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**Table 15.2.6.18.1 Summary of Cough Assessments by Study Day - Safety Population**

Study Day/ Variable	Statistic	THS 2.2 Menthol - mCC (N=22)	mCC - THS 2.2 Menthol (N=22)	THS 2.2 Menthol - NRT gum (N=9)	NRT gum - THS 2.2 Menthol (N=9)	Enrolled Not Randomized (N=11)	Overall Safety (N=73)
Amount of sputum produced							
No sputum	n (%)	6 ( 27.3%)	6 ( 27.3%)	2 ( 22.2%)	2 ( 22.2%)		16 ( 21.9%)
A moderate amount of sputum	n (%)	0	1 ( 4.5%)	3 ( 33.3%)	1 ( 11.1%)		5 ( 6.8%)
A larger amount of sputum	n (%)	0	0	0	0		0
A very large amount of sputum	n (%)	0	0	0	0		0

Note: mCC = menthol conventional cigarettes; NRT gum = Nicotine Replacement Therapy gum; 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 Menthol or NRT gum.

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

Note: The assessments performed at Day 0 to Day 4 are used to evaluate cough at Day -1 to Day 3, respectively.

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