

# S1 Appendix: Model Formulation

## S1.1 Conceptual Model Description

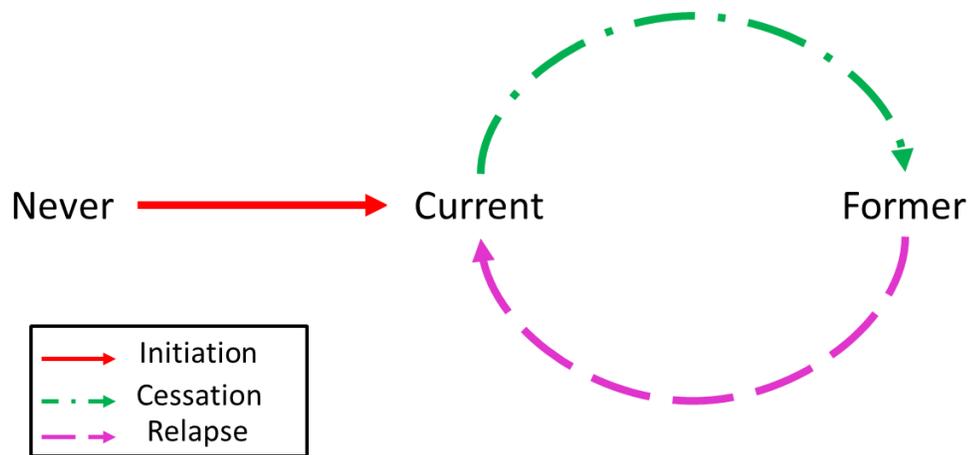
This section details the conceptual formulation for the dynamic, multi-state model for assessing the effects of tobacco product use behaviors on population health. Up to any combination of  $N$  tobacco products are being consumed within a specified population, and the model can be used to track the number of people in subpopulations over time. Subpopulations are defined to be groups of individuals that share a common set of the following attributes:

- Age, denoted by  $a = 0, 1, 2, \dots, G, G^+$ .  $G$  represents the maximum individual age tracked and  $G^+$  represents the group of ages greater than  $G$ . For example, model formulations frequently set  $G = 99$  and  $G^+$  represents  $a > 99$ .
- Sex, denoted by  $s = male, female$ .
- Tobacco product use status. An individual's tobacco product use status with respect to the  $N$  tobacco products is denoted with an  $N$ -tuple,  $u = (u_1, u_2, \dots, u_N)$ , where  $u_k, k = 1, \dots, N$ , is defined to be

$$u_k = \begin{cases} n, & \text{never user of product } k \\ c, & \text{current user of product } k \\ f_j, & \text{former user of product } k \text{ who quit } j \text{ years ago, } j = 1, 2, \dots, M_k \end{cases}$$

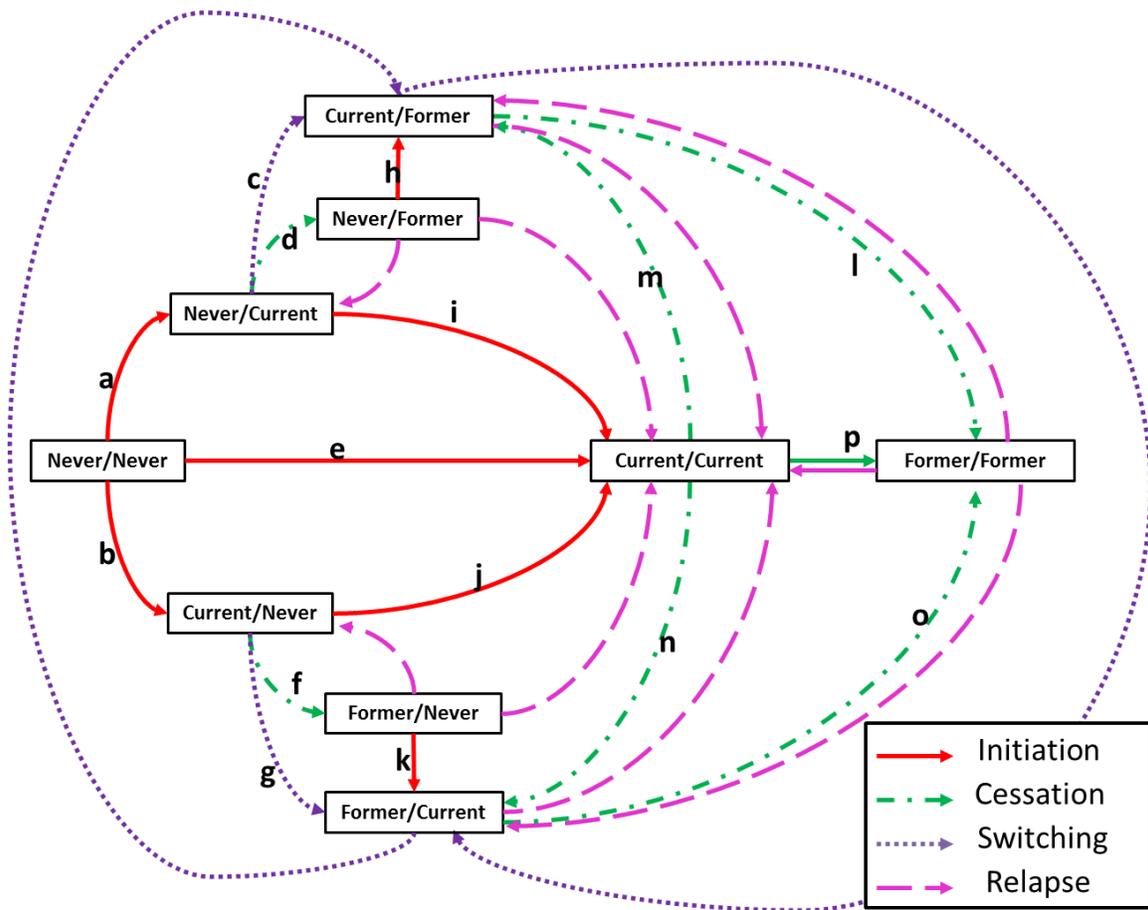
where  $M_k$  denotes the maximum number of years tracked for quitters of product  $k$ . This manuscript uses the notation  $u_{never}$  to denote the tobacco product use status of never use for all  $N$  products.

Over time, individuals age during each modeled time step; individuals may die; and individuals may change their tobacco product use status, e.g., they may initiate new products, quit using products, relapse and take up products they had previously quit, and so on. Changes in age and tobacco use status cause individuals to move from one subpopulation to another. The probability that an individual transitions from one tobacco use status to another is determined by the individual's current age, sex, and tobacco use status. Fig. A and B show the set of tobacco-use-status transitions for one- and two-product model formulations, respectively. This model can be extended to include additional products.



**Figure A: Tobacco-use-status transitions for a single product model formulation.**

**Figure Note:** In the current model transition takes place among subpopulations defined by age, sex, and tobacco use status.



**Figure B: Tobacco-use-status transitions for a two-product model**

**Figure Note:** Transitions are categorized into four different behavior groups: initiation behaviors, cessation behaviors, switching behaviors, and relapse behaviors. The boxes represent the nine possible use statuses, with the first and second terms corresponding to the first and second tobacco products, respectively.

Deaths are calculated using all-cause mortality probabilities. Mortality probabilities vary by age, sex, and tobacco product use status. It is assumed that current and former tobacco product use may result in an elevated mortality risk, relative to never use of the products. Deaths decrease the overall size of the population (and subpopulations), and births contribute to the population size. Net international migration can increase or decrease the population size, depending on whether immigration or emigration is larger.

The model tracks the number of individuals in each subpopulation over time. At each time step<sup>1</sup> and for each subpopulation, the model tracks the number of individuals within the subpopulation, the number of individuals that leave the subpopulation and join another subpopulation, and the number of individuals within the subpopulation who die. Key model output measures include prevalence of use for each tobacco product, mortality, and mortality attributable to tobacco product use.

## S1.2 Mathematical Model Description

Let  $A = \{0, 1, \dots, G, G^+\}$  be the set of ages,  $S = \{male, female\}$  be the set of sexes, and  $U$  be the set of all possible tobacco product use status  $N$ -tuples considered in the model. Furthermore, let  $F$  denote the set of childbearing ages for females. Then Equations (1) through (9) provide a mathematical representation of the model, and Table A defines key model parameters and variables. Given that much of the data necessary to develop parameters is provided on an annual basis, time steps are generally taken to be one year increments, and parameters and variables are defined and described accordingly. A different sized time step could certainly be used, and interpretation of parameter and variable definition would need to change accordingly.

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<sup>1</sup> In its most general form, the model can use any specified time step for which appropriate parameters can be determined. In practice, annual time steps are most commonly used.

$$Pop(0, s, u_{never}, t_{i+1}) = b(s, t_{i+1}) + m(0, s, u_{never}, t_{i+1}) \quad (1)$$

$$Pop(0, s, u, t_{i+1}) = 0, u \neq u_{never} \quad (2)$$

$$Pop(a+1, s, u, t_{i+1}) = \sum_{x \in U} Pop(a, s, x, t_i) \times p(x \rightarrow u | a, s, x, t_{i+1}) \times [1 - p(death | a, s, u, t_{i+1})] \\ + m(a+1, s, u, t_{i+1}), a = 0, \dots, G-1 \quad (3)$$

$$Pop(G^+, s, u, t_{i+1}) = \sum_{x \in U} Pop(G, s, x, t_i) \times p(x \rightarrow u | G, s, x, t_{i+1}) \times [1 - p(death | G, s, u, t_{i+1})] \\ + \sum_{x \in U} Pop(G^+, s, x, t_i) \times p(x \rightarrow u | G^+, s, x, t_{i+1}) \times [1 - p(death | G^+, s, u, t_{i+1})] \\ + m(G^+, s, u, t_{i+1}) \quad (4)$$

$$m(a, s, u, t_{i+1}) = m_{frac}(a, s, u, t_{i+1}) \times m_{rate}(t_{i+1}) \times \sum_{a_1 \in A} \sum_{s_1 \in S} \sum_{u_1 \in U} Pop(a_1, s_1, u_1, t_i) \quad (5)$$

$$b(s, t_{i+1}) = b_{frac}(s, t_{i+1}) \times b_{rate}(t_{i+1}) \times \sum_{a \in F} \sum_{u \in U} [Pop(a, female, u, t_i) + m(a, female, u, t_{i+1})] \quad (6)$$

$$p(death | a, s, u, t_i) = RR(a, s, u, t_i) \times p(death | a, s, u_{never}, t_i) \quad (7)$$

$$prev(A_1, S_1, U_1, t_i) = \frac{\sum_{a \in A_1} \sum_{s \in S_1} \sum_{u \in U_1} Pop(a, s, u, t_i)}{\sum_{a \in A_1} \sum_{s \in S_1} \sum_{u \in U} Pop(a, s, u, t_i)}, A_1 \subset A, S_1 \subset S, U_1 \subset U \quad (8)$$

$$AD(A_1, S_1, t_{i+1}) = \sum_{a \in A_1} \sum_{s \in S_1} \sum_{x, u \in U} Pop(a, s, x, t_i) \times p(x \rightarrow u | a, s, x, t_{i+1}) \times [RR(a, s, u, t_{i+1}) - 1] \times p(death | a, s, u_{never}, t_{i+1}), \\ A_1 \subset A, S_1 \subset S \quad (9)$$

**Table A: Model parameters and variables**

Parameter	Description	Input Parameter or Output Variable
$Pop(a, s, u, t_i)$	Number of individuals of age $a$ , sex $s$ , and tobacco use status $u$ at year $t_i$ . The population at year $t_0$ is defined to be the initial population.	output (except for the initial population which is an input parameter)
$p(x \rightarrow u   a, s, x, t_{i+1})$	Annual proportion of individuals of age $a$ , sex $s$ , and tobacco use status $x$ that transition to tobacco use status $u$ in the time interval $(t_i, t_{i+1}]$ . When $x = u$ , this parameter represents the rate at which individuals maintain and do not change their tobacco use status.	input
$p(death   a, s, u, t_{i+1})$	Annual proportion of individuals with age $a$ , sex $s$ , and tobacco use status $u$ that die in the time interval $(t_i, t_{i+1}]$	input
$b_{rate}(t_{i+1})$	Annual number of births per female of childbearing age during the time interval $(t_i, t_{i+1}]$	input
$b_{frac}(s, t_{i+1})$	Fraction of births that are of sex $s$ during the time interval $(t_i, t_{i+1}]$	input
$b(s, t_{i+1})$	Number of births of sex $s$ during the time interval $(t_i, t_{i+1}]$	output
$m_{rate}(t_{i+1})$	Annual number of net international migrants per person in the population entering/leaving during the time interval $(t_i, t_{i+1}]$	input
$m_{frac}(a, s, u, t_{i+1})$	Fraction of net international migrants entering/leaving the population during the time interval $(t_i, t_{i+1}]$ of age $a$ , sex $s$ , and tobacco use status $u$	input
$m(a, s, u, t_{i+1})$	Number of net international migrants of age $a$ ,	output

Parameter	Description	Input Parameter or Output Variable
	sex $s$ , and tobacco use status $u$ entering/leaving the population during the time interval $(t_i, t_{i+1}]$	
$prev(A_1, S_1, U_1, t_i)$	Prevalence of individuals with tobacco use status in the subset $U_1 \subset U$ among the population of age in the subset $A_1 \subset A$ and sex in the subset $S_1 \subset S$ at year $t_i$	output
$AD(A_1, S_1, t_{i+1})$	Attributable deaths (from all products) among the population of age in the subset $A_1 \subset A$ and sex in the subset $S_1 \subset S$ during the time interval $(t_i, t_{i+1}]$	output

Equations [1] – [4] describe how subpopulations are tracked. The number of newborns (age 0) is determined solely by births and migration of infants less than age 1 (Equation [1]), and all newborns are assumed to have never used any of the tobacco products (Equation [2]). For older ages (Equations [3] and [4]), the size of a subpopulation is calculated by determining the number of people from the previous year who transition into a particular subpopulation and do not die and the number of net international migrants entering or leave the subpopulation.

Equations [5] and [6] calculate the number of net international migrants and births, respectively. Births are assumed to be linearly proportional to the number of females of childbearing ages in the population, and the number of immigrants is assumed to be linearly proportional to the entire population. These assumptions are made for consistency with U.S. Census Bureau projection methods. However, it should be noted that alternative birth and

immigration functions could be easily substituted for the linear functions used in Equations [5] and [6].

The model assumes that the all-cause mortality proportion for a subpopulation is the product of the all-cause mortality relative risk ( $RR$ ) for this subpopulation and the base mortality probability of dying for never users of any of the  $N$  tobacco products by sex and age (Equation [7]). (The base mortality probability for never users is specified in S2 Appendix). Because use of any combination of the  $N$  tobacco products is assumed to have no protective effect (relative to never use), all relative risks are greater than or equal to 1. Similar to births and migration, an alternative risk formulation could be substituted for this calculation.

Equations [8] and [9] describe how to calculate product use prevalence among a population and deaths attributable to use of any combination of the  $N$  tobacco products, respectively. The prevalence calculation is straightforward.

# S2 Appendix: Data Inputs and Assumptions

## S2.1. Baseline One-Product Parameter Inputs and Assumptions

### S2.1.1 Baseline Scenario

The baseline one-product scenario projects the effect of tobacco use and harm in the US population over time where the single product is cigarette smoking and all parameters, except for the mortality scale factors, remain constant from 2000 through 2050. Fig. A shows the possible product use states and transitions involving the single product. Table A provides a summary of the one-product simulation parameter values that are used in our baseline scenario projections.

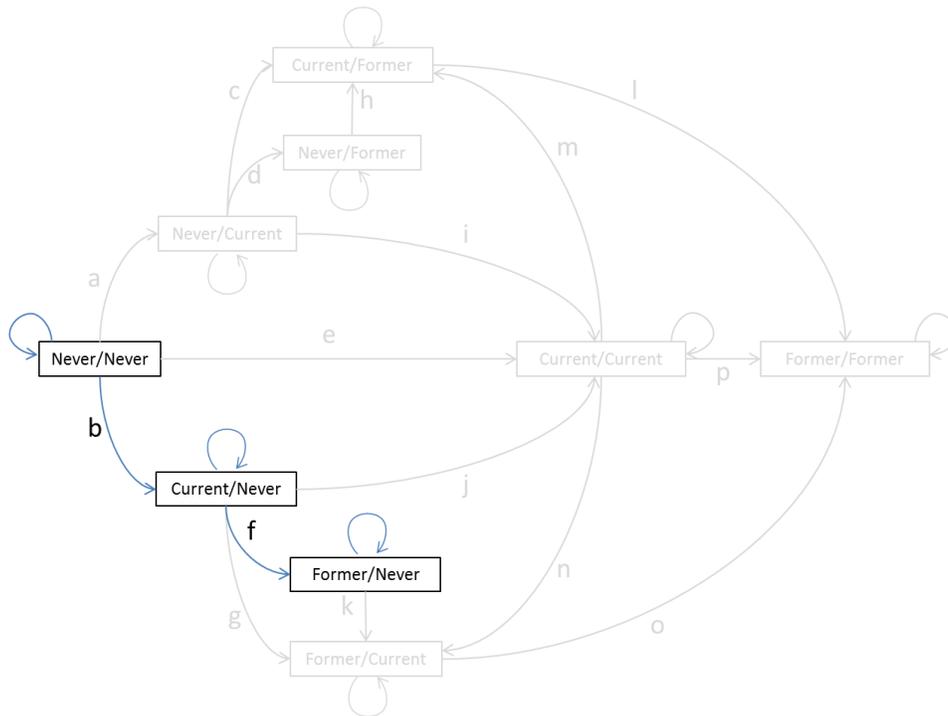


Figure A: One-product use states and transitions

**Table A: Summary of baseline parameter inputs and data sources**

Model Component	Model Parameter	Data Source	Notes
<b>Initial Population</b>	Population distribution by sex and age	US Census National Population Estimates for 2000 [1]: <a href="http://www.census.gov/population/projections/data/national/2008/downloadablefiles.html">http://www.census.gov/population/projections/data/national/2008/downloadablefiles.html</a> , Table 1.	
	Cigarette smoking status (current, former, and never smoker) by sex, age, and time since cessation	National Center for Health Statistics (NCHS), National Health Interview Survey (NHIS) data from 2000 [2].	
<b>Births</b>	Birth rate by sex	Computed from US Census National Population Projections 2008 [1]: <a href="http://www.census.gov/population/projections/data/national/2008/downloadablefiles.html">http://www.census.gov/population/projections/data/national/2008/downloadablefiles.html</a> , Table 2.	Birth rates were calculated by dividing the projected number of births by sex by the projected number of females of ages 15-34 years for 2000-2050 from Census projections and applying these ratios to the number of females of ages 15-34 years projected for 2000-2050 by our model.
<b>Net International Migration</b>	Net migration rate by sex	Computed from US Census National Population Projections 2008 [1] - <a href="http://www.census.gov/population/projections/data/national/2008/downloadablefiles.html">http://www.census.gov/population/projections/data/national/2008/downloadablefiles.html</a> , Table 4.	Net migration rates by sex were calculated by dividing the projected number of net international migrants by the projected population size by sex for 2000-2050 from Census projections and applying these ratios to the population sizes by sex projected for 2000-2050 by our model.
	Immigrant age distribution	US Census Bureau, The 2012 Statistical Abstract, the National Data Book, Table 40, 2012 [3]- <a href="http://www.census.gov/compendia/statab/cats/population/native_and_foreign-born_populations.html">http://www.census.gov/compendia/statab/cats/population/native_and_foreign-born_populations.html</a> , Table 40.	
	Immigrant smoking prevalence by sex	NHIS data from 2007-2011 [2].	Smoking prevalence was calculated for immigrants to the US of ages 18 years and over who had been in the US less than five years.

**Table A: Summary of baseline parameter inputs and data sources**

Model Component	Model Parameter	Data Source	Notes
Deaths	Never-smoker death rate by sex and age	US Census Bureau, The 2012 Statistical Abstract, the National Data Book, Table 110, 2012 - death rates from 2000 for ages under 35 [3] and death rates calculated from National Health Interview Survey – Linked Mortality Files (NHIS-LMF) [4] data from 2002-2006 for ages 35 and over.	US death rates were used as death rates for never smokers of ages less than 35 years, given that smoking-attributable mortality is low at these ages. Never-smoker death rates for ages 35 years and over were estimated from NHIS-LMF data for NHIS Sample Adult Questionnaire participants from 1997-2004 followed for mortality through linkage with the National Death Index from 2002-2006.
	Mortality adjustment factor by sex and age	<p>US Vital Statistics and NHIS-LMF death rates from 2002-2006. US Vital Statistics death rates come from annual mortality reports.</p> <ul style="list-style-type: none"> <li>National Vital Statistics Report, Deaths: Final Data for 2002, Table 5 [5] <a href="http://www.cdc.gov/nchs/data/nvsr/nvsr53/nvsr53_05acc.pdf">http://www.cdc.gov/nchs/data/nvsr/nvsr53/nvsr53_05acc.pdf</a></li> <li>National Vital Statistics Report, Deaths: Final Data for 2003, Table 5 [6] <a href="http://www.cdc.gov/nchs/data/nvsr/nvsr54/nvsr54_13.pdf">http://www.cdc.gov/nchs/data/nvsr/nvsr54/nvsr54_13.pdf</a></li> <li>National Vital Statistics Report, Deaths: Final Data for 2004, Table 5 [7] <a href="http://www.cdc.gov/nchs/data/nvsr/nvsr55/nvsr55_19.pdf">http://www.cdc.gov/nchs/data/nvsr/nvsr55/nvsr55_19.pdf</a></li> <li>National Vital Statistics Report, Deaths: Final Data for 2005, Table 5 [8] <a href="http://www.cdc.gov/nchs/data/nvsr/nvsr56/nvsr56_10.pdf">http://www.cdc.gov/nchs/data/nvsr/nvsr56/nvsr56_10.pdf</a></li> <li>National Vital Statistics Report, Deaths: Final Data for 2006, Table 5 [9] <a href="http://www.cdc.gov/nchs/data/nvsr/nvsr57/nvsr57_14.pdf">http://www.cdc.gov/nchs/data/nvsr/nvsr57/nvsr57_14.pdf</a></li> </ul>	NHIS-LMF never-smoker death rates were adjusted for low mortality in the NHIS’s civilian non-institutionalized population by multiplying the rates by the ratio of US Vital Statistics death rates divided by NHIS-LMF death rates by sex and age.
	Mortality scaling factors	Calculated from U.S. death rates using Lee-Carter Method [10]. US death rates come from Human Mortality Database data.	Mortality scaling factors for 2000-2050 were calculated from US death rates for 1950-1999 using the Lee-Carter mortality projection method to account for expected age-specific changes in mortality over time.

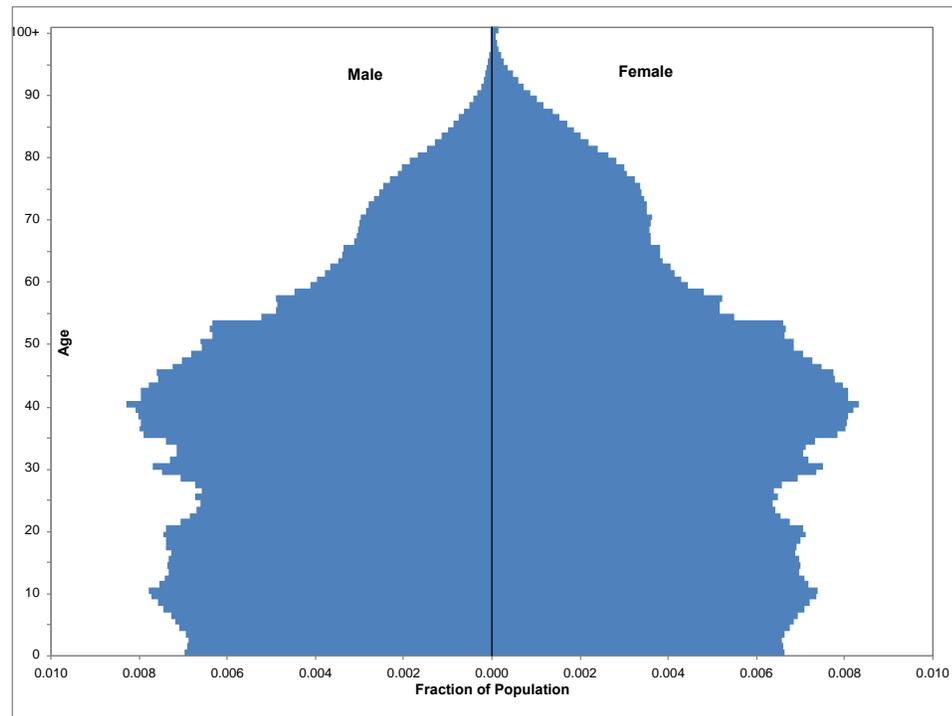
Model Component	Model Parameter	Data Source	Notes
	Relative risk by sex, age, smoking status, and age at cessation for former smokers	Table A: Summary of baseline parameter inputs and data sources	Hazard ratios and relative risks were estimated for 1997-2004 NHIS Sample Adult Questionnaire participants followed for mortality through linkage with the National Death Index through the end of 2006.
Cigarette Smoking Transition Behaviors	Sex and age-specific initiation rates	Reconstructions of cohort smoking histories from NHIS data [11].	
	Sex and age-specific cessation rates	Reconstructions of cohort smoking histories from NHIS data [11].	
	Sex and age-specific relapse rates	Set to 0.	

## S2.1.2 Initial Population Estimates

The initial population in our model is the US population by sex, age, and smoking status for 2000, which is the initial year in our population projections.

### 1. Year 2000 Population Distribution by Sex and Age

US population estimates for 2000 by sex and age come from US Census Bureau estimates [1] and are shown in Fig. B.



**Figure B: US Population distribution by sex and age for 2000 from US Census Bureau estimates**

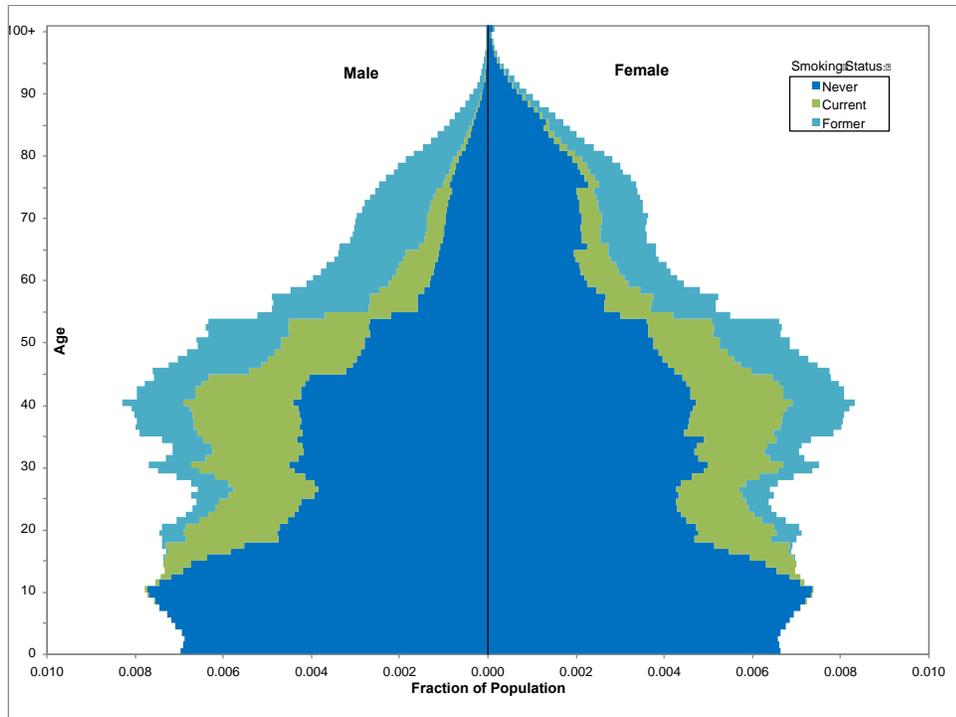
### 2. Year 2000 Smoking Prevalence by Sex, Age and Time since Cessation

US adult cigarette smoking prevalence for 2000 was estimated using National Health Interview Survey (NHIS) Sample Adult Questionnaire data [2]. NHIS is a national health survey of the US civilian non-institutionalized population that is conducted by the National Center for

Health Statistics (NCHS) on an annual basis. NHIS data are used by the Centers for Disease Control and Prevention (CDC) to estimate smoking prevalence for the US adult population[12]. Prevalence was estimated for current and former smokers by sex and age, as shown in Table B. The proportional distribution of former smokers by time since cessation was also estimated, as shown in Tables C (Males) and D (Females). Smoking status was defined using the categories commonly used by the Centers for Disease Control and Prevention [12]. Current smokers reported having smoked at least 100 cigarettes in their lives and currently smoking every day or some days; former smokers reported having smoked at least 100 cigarettes in their lives and not currently smoking at all; and never smokers reported not having smoked at least 100 cigarettes in their lives. All NHIS estimates were calculated using SAS version 9.3 using the appropriate NHIS sample weights and taking into account the NHIS complex survey design in accordance with analytic guidelines provided by NCHS [24].

The NHIS Sample Child Questionnaire does not ask about tobacco use. Smoking prevalence for the US in 2000 for ages of less than 18 was obtained from estimates of cohort smoking histories for 1980-1984 birth cohorts that were reconstructed from cross-sectional NHIS data [11]. Current smoking prevalence by age for these cohorts was estimated by multiplying the estimated ever smoking prevalence at each age by the cumulative proportion of smokers who had not quit smoking by that age.

Figure C shows the initial population distribution in our model by sex, age, and smoking status which was generated using US Census Bureau population estimates and NHIS smoking prevalence data.



**Figure C: US population distribution by sex, age, and smoking status for 2000 from US Census Bureau estimates and the National Health Interview Survey**

**Table B: Adult smoking prevalence by sex and age, National Health Interview Survey, 2000**

Age	Males			
	Current	95% CI	Former	95% CI
<b>18-24</b>	28.5%	25.9%, 31.1%	7.4%	5.9%, 9.0%
<b>25-34</b>	29.0%	27.0%, 31.1%	12.3%	10.9%, 13.7%
<b>35-44</b>	30.2%	28.2%, 32.2%	16.5%	15.0%, 17.9%
<b>45-54</b>	28.8%	26.8%, 30.9%	28.9%	26.6%, 31.2%
<b>55-64</b>	22.6%	20.4%, 24.8%	45.0%	42.3%, 47.7%
<b>65-74</b>	13.7%	11.7%, 15.6%	53.9%	50.9%, 56.9%
<b>75-84</b>	5.7%	4.0%, 7.3%	58.7%	54.8%, 62.6%
<b>85+</b>	3.8%	1.0%, 6.6%	58.3%	49.8%, 66.7%
<b>Total</b>	25.7%	24.8%, 26.6%	25.8%	24.9%, 26.7%

Females			
Current	95% CI	Former	95% CI
25.1%	22.6%, 27.5%	8.0%	6.6%, 9.4%
22.5%	20.9%, 24.1%	10.9%	9.6%, 12.2%
26.2%	24.6%, 27.7%	17.1%	15.6%, 18.5%
22.2%	20.5%, 23.8%	23.1%	21.3%, 24.9%
20.8%	18.9%, 22.8%	28.3%	26.1%, 30.5%
12.2%	10.5%, 13.8%	28.8%	26.3%, 31.3%
6.9%	5.6%, 8.2%	25.1%	22.6%, 27.6%
3.8%	2.1%, 5.6%	19.3%	14.5%, 23.7%
21.0%	20.2%, 21.8%	18.9%	18.2%, 19.6%

**Table C: Proportional distribution of male former smokers by time since cessation by sex and age, National Health Interview Survey, 2000**

Age	Years since Cessation														
	0-4	5-9	10-14	15-19	20-24	25-29	30-34	35-39	40-44	45-49	50-54	55-59	60-64	65-69	70+
<b>18-24</b>	88.6%	11.4%	-	-	-	-	-	-	-	-	-	-	-	-	-
<b>25-34</b>	58.7%	24.0%	15.0%	2.3%	-	-	-	-	-	-	-	-	-	-	-
<b>35-44</b>	31.9%	18.1%	19.9%	13.8%	14.2%	2.1%	-	-	-	-	-	-	-	-	-
<b>45-54</b>	18.5%	13.4%	16.4%	15.1%	16.2%	11.5%	7.9%	1.0%	-	-	-	-	-	-	-
<b>55-64</b>	16.6%	10.5%	14.5%	14.3%	13.5%	12.1%	8.9%	5.0%	4.4%	0.1%	-	-	-	-	-
<b>65-74</b>	9.7%	8.3%	11.0%	7.9%	13.9%	8.2%	15.4%	7.9%	10.9%	4.7%	1.9%	0.2%	-	-	-
<b>75-84</b>	6.4%	7.3%	8.5%	9.5%	14.1%	9.6%	12.8%	6.3%	12.9%	4.2%	6.2%	0.2%	1.5%	0.2%	-
<b>85+</b>	6.6%	7.0%	8.2%	8.2%	10.4%	11.1%	11.5%	5.8%	13.6%	2.5%	2.4%	4.5%	3.1%	1.6%	3.5%

**Table D: Proportional distribution of female former smokers by time since cessation by sex and age, National Health Interview Survey, 2000**

Age	Years since Cessation														
	0-4	5-9	10-14	15-19	20-24	25-29	30-34	35-39	40-44	45-49	50-54	55-59	60-64	65-69	70+
<b>18-24</b>	92.6%	7.4%	-	-	-	-	-	-	-	-	-	-	-	-	-
<b>25-34</b>	60.1%	24.6%	13.6%	1.7%	-	-	-	-	-	-	-	-	-	-	-
<b>35-44</b>	27.8%	18.2%	21.9%	16.4%	13.4%	2.3%	-	-	-	-	-	-	-	-	-
<b>45-54</b>	22.6%	11.5%	13.5%	18.0%	15.3%	10.9%	8.0%	0.3%	-	-	-	-	-	-	-
<b>55-64</b>	19.0%	9.9%	13.2%	13.7%	13.4%	9.4%	11.5%	5.7%	3.4%	0.8%	-	-	-	-	-
<b>65-74</b>	13.1%	14.0%	13.5%	11.6%	11.1%	9.3%	10.9%	5.0%	6.0%	1.9%	3.5%	-	-	-	-
<b>75-84</b>	9.5%	8.8%	11.3%	10.8%	15.0%	8.5%	11.6%	6.4%	7.6%	2.8%	4.1%	1.6%	1.9%	-	-
<b>85+</b>	14.4%	3.4%	16.5%	4.9%	11.6%	4.6%	11.7%	3.0%	4.0%	6.7%	7.5%	4.0%	5.0%	2.7%	-

### S2.1.3 Births and Net International Migration

The US Census Bureau provides birth, net international migration, and population projections for the US population from 2001-2050 [1]. We use these projections to parameterize our model with birth and net migration rates. We limit fertility to ages less than 35 years to prevent any effects of tobacco-attributable mortality on fertility in different scenarios, given that our model introduces tobacco-attributable mortality at age 35. We compute an annual birth rate ( $BR$ ) by sex of infant ( $s$ ) for the time interval  $t$  to  $t + 1$  for use in the model by dividing the projected birth count by the projected female population count for ages 15-34 years for each year from 2001-2050.

$$\begin{aligned}
 B_{t,t+1}(s) &= \text{Census projected births of sex}=s \text{ in the interval } t \text{ to } t + 1 \\
 \sum_{a=15}^{34} P_t(\text{female}, a) &= \text{Census projected female population ages 15-34 at time } t \\
 BR_{t,t+1}(s) &= \frac{B_{t,t+1}(s)}{\sum_{a=15}^{34} P_t(\text{female}, a)}
 \end{aligned}$$

Birth rates used in the model for the period from 2001-2050 range between 5.2 and 5.4 male births and 5.0 and 5.1 female births per 100 women. We then use the calculated birth rates to project the number of births by sex in the interval  $t$  to  $t + 1$  by applying the rate to the projected female population ages 15-34 years in our model at time  $t$ , where  $u$  tracks the tobacco use state.

$$\begin{aligned}
 \sum_{a=15}^{34} \sum_u P_t(\text{female}, a, u) &= \text{model projected female population ages of 15-34 at time } t \\
 B_{t,t+1}(s, \text{never}) &= BR_{t,t+1}(s) \cdot \sum_{a=15}^{34} \sum_u P_t(\text{female}, a, u)
 \end{aligned}$$

The Census Bureau also provides net international migration and total population projections by year from 2001-2050 [1]. We compute a net migration rate ( $MR$ ) by sex ( $s$ ) for

the interval  $t$  to  $t + 1$  for use in our model by dividing the net international migration projections by the total population projections by year.

$$\begin{aligned} \sum_a M_{t,t+1}(s, a) &= \text{Census projected net migrants with sex}=s \text{ for the interval } t \text{ to } t + 1 \\ \sum_{s,a} P_t(s, a) &= \text{Census projected total population at time } t \\ MR_{t,t+1}(s) &= \frac{\sum_a M_{t,t+1}(s, a)}{\sum_{s,a} P_t(s, a)} \end{aligned}$$

Projected net migration for the US from 2001-2050 is consistently positive, indicating greater immigration than emigration. Calculated net international migration rates used in our model are between 1.2 and 2.2 per 1000 for males and between 1.5 and 2.5 per 1000 for females.

We use these calculated net migration rates in our model to project the net number of immigrants by sex ( $s$ ) in the interval  $t$  to  $t + 1$  by applying them to the projected total population in the model at time  $t$ .

$$\begin{aligned} \sum_{s,a,u} P_t(s, a, u) &= \text{model projected total population at time } t \\ M_{t,t+1}(s) &= MR_{t,t+1}(s) \cdot \sum_{s,a,u} P_t(s, a, u) \end{aligned}$$

We obtained the age distribution of immigrants,  $M_{t,t+1}(s, a)$ , from US Census Bureau estimates of the age distribution of immigrants in the US in 2010 who had arrived in the US between 2000 and 2010 [3].

Smoking prevalence for the incoming net international migrant population,  $M_{t,t+1}(s, a, u)$ , is based on an analysis of 2007-2011 NHIS data for adult immigrants who had been in the US less than five years. These years were the most recent with NHIS data available, and smoking prevalence was estimated from five years of data for immigrants who had been in the US for less than five years to increase the precision of the estimates. The current smoking

prevalence for these immigrants was 19.1% for males and 4.4% for females and the former smoking prevalence was 13.1% for males and 5.6% for females (NHIS). These estimates are used in the model for incoming immigrants ages 18 years and over throughout the projection period from 2001-2050. We assigned years since cessation to the former smoker immigrant population using the distributions by sex and age observed for the US in 2000 NHIS data, as described above.

## S2.1.4 Deaths

### 3. S2.1.4.1 Death Rates for Never Smokers

Never-smoker death rates provide the base for estimates of mortality and smoking-attributable mortality in our model. These rates are projected for the period from 2000-2050. US death rates for the year 2000 by sex and age from vital statistics data are used to project never-smoker death rates for ages of less than 35 years [13], given that smoking-attributable mortality is low at these ages. Never-smoker death rates for ages of 35 years and over are projected from estimates from National Health Interview Survey – Linked Mortality Files (NHIS-LMF) data [4]. NHIS-LMF data provide recent, nationally representative estimates of mortality risks and have been used extensively in the analysis of mortality risks by smoking status [14,15] and in modeling smoking-attributable mortality [16]. The NHIS-LMF data used in our analysis come from NHIS Sample Adult Questionnaire participants from 1997-2004 followed for mortality through linkage with the National Death Index through the end of 2006. The NHIS-LMF never-smoker death rates ( $m$ ) are estimated by ten-year age group as the ratio of deaths ( $d$ ) to person-time ( $L$ ) for never smokers ( $u = ns$ ) during mortality follow-up from 2002-2006.

$$NHIS - LMF m(s, a, ns) = \frac{d(s, a, ns)}{L(s, a, ns)}$$

The NHIS-LMF never-smoker death rates are shown in Table E. NHIS-LMF death rates tend to be somewhat lower than US death rates, given that the NHIS considers the US civilian non-institutionalized population, which excludes individuals in long-term care facilities and nursing homes. The NHIS-LMF never-smoker death rates were therefore adjusted by multiplying them by the ratio of US death rates from vital statistics data divided by NHIS-LMF death rates by sex and age for 2002-2006.

$$NHIS - LMF_{adj} m(s, a, ns) = NHIS - LMF m(s, a, ns) \cdot \frac{US m(s, a)}{NHIS - LMF m(s, a)}$$

These ratios are shown in Table F (Males) and Table G (Females).

**Table E: Never-smoker death rates per 100,000 person-years from mortality follow-up from 2002-2006 for 1997-2004 NHIS Sample Adult Questionnaire participants**

Age	Males		Females	
	Death Rate per 100,000 person years	Standard Error	Death Rate per 100,000 person years	Standard Error
35-44	143	18	83	12
45-54	361	30	201	21
55-64	564	54	489	40
65-74	1,731	127	1,055	58
75-84	4,367	255	3,198	112
85+	14,151	707	10,498	267

**Table F: Male death rates from follow-up from 2002-2006 for 1997-2004 NHIS Sample Adult Questionnaire participants and mean 2002-2006 US death rates by sex and age**

Age	Death Rates			Ratio
	NHIS-LMF Rate	Standard Error	US Rate	
35-44	195	16	248	1.268
45-54	530	25	547	1.031
55-64	1,010	44	1,144	1.133
65-74	2,665	92	2,680	1.005
75-84	5,918	156	6,465	1.092
85+	15,323	483	15,256	0.996

**Table G: Female death rates from follow-up from 2002-2006 for 1997-2004 NHIS Sample Adult Questionnaire participants and mean 2002-2006 US death rates by sex and age**

Age	Death Rates			Ratio
	NHIS-LMF Rate	Standard Error	US Rate	
35-44	122	12	146	1.192
45-54	278	17	317	1.141
55-64	673	30	713	1.060
65-74	1,540	57	1,773	1.151
75-84	4,027	103	4,573	1.135
85+	11,075	259	13,522	1.221

#### 4. S2.1.4.2 Mortality Projections

Our model projects never-smoker death rates from 2000-2050. Rates are projected using mortality scaling factors obtained from the Lee-Carter mortality forecasting method [10] as implemented in the demography package version 1.16 for R [17]. This method was used to

project expected age-specific changes in mortality over time based on observed death rates. The Lee-Carter models death rates as a function of age ( $x$ ) and time ( $t$ ) as:

$$\ln(m_{x,t}) = a_x + b_x k_t + e_{x,t}$$

where  $a_x$  is an age coefficient that defines the basic mortality level by age,  $b_x$  is an age coefficient that identifies change in mortality over time,  $k_t$  is a parameter for mortality change over time, and  $e_{x,t}$  is an error term. The model is fit to data using a least-squares solution found from the singular value decomposition. Death rates are then forecast by modeling  $k_t$  as a random walk with drift:

$$k_t = c + k_{t-1} + u_t$$

where  $c$  is a drift term representing the average annual change in  $k_t$  and  $u_t$  is an error term.

The Lee-Carter method was used to project US death rates from 2000-2050 by sex and age using observed US death rates from 1950-1999. The resulting projected rates were centered around 2000 for ages of less than 35 years and around 2004 for ages of 35 years and over. Mortality scaling factors by sex and age were calculated as the ratio of projected rates for 2000-2050 to the projected rates for 2000 for ages of less than 35 years and for 2004 for ages of 35 years and over. These scaling factors were then applied to the never-smoker death rates explained above for 2000 for ages of less than 35 years and for 2004 for ages of 35 years and over to produce projected never-smoker death rates by sex and age for 2000-2050.

$$msf(s, a < 35, t = 2000 - 2050) = \frac{\text{projected US } m(s, a, t)}{\text{projected US } m(s, a, t = 2000)}$$

$$msf(s, a \geq 35, t = 2000 - 2050) = \frac{\text{projected US } m(s, a, t)}{\text{projected US } m(s, a, t = 2004)}$$

$$NHIS - LMF_{adj} m(s, a, ns, t) = NHIS - LMF_{adj} m(s, a, ns) \cdot msf(s, a, t)$$

The scaling factors by ten-year intervals are shown in Table H.

**Table H: Mortality scaling factors obtained from Lee-Carter mortality forecasting method for US from 2000-2050 by sex and age**

Age	Males						Females					
	2000	2010	2020	2030	2040	2050	2000	2010	2020	2030	2040	2050
0	1	0.719	0.516	0.371	0.267	0.192	1	0.752	0.565	0.425	0.320	0.240
1	1	0.769	0.592	0.455	0.350	0.269	1	0.779	0.607	0.473	0.369	0.288
5	1	0.774	0.599	0.463	0.359	0.278	1	0.800	0.640	0.512	0.409	0.327
10	1	0.830	0.689	0.572	0.475	0.394	1	0.851	0.725	0.617	0.525	0.447
15	1	0.942	0.887	0.835	0.786	0.740	1	0.921	0.848	0.781	0.719	0.662
20	1	0.934	0.872	0.814	0.760	0.709	1	0.895	0.800	0.716	0.641	0.573
25	1	0.957	0.917	0.877	0.840	0.804	1	0.886	0.785	0.695	0.616	0.545
30	1	0.972	0.944	0.917	0.891	0.866	1	0.878	0.771	0.677	0.595	0.522
35	1.024	0.966	0.911	0.859	0.810	0.764	1.057	0.920	0.801	0.697	0.606	0.528
40	1.043	0.939	0.845	0.761	0.685	0.617	1.060	0.916	0.791	0.683	0.590	0.509
45	1.056	0.921	0.803	0.701	0.611	0.533	1.057	0.921	0.802	0.699	0.609	0.530
50	1.064	0.911	0.780	0.668	0.572	0.490	1.051	0.928	0.819	0.722	0.637	0.562
55	1.058	0.919	0.797	0.692	0.601	0.521	1.041	0.942	0.852	0.771	0.698	0.631
60	1.053	0.925	0.813	0.714	0.627	0.551	1.039	0.945	0.860	0.782	0.712	0.648
65	1.048	0.933	0.831	0.740	0.659	0.586	1.041	0.942	0.853	0.772	0.699	0.633
70	1.039	0.944	0.857	0.778	0.706	0.641	1.044	0.937	0.841	0.754	0.677	0.607
75	1.037	0.947	0.865	0.790	0.722	0.660	1.052	0.927	0.818	0.721	0.636	0.561
80	1.029	0.958	0.893	0.832	0.775	0.722	1.045	0.936	0.839	0.752	0.673	0.603
85	1.017	0.976	0.936	0.899	0.863	0.828	1.025	0.964	0.906	0.853	0.802	0.754

### 5. S2.1.4.3 Mortality Probability Conversion and Calculations

Age-specific death rates by sex and time were assumed to be equal to these adjusted and projected death rates by sex, time, and ten-year age group. These age-specific death rates ( $m$ ) which have person-time as their denominator, were then converted to age-specific probabilities of dying ( $q$ ), which have persons-at-risk as their denominator, for use in calculating mortality in the population model. The conversion was made using the standard demographic formula [18]:

$$q_x = \frac{m_x}{1 + (1 - a_x)m_x}$$

where  $x$  is year of age and  $a_x$  is the average length of time lived at that age by people who die at that age. Values for  $a_x$  at young ages were taken from Coale and Demeny model life tables and were  $0.045 + 2.684 * m_0$  for males and  $0.053 + 2.800 * m_0$  for females at age 0 and  $(1.651 - 2.816 * m_0) / 4$  for males and  $(1.522 - 1.518 * m_0) / 4$  for females at ages of 1-4 years [18]. Values for  $a_x$  at ages of 5 years and over were set equal to 0.5.

Never-smoker probabilities of dying were then multiplied by relative risks ( $RR$ ) for current and former smokers compared to never smokers by sex and age to produce probabilities of dying for current and former smokers for use in the population model.

$$q(s, a, u, t) = NHIS - LMF_{adj} q(s, a, ns, t) \cdot RR(s, a, u, t)$$

### 6. S2.1.4.4 Smoking Relative Risks

Relative risks by smoking status were estimated as hazard ratios using NHIS-LMF data for 1997-2004 NHIS Sample Adult Questionnaire data with mortality follow-up through the end of 2006. The hazard ratios compare mortality risks for current and former smokers to those of never smokers; this comparison is shown in Table I (current smokers), Table J (male former smokers observed), Table K (female former smokers observed), and Table L (former smokers used in modeling). The hazard ratios were estimated by sex and age using age as the time

scale and adjusting for race/ethnicity, educational attainment, alcohol consumption, and body mass index. Hazard ratios for former smokers were also estimated by age at cessation. Estimated hazard ratios for the most recent quitters among former smokers were often quite high, sometimes higher than estimated hazard ratios for current smokers of the same age. Similar results have been observed previously for recent quitters [19,20], and the most common explanation is that some of these individuals quit smoking due to symptoms or diagnoses of smoking-related illnesses but still had increased mortality risks after smoking cessation. To prevent increases in mortality from occurring in scenarios with increased smoking cessation, we set the maximum hazard ratio for former smokers equal to the hazard ratio for current smokers of the same age and sex. Estimated hazard ratios for former smokers who quit before the age of 40 were generally slightly above or below 1.0, consistent with previous research that has found limited increased mortality risks for former smokers who stopped smoking before this age [14]. We therefore set these hazard ratios to 1.0 for use in the population model.

**Table I: Current-smoker mortality hazard ratios from follow-up from 1997-2006 for 1997-2004 NHIS Sample Adult Questionnaire participants by sex, age, and smoking status**

Age	Males		Females	
	HR	95% CI	HR	95% CI
35-44	1.84	(1.34, 2.52)	3.34	(2.37, 4.72)
45-54	2.04	(1.62, 2.58)	2.42	(1.87, 3.12)
55-64	2.79	(2.28, 3.40)	2.35	(1.92, 2.87)
65-74	2.77	(2.34, 3.29)	2.90	(2.48, 3.39)
75-84	2.19	(1.88, 2.54)	2.57	(2.28, 2.90)
85+	1.36	(1.01, 1.84)	1.56	(1.28, 1.90)

**Table J: Male former smoker observed mortality hazard ratios from follow-up from 1997-2006 for 1997-2004 NHIS Sample Adult Questionnaire participants by sex, age, and smoking status**

Age	Age at Cessation							
	< 40		40-49		50-59		60+	
	HR	95% CI	HR	95% CI	HR	95% CI	HR	95% CI
35-44	1.05	(0.67, 1.64)	5.78	(2.01, 16.59)				
45-54	0.99	(0.73, 1.35)	1.68	(1.18, 2.39)	4.60	(2.39, 8.86)		
55-64	0.96	(0.71, 1.28)	1.54	(1.16, 2.06)	2.44	(1.77, 3.35)	2.56	(1.22, 5.39)
65-74	0.76	(0.60, 0.95)	1.19	(0.96, 1.48)	1.87	(1.52, 2.30)	2.20	(1.82, 2.67)
75-84	1.02	(0.87, 1.21)	1.43	(1.20, 1.69)	1.50	(1.29, 1.74)	1.99	(1.75, 2.77)
85+	1.07	(0.89, 1.29)	1.12	(0.93, 1.36)	1.04	(0.86, 1.26)	1.35	(1.15, 1.58)

**Table K: Female former smoker observed mortality hazard ratios from follow-up from 1997-2006 for 1997-2004 NHIS Sample Adult Questionnaire participants by sex, age, and smoking status**

Age	Age at Cessation							
	< 40		40-49		50-59		60+	
	HR	95% CI	HR	95% CI	HR	95% CI	HR	95% CI
35-44	1.67	(1.00, 2.81)	3.70	(0.69, 19.97)				
45-54	1.05	(0.70, 1.57)	1.57	(0.96, 2.56)	6.06	(2.08, 17.59)		
55-64	0.81	(0.52, 1.25)	1.71	(1.18, 2.47)	2.56	(1.92, 3.41)	6.30	(3.39, 11.73)
65-74	1.06	(0.76, 1.47)	1.56	(1.19, 2.05)	1.94	(1.58, 2.38)	3.20	(2.54, 4.03)
75-84	1.09	(0.85, 1.39)	1.38	(1.11, 1.70)	1.52	(1.26, 1.83)	2.23	(1.99, 2.50)
85+	1.02	(0.79, 1.31)	1.09	(0.82, 1.44)	1.37	(1.11, 1.70)	1.54	(1.35, 1.75)

**Table L: Former-smoker mortality hazard ratios used in modeling**

Age	Males				Females			
	Age at Cessation				Age at Cessation			
	< 40	40-49	50-59	60+	< 40	40-49	50-59	60+
35-44	1.00	1.84			1.00	3.34		
45-54	1.00	1.68	2.04		1.00	1.57	2.42	
55-64	1.00	1.54	2.44	2.56	1.00	1.71	2.35	2.35
65-74	1.00	1.19	1.87	2.20	1.00	1.56	1.94	2.90
75-84	1.00	1.43	1.50	1.99	1.00	1.38	1.52	2.23
85+	1.00	1.00	1.00	1.35	1.00	1.00	1.37	1.54

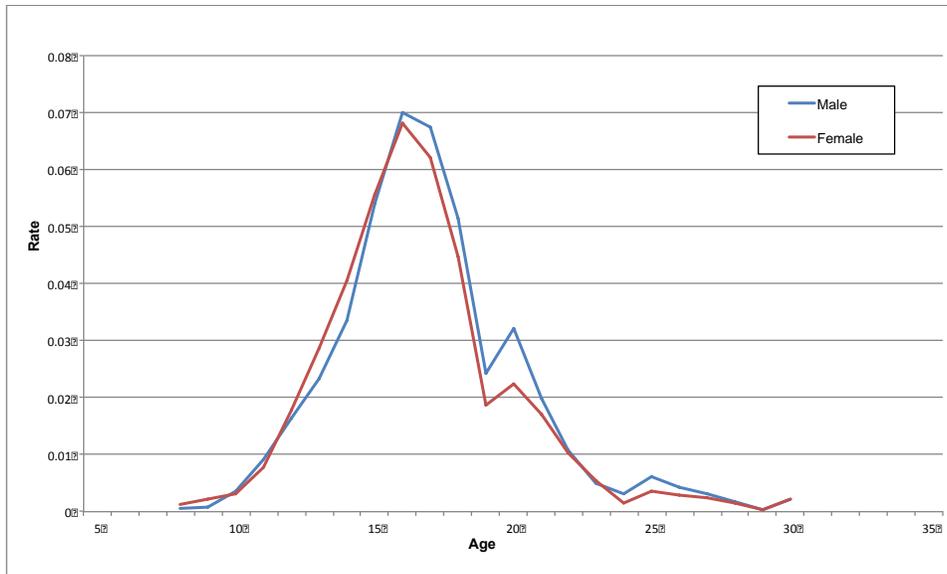
### S2.1.5 Smoking Initiation and Cessation

We use sex- and age-specific cigarette initiation and cessation rates in our model that were produced from US birth cohort smoking history data. The estimates were generated as part of the CISNET (Cancer Intervention and Surveillance Modeling Network) Lung Working Group, which is sponsored by the National Cancer Institute, and have been used to model and quantify the effect of reduced cigarette smoking on lung cancer mortality in the US from 1975 to 2000 [21]. Working group researchers estimated cohort smoking histories using 25 NHIS surveys administered from 1965 to 2001 [11]. The researchers estimated current- and ever-smoking prevalence and initiation and cessation rates by age and sex for five-year birth cohorts for individuals born between 1900 and 1984. A detailed explanation of their methodological approach and the resulting data are available on-line [22].

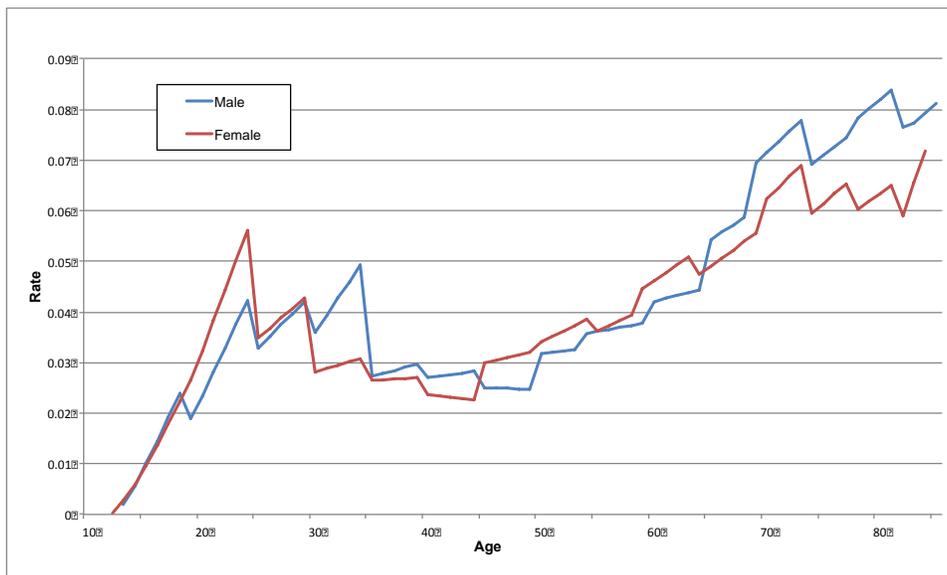
We use cohort age-specific smoking initiation rates for the period closest to 2000, the initial year in our population model projections. We use initiation rates for ages 0-19 years from 1980-1984 birth cohorts, rates for ages 20-24 from 1975-1979 cohorts, rates for ages 25-29 from

1970-1974 cohorts, and rates for age 30 from 1965-1969 cohorts. We do not include cigarette initiation beyond the age of 30 in our model, given that the overwhelming majority of smoking initiation has occurred by this age [23]. Figure D shows the estimated initiation rates by sex and age that are used in the model. The rates reflect initiation to established use as indicated by NHIS participants reporting if they had smoked at least 100 cigarettes in their lives. These age-specific initiation rates are assumed to remain constant throughout the projection period in the baseline scenario.

We also use smoking cessation rates in the model that come from CISNET cohort data. Rates for ages 1-18 years come from 1980-1984 birth cohorts, rates for ages 19-24 come from 1975-1979 cohorts, rates for ages 25-29 come from 1970-1974 cohorts, and so on through ages 83-85 from 1900-1904 cohorts. Cessation rates are available from the cohort data through age 85 for males and age 84 for females. Beyond these ages, cessation rates in the model are assumed to remain equal to the last available estimate from the cohort data. Figure E shows the estimated cessation rates by sex and age that are used in the model. The model allows for relapse, but transition probabilities for relapse behaviors are set to zero in the modeling simulations presented here because the CISNET cessation rates reflect successful smoking cessation for at least two years. These age-specific cessation rates are assumed to remain constant throughout the projection period in the baseline scenario.



**Figure D: Cigarette Smoking Initiation Rates by Sex and Age from CISNET Cohort Smoking Histories**



**Figure E: Cigarette Smoking Cessation Rates by Sex and Age from CISNET Cohort Smoking Histories**

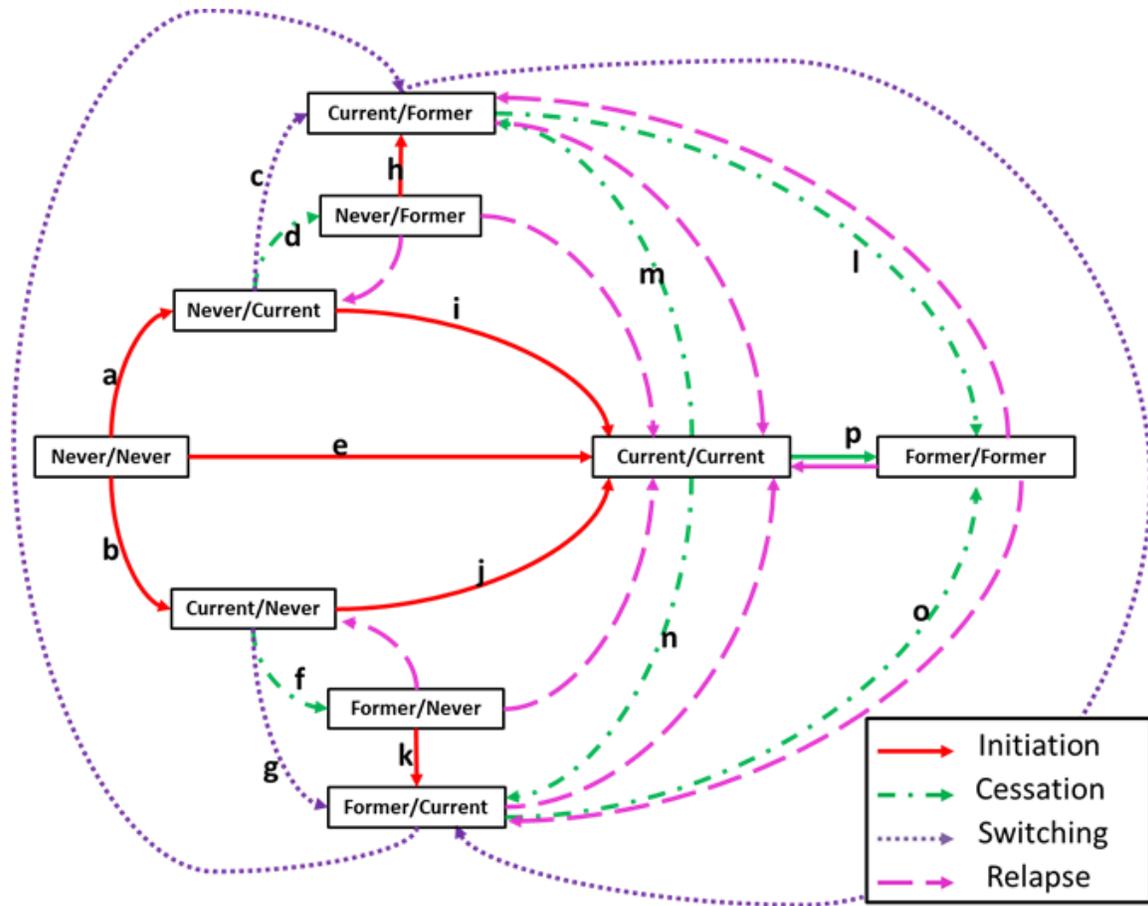
## S2.2. Alternative Two-Product Scenario Parameter Inputs and Assumptions

### S2.2.1 Two-Product Scenarios

The alternative two-product scenarios project the effect of introduction of a hypothetical new tobacco product on tobacco use and harm in the US population over time. The scenario begins in 2000 with cigarettes as the only product, using the input values described in Section S2.1. The hypothetical new product is introduced in 2003. Figure F shows the possible product use states and transitions involving the two products. Table M provides a summary of the two-product simulation parameter values that are allowed to vary in the model projections and their implementation in the model. Sex- and age-specific initiation rates and excess relative risk for the new product are defined as proportions of initiation rates and excess relative risk for cigarettes. Initiation for the hypothetical new tobacco product and switching from this new product to cigarettes is not allowed after age 30, given that most tobacco initiation occurs among young people. Initiation rates for a product for former users of the other product (transitions **h** and **k** in Figure F) are set equal to initiation rates for this product for never users of the other product. Initiation to dual use in a single year is assumed to be minimal, so these initiation rates (transition **e**) are set equal to 0. Cessation rates for new product users (transition **d**) are set equal to baseline smoking cessation rates and cessation rates for multiple product (any combination of current or former for both products) users (transitions **l**, **m**, **n**, **o**, and **p**) are also set equal to these smoking cessation rates.

The new product is assumed to have an excess relative risk that is equal to a proportion of the excess relative risk for cigarette smoking. Relative risks for current dual

users are set equal to the maximum of the relative risks for the individual products. Former cigarette smokers who are current new product users are assumed to have lower ERR than current smokers, but higher ERR than they would have had had they quit tobacco products entirely. Table N details the scenario relative risk calculations and assumptions for all tobacco product use statuses.



**Figure F: Alternative two-product use states and transitions**

**Figure Note:** Product Use #1/Product Use #2 represents users' use of Product #1 (cigarettes) and Product #2 (hypothetical new product). Product use relapse is incorporated into the model but not included in the scenarios described in this manuscript.

**Table M: Summary of two-product parameter inputs and implementation in model**

Parameter	Variable	Base Value	Range	Implementation in Model
<b>Health Risk of the New Product</b>				
Excess relative risk factor- a multiplicative factor that defines new product risks in terms of cigarette risks	$\alpha$	0.25	[0.01,0.50]	All-cause mortality relative risk for a never smoker-current user of the new product is $RR_{current\ new} = 1 + \alpha * (RR_{current\ smoker} - 1)$  All-cause mortality relative risk for a former smoker that switches to the new product is $RR_{switch} = RR_{former\ smoker} + \alpha * [RR_{current\ smoker} - RR_{former\ smoker}]$
<b>Impact on Current Smokers</b>				
Proportion of cigarette smokers who switch to new product use on an annual basis	$\beta$	0.015 (A)*	[0,0.03]	$g(2003^+) = \beta$
		0.010 (B)**		
Proportion of cigarette smokers who transition to dual product use on an annual basis	$\gamma$	0.015 (A)*	[0,0.03]	$j(2003^+) = \gamma$
		0.020 (B)**		
Proportion of switchers and dual users coming from smokers who would have otherwise quit smoking that year	$\delta$	0.25	[0,0.5]	$f(2003^+) = f(2000) - \delta * \beta - \delta * \gamma$
<b>Impact on Never Smokers</b>				
New product initiation factor – represents the initiation rate for the new product among never smokers as a proportion of the cigarette smoking initiation rate	$\varepsilon$	0.5	[0.25,0.75]	$a(2003^+) = \varepsilon * b(2000)$
Proportion of new product initiates who would have otherwise initiated cigarettes that year	$\eta$	0.5	[0.25,0.75]	$b(2003^+) = (1 - \varepsilon * \eta) * b(2000^+)$
Proportion of new product users who switch to cigarette use on an annual basis	$\theta$	0.05	[0,0.1]	$c(2003^+) = \theta$
Proportion of new product users who transition to dual produce use on an annual basis	$\iota$	0.05	[0,0.1]	$i(2003^+) = \iota$
* Hypothetical scenario A in which transition rate to dual use and switching are equivalent				

\*\* Hypothetical scenario B in which transition rate to dual use is greater than switching completely

Table Note:  $x(Y^+)$  represents the rate of transition with label  $x$  from Figure S2.6. for year  $Y$  and subsequent years in the projection period.

**Table N: Relative risk (RR) scenario assumption**

<b>Tobacco Use Status*</b>	<b>Relative Risk Calculation</b>	<b>Interpretation</b>
<b>NN</b>	$RR = 1$	Never/Never use represents the minimum risk state.
<b>CN</b>	RR for current smoker	CN is state for current smokers who do not use the new product.
<b>FN</b>	RR for former smoker	FN is state for former smokers who do not use the new product.
<b>NC</b>	$RR = 1 + \alpha * [ERR \text{ for current smoker}]$	Scenario assumes excess relative risk (ERR) for current user of the new product is a proportion of current smoker ERR defined by the excess relative risk factor $\alpha$ (from Table B.8).
<b>CC</b>	$RR = \text{maximum}[RR \text{ for current smoker, } RR \text{ for current new product use}]$	RR for dual use is the maximum of the individual RRs.
<b>FC</b>	$RR = RR \text{ for former smoker} + \alpha * [RR \text{ for current smoker} - RR \text{ for former smoker}]$	FC is state for former smokers who are current users of the new product. Use of the new product is assumed to result in a lower RR than smoking but a higher RR than the FN population that quits tobacco entirely. We again use the excess relative risk factor $\alpha$ to define the RR in terms of the RR for current and former smokers.
<b>NF</b>	$RR = 1 + \alpha * [ERR \text{ for former smoker}]$	Scenario assumes ERR for former user of the new product is a proportion of former smoker ERR
<b>CF</b>	RR = Current-Smoker Risk	CF is state for current smokers who are former users of the new product. These individuals are assigned an RR equal to current smokers' RR.
<b>FF</b>	$RR = \text{max}(\text{former smoker, former user})$	FF is state for former smokers and former users of the new product. The individuals retain the RR from the highest risk behavior.

\*First letter denotes cigarette use, and second letter denotes new product use. N=never, C=current, and F=former

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# **S3 Appendix: Model Diagnostics and Validation**

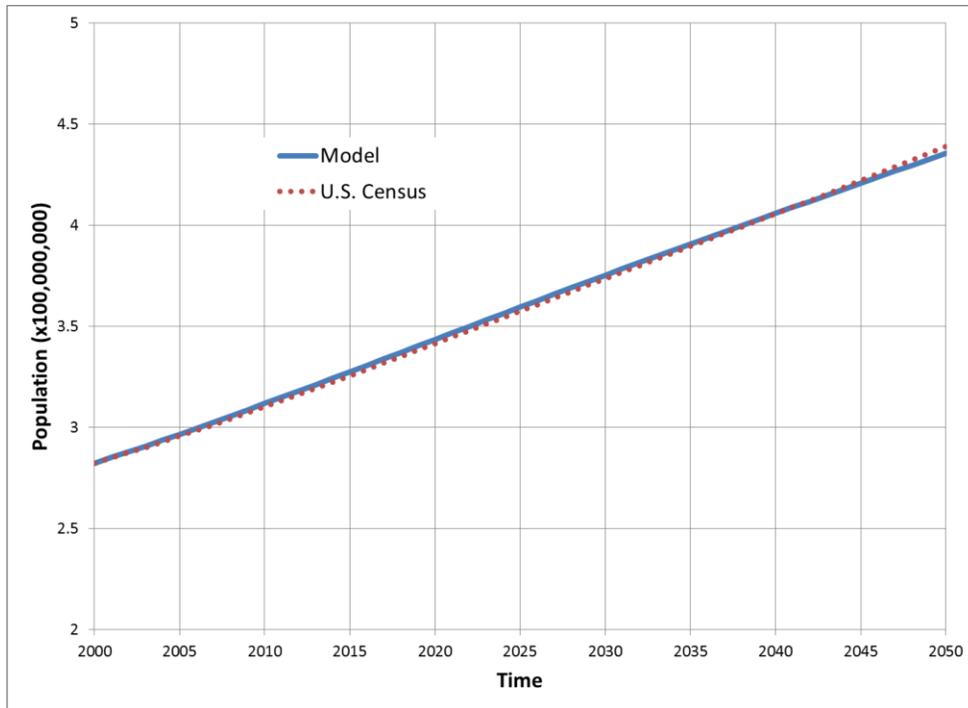
## **S3.1. Method**

To aid in model evaluation and validation, we present comparisons of results from our model and published estimates and projections. We first compare US population and mortality projections from our model with US Census Bureau projections for the period from 2000 to 2050. We also compare US adult cigarette smoking prevalence estimates from our model to estimates from the National Health Interview Survey (NHIS) for the period from 2000 to 2012. NHIS estimates are used by the Centers for Disease Control and Prevention (CDC) as estimates of smoking prevalence for the US population. We also present estimates of smoking-attributable deaths from our model in comparison with those from the CDC.

## **S3.2. Results**

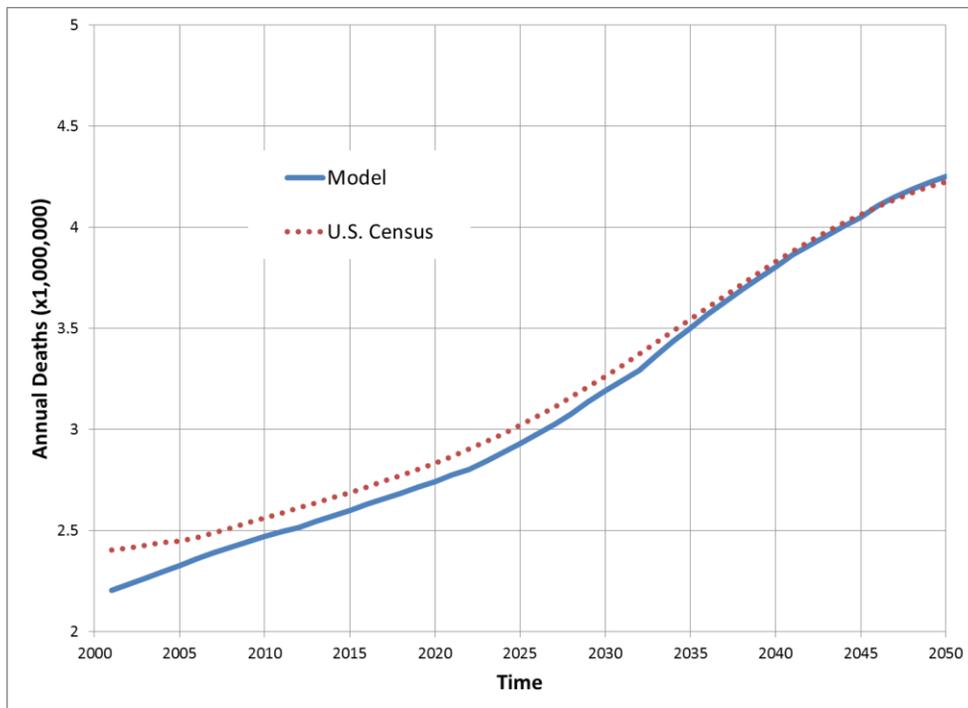
### **B. S3.2.1. Compare Population and Mortality Projections to Census**

Fig. A presents US Census and model projections for the total US population from 2000 until 2050. The two sets of estimates are always very close, and the relative difference in a given year between the two estimates is always less than 0.8%. Fig. B contains projections of annual deaths in the US population from 2000 to 2050. The model estimates are initially lower than national estimates, but model and Census projections of deaths converge to a difference of less than 0.5% by 2050.



**Figure A: US total population projections**

Figure Note: Data Source: US Census Bureau [1]

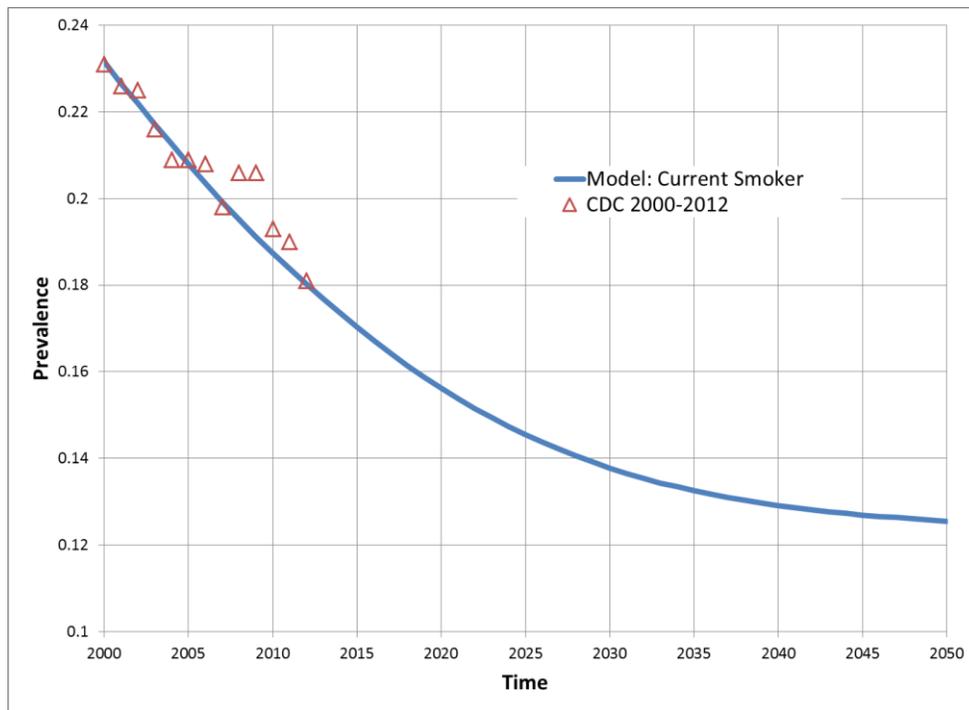


**Figure B: Projected US annual deaths**

**Figure Note:** Data Source: US Census Bureau [2]

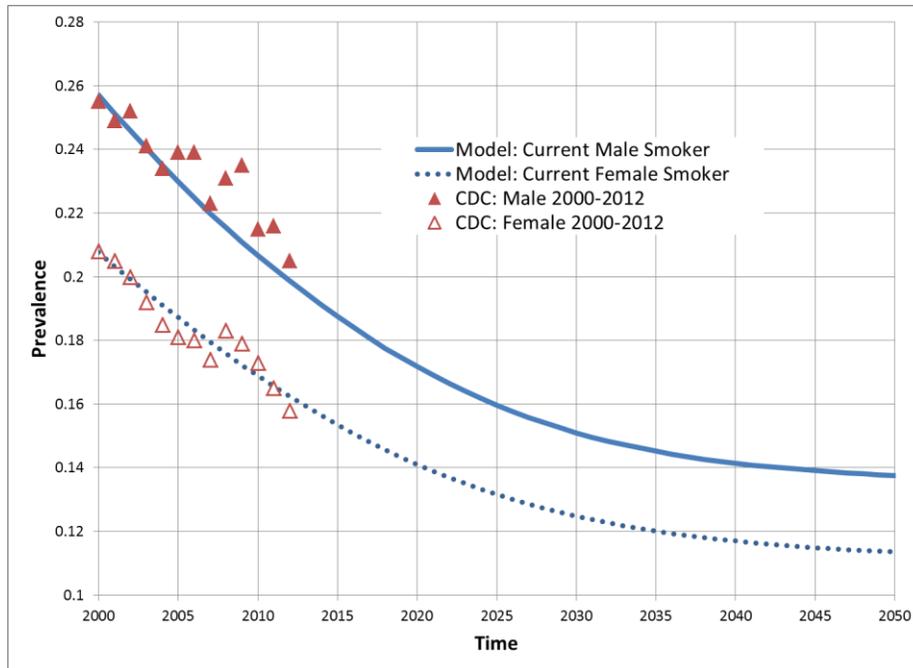
### C. S3.2.2. Smoking Prevalence

Fig. C contains model projections from 2000-2050 and CDC observed estimates from 2000-2012 for smoking prevalence among US adults ages 18 years and older. Model estimates for 2000-2012 are comparable in magnitude to CDC estimates and show a similar decline over time. Fig. D presents estimates of US adult smoking prevalence by sex. Again, model projections and CDC estimates are comparable in magnitude and show similar decreases over time.



**Figure C: US adult smoking prevalence**

**Figure Note:** Data source: CDC [3]



**Figure D: US adult smoking prevalence by sex**

**Figure Note:** Data source: CDC [3].

Table A shows estimated smoking prevalence by age group. Model projections are again similar to observed national estimates.

**Table A: US smoking prevalence by age**

	Year	2000	2005	2010
<b>18-44</b>	Model	0.270	0.245	0.227
	CDC	0.267	0.241	0.215
<b>45-64</b>	Model	0.240	0.213	0.189
	CDC	0.237	0.219	0.211
<b>65-74</b>	Model	0.129	0.125	0.116
	CDC		0.111	0.130
<b>75+</b>	Model	0.058	0.052	0.039
	CDC		0.058	0.051
<b>65+</b>	Model	0.095	0.088	0.079
	CDC	0.096		

**Table Note:** Data source: CDC [3]. In 2000 and 2001, CDC reported smoking prevalence for ages 65 years and older. In 2003, CDC changed their reporting format to report smoking prevalence for ages 65 to 74 years and 75 years and older.

#### **D. S3.2.3. Smoking-Attributable Mortality**

Our model can be used to estimate smoking-attributable mortality (SAM) and the smoking-attributable fraction of deaths in the US population. Most estimates of current smoking-attributable mortality for the US are generally in the range of 400,000 to 500,000 deaths per year [4], [5], [6], [7]. Table B lists smoking-attributable mortality estimates from the model and from the CDC [5] for the years 2000 to 2004 and 2005 to 2009.

Estimates from our model and the CDC are generally comparable. The CDC updated the data and methodology that it uses to estimate smoking-attributable mortality for 2005 to 2009, thus explaining the increase in smoking-attributable deaths from the earlier period in its estimates. The CDC previously used relative risks obtained from American Cancer Society Cancer Prevention Study II (CPS-II) data in its estimation of smoking-attributable mortality for the US [4]. CPS-II was a large longitudinal cohort study with over one million participants that was initiated in 1982. The CDC used mortality follow-up from the CPS-II from 1982 to 1988 in its Smoking-Attributable Mortality, Morbidity, and Economic Costs (SAMMEC) methodology [8]. Research has shown that relative risks for smoking have changed since this time and that risks for women have come to be similar to those for men [9]. The CDC updated its data and methods for estimating smoking-attributable mortality in the 2014 Surgeon General's Report on the health effects of smoking in producing its estimates of smoking-attributable mortality for the US from 2005 to 2009 [5]. The estimates are obtained from relative risks from mortality follow-up for a pooled analysis of several more recent cohort studies, and the estimates are calculated for more specific age groups for some causes of death. Our method and the CDC method estimate smoking-attributable mortality in somewhat different ways, given that we estimate smoking-attributable mortality from all-cause mortality whereas the CDC estimates smoking-

attributable mortality for a set of smoking-related causes, but the two sets of estimates are generally consistent. We use relative risks obtained from NHIS-LMF data, given that this data source provides recent, nationally representative estimates from data collected and made publicly available by the CDC's National Center for Health Statistics, the federal government's principal health statistics agency. Our estimates for US smoking-attributable mortality from 2000-2009 are consistent with published estimates for this period calculated with NHIS-LMF relative risks using the SAMMEC methodology previously used by CDC [6].

Table B also includes estimated smoking-attributable fractions of deaths for the US, which is calculated by dividing the number of smoking-attributable deaths by the total numbers for an age group in a time period. Model estimates are again comparable to CDC estimates and the smoking-attributable fractions of deaths reported by researchers using a variety of estimation methods (see Table 4 in Fenelon and Preston [10] for a collection of different approaches and corresponding estimates of smoking-attributable fraction of deaths).

**Table B: Smoking-attributable mortality and smoking-attributable fraction of deaths: ages 35 years and older**

Year	Total Smoking-Attributable Mortality		Average Annual Smoking-Attributable Mortality <sup>+</sup>		Average Annual Smoking-Attributable Fraction of Deaths	
	Model	CDC <sup>*,**</sup>	Model	CDC	Model	CDC <sup>+,**</sup>
2000-2004	2,080,000	1,960,000	416,000	392,000	0.18	0.17
2005-2009	1,950,000	2,190,000	390,000	438,000	0.16	0.19

\*Data Source: US Department of Health and Human Services [5].

\*\*Model projections of smoking-attributable mortality do not include deaths from secondhand smoke exposure and residential fires and perinatal deaths. For purposes of comparison, the CDC estimates reported in the table do not include these categories of deaths as well.

<sup>+</sup>Average Annual Smoking-Attributable Mortality = (Total Smoking-Attributable Mortality)/5 years

<sup>++</sup>Average Annual Smoking-Attributable Fraction of Deaths =  
 (Smoking-Attributable Mortality in 5 year period)/(Total Number of Deaths in 5 year period)

<sup>+++</sup>Data source for total number of deaths: CDC [11].

### S3.3 Conclusion

Model projections for the US are similar to national estimates and projections for population size, deaths, cigarette smoking prevalence, and smoking-attributable mortality.

### S3.4 References

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