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German EstSmoke: Estimating adult smoking-related costs and consequences of smoking cessation for Germany

Running head: German EstSmoke: Smoking-related costs in Germany

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10

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12

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24 The authors have no conflict of interest to disclose. The authors have no financial relationships relevant to this
25 article to disclose.

26 **Abstract**

27 **Aims.** We compared predicted lifetime health care costs for current, never and ex-smokers in
28 Germany under the current set of tobacco control polices. We compared these economic
29 consequences of the current situation with an alternative in which Germany were to
30 implement more comprehensive tobacco control policies consistent with the WHO
31 Framework Convention for Tobacco Control (FCTC) guidelines.

32 **Design.** German EstSmoke, an adapted version of the UK EstSmoke simulation model, applies
33 the Markov modelling approach. Transition probabilities for (re-)currence of smoking-related
34 disease were calculated from large German disease-specific registries and the German Health
35 Update (GEDA 2010). Estimations of both health care costs and effect sizes of smoking
36 cessation policies were taken from recent German studies and discounted at 3.5%/year.

37 **Setting.** Germany

38 **Participants.** German population of prevalent current, never and ex-smokers in 2009:
39 81 million

40 **Measurement.** Lifetime cost and outcomes in current, never and ex-smokers

41 **Findings.** If tobacco control policies are not strengthened, the German smoking population
42 will incur €41.56 billion lifetime excess costs compared to never smokers. Implementing
43 tobacco control policies consistent with WHO FCTC guidelines would reduce the difference
44 of lifetime costs between current smokers and ex-smokers by at least €1.7 billion.

45 **Conclusions.** Modelling suggests that the lifetime healthcare costs of people in Germany who
46 smoke are substantially greater than those of people who have never smoked. However, more
47 comprehensive tobacco control policies could reduce healthcare expenditure for current
48 smokers by at least 4%.

49

50

51 **Keywords:** smoking cessation, lifetime health care costs, smoking-attributable diseases, cost
52 savings, tobacco control policies, WHO FCTC

53

54 **What is already known about this topic**

- 55 • Since health care expenditures increase with age and smoking-related conditions have
56 long latency periods, lifetime estimations of cost are essential to make informed
57 decisions on resource allocation.
- 58 • Compared to never smokers, current smokers have higher lifetime and annual health
59 care expenditures.

60

61 **What important knowledge gaps exist on this topic**

- 62 • Dynamic modelling frameworks that estimate lifetime benefits of smoking cessation
63 depending on the number of years since quitting.
- 64 • Estimations of lifetime economic consequences associated with smoking cessation if
65 Germany implements stronger, more comprehensive or new tobacco control policies
66 consistent with WHO FCTC guidelines.

67

68 **What this study adds**

- 69 • The number of years since quitting strongly influences mortality and morbidity risks
70 in ex-smokers and the magnitude of cost savings over lifetime.
- 71 • Even tobacco control policies with low effect sizes (3%-4%) result in sizeable future
72 cost savings that outweigh implementation costs.
- 73 • German EstSmoke supports public policy planners in making informed decisions how
74 to best allocate resources to implement both federal and state tobacco control policies.

75

76

77 **1. Introduction**

78 Since smoking is the leading cause of chronic and cost-intensive illnesses, the World Health
79 Organisation (WHO) set out the WHO Framework Convention for Tobacco Control (FCTC)
80 to reduce the health, social and economic consequences of smoking (1). To assist
81 governments to implement the WHO FCTC in a timely and effective manner, WHO
82 introduced MPOWER, a tool to measure tobacco control policy performance based on six
83 policies. These are: 1) monitoring tobacco use and prevention policies, 2) protecting people
84 from tobacco smoke, 3) offering help to quit tobacco use, 4) warning about the dangers of
85 tobacco, 5) enforcing bans on tobacco advertising, promotion and sponsorship, and 6) raising
86 taxes on tobacco (2). However, uniform implementation and application of these policies is
87 currently lacking in the European Union. A large disparity among European countries has
88 been observed by the most recent estimations of the Tobacco Control Scale 2013 (TCS),
89 which quantifies the implementation of tobacco control policies at country level. While
90 countries like the UK have implemented a comprehensive set of tobacco control policies,
91 Germany has been ranked among the most inactive countries in terms of tobacco control (3,
92 4). Specifically, Germany has largely failed in the TCS categories of “bans on advertising and
93 promotion”, “large health warning labels” and “treatment to help smokers quit”. Indeed,
94 tobacco advertising is only prohibited on TV, on the radio and in most print publications
95 (based on 2003/33/EU (5)). In contrast, point-of-sale, billboard and outdoor advertising and
96 promotion are allowed (3). Moreover, cessation treatment policies like pharmacotherapies and
97 behavioural interventions involving advice and support by health care providers are not
98 covered by the statutory health insurance in Germany. More importantly, Germany has failed
99 to introduce stronger, more comprehensive or new tobacco control policies following the
100 WHO FCTC. This is mainly because both state and federal governments are responsible for
101 implementing tobacco control policies, resulting in unclear regulations and numerous
102 loopholes (6). This lack of clear jurisdiction may have led to Germany’s current smoking
103 rates, which are stagnating at a high level after decades of decline (3, 7). Moreover, the high
104 prevalence of smoking in Germany imposes substantial health-related economic costs on the
105 country’s health care sector (8) (9) (10). In particular, a German study (8) reported 20%
106 higher annual costs in 2009 for smokers than for never smokers, which amounts to €17.9
107 billion per year. However, the long-term health and economic consequences of Germany’s
108 inactivity are unclear. Similarly, the potential cost savings that Germany would incur if it
109 implemented tobacco control policies in alignment with WHO FCTC guidelines are still not
110 known.

111 In fact, the only study (11) to date to consider the long-term health consequences of current
112 German tobacco control policies has estimated a decline of smoking prevalence by 17%,
113 which fails to meet WHO FCTC guidelines. However, this study does not consider the long-
114 term economic consequences on the German health care sector. Only a few international cost-
115 of-illness (COI) studies (12) (13) (14) (15) have investigated the long-term economic
116 consequences of smoking in the US and European countries - and their results differ. While
117 the majority of these studies found higher average lifetime health care costs in smokers and
118 ex-smokers compared to never smokers, the study by Barendregt et al. (13) found the
119 opposite. It showed that higher health care costs later in life outweigh the economic benefits
120 of smoking cessation (13). However, this study assumed that costs for all diseases other than
121 heart disease, stroke, chronic pulmonary disease (COPD) and cancers are not related to
122 smoking and does not consider that smokers are also at a higher risk for other cost-intensive
123 diseases (16). Due to differences in the prevalence of smoking, the related costs of health
124 services and the implementation of tobacco control policies in European countries, the
125 economic results of international COI studies are not easily transferable to Germany.
126 Therefore, the primary aim of this study is to predict lifetime health care costs of current
127 smokers, never smokers and ex-smokers in Germany. A second objective is to estimate
128 lifetime economic consequences if Germany strengthens its efforts in implementing WHO
129 FCTC policies, such as marketing bans, health warnings and cessation treatment.

130

131 **2. Method**

132 In this subsection, we discuss the model parameters, modelling approach and design of
133 German EstSmoke.

134

135 **2.1. Design of German EstSmoke**

136 German EstSmoke is an adapted version of the UK EstSmoke, which estimates the lifetime
137 health care cost of smoking in the UK by using Markov modelling (12). Markov modelling is
138 a cohort simulation approach commonly used to model the clinical development of the risks
139 and trajectories of chronic diseases over the course of a lifetime (17). This is particularly
140 relevant for the progression of smoking-related diseases, which are characterised by long
141 latency periods after smoking onset. German EstSmoke was developed in Microsoft Excel
142 2010 (Microsoft Corporation, Redmond, WA, USA). While Monte Carlo Simulations were

143 programmed in Visual Basic for Application (VBA), further statistical analyses were
144 conducted using Stata version 12.1 (StataCorp. 2011, College Station, TX: StataCorp LP.).

145 German EstSmoke focuses on the four smoking-related conditions known to have the highest
146 economic and health-related consequences associated with smoking (18, 19): myocardial
147 infraction (MI), stroke, COPD and lung cancer. To model their lifetime trajectories depending
148 on smoking status, we created three base cohorts, one each for current smokers, never
149 smokers and ex-smokers. Modelling clinical pathways separately for each smoking status is in
150 line with other smoking models and a common approach to avoid influences from individual
151 smoking histories (14, 20). In the following, we examine briefly each clinical pathway.

152 As shown in Figure 1, the base cohort includes healthy adults, i.e. they do not have any of the
153 four clinical conditions explicitly modelled here. During a Markov cycle (here one year),
154 individuals in the base cohort may develop a first-ever acute MI event. Such an event may
155 either be fatal or non-fatal. If fatal, individuals move to the absorbing state “fatal MI”; if non-
156 fatal, they move to the state “non-fatal MI (year 1)”. In the next cycle, individuals may move
157 to the state “non-fatal MI (year 1+)”, have another acute MI event or die from other causes.
158 Beyond year 1 post-MI, individuals may remain in the same health state, have another acute
159 MI or die from other diseases. Individuals who have a second acute MI will follow the same
160 pathway as the first acute MI. Figure 1 shows that the stroke pathway is similar to the MI
161 pathway. By contrast, the COPD pathway has a chronic course and often leads to death.
162 Similarly, once individuals have developed lung cancer, they tend to die from the cancer itself
163 or an opportunistic disease. The model incorporates the increased risk of mortality from lung
164 cancer in individuals with COPD.

165 We simulated each cohort of current smokers, never smokers and ex-smokers from age 35
166 until the age 100 to estimate costs and outcomes for each clinical pathway, stratified by sex.
167 Lifetime excess costs of smoking per person were estimated as the difference in projected
168 lifetime costs between a 35-year-old (continuing) smoker and a 35-year-old never smoker.
169 Similarly, the economic consequences of smoking cessation were calculated as the difference
170 in lifetime costs between a 35-year-old smoker and a 35-year-old ex-smoker at given age of
171 quitting for varying smoking cessation rates. The model starts at the age of 35 for three major
172 reasons. Firstly, the absolute risks of developing or dying from MI, stroke, lung cancer and
173 COPD are very low in the general population of adolescence and young adulthood (21).
174 Secondly, the increased relative risk of smoking-attributable conditions accumulates over
175 time, i.e. it has long latency periods and increases with the number of smoking years. Hence,

176 this relative risk is low below the age of 35. Thirdly, the impact of smoking on health care
177 costs is negligible in younger ages due to the long latency periods of smoking-related diseases
178 (15, 20). Finally, modelling the economic consequences of the general adult population (71%
179 of the total population in Germany is ≥ 35 years) is in line with other model-based economic
180 simulations (22).

181

182 **2.2 Model Parameters**

183 German EstSmoke uses data on smoking prevalence obtained from the German Health Update
184 (GEDA 2010), which is a nationally representative health survey of 25,000 German adults (age
185 >18) (23) (Table 1). Estimations of mortality rates, incidence rates and relative risks of
186 smoking-related events are taken from the literature, as described below. We evaluated the
187 quality of these data using the STROBE checklist for observational data (24) and the
188 CONSORT checklist for randomised controlled trials (25).

189

190 State transition probabilities for MI, stroke, lung cancer and COPD

191 For state transition probabilities of smoking-related diseases in the general population, we
192 used population-based data from nationally representative German registries for i) MI (26), ii)
193 stroke (27), iii) lung cancer (28) and iv) COPD (29). Following Briggs et al. (30), one-year
194 probabilities of first-ever acute MI, stroke, lung cancer and COPD in the general population
195 were derived using these data (Appendix 1). Since these data include all three smoking-
196 related groups, i.e. current, never and ex-smokers, we subsequently calculated state transition
197 probabilities for each group (Appendices 2a-2d) by applying the previously described
198 approach in our companion paper (12). Specifically, transition probabilities for first-ever MI,
199 stroke, lung cancer and COPD for each group were calculated using relative risk estimations
200 from i) the INTERHEART study (31, 32), which investigated the risk factors associated with
201 first-ever acute MI based on data from 52 countries, including Germany, ii) the two largest
202 and longest running studies of risk factors associated with primary stroke in women (Nurses'
203 Health Studies) and men (Health Professional Follow-up Study) (33), iii) the SYNERGY
204 project (34) using lung cancer data from eight European case-control studies in 11 countries,
205 including Germany, between 1985-2005 and iv) the European Community Respiratory Health
206 Survey (ECRHS), which examines incidence rates for COPD for 12 countries, including
207 Germany (29) (Table 2).

208 Similar to transition probabilities of first-ever events, we calculated transition probabilities of
209 recurrent MI and stroke among current, never and ex-smokers based on two nation-wide
210 German registries (recurrent MI) (35) and a large German cohort study (recurrent stroke) (36)
211 (Table 2). Beyond year 1, the probability of acute MI or stroke was based on a systematic
212 review by Lip and Kalra (2010) (37). Finally, for state transition probabilities of fatal events
213 (Appendix 3), we used population-based data from i) the MONICA/KORA MI Registry (fatal
214 MI) (26), ii) Erlangen Stroke Project (ESPrO) (fatal stroke) (27), iii) the Association of
215 Population-based Cancer Registries in Germany (GEKID Atlas) (death from lung cancer) (28)
216 and iv) the WHO health statistics (death from COPD) (38). Since patients with lung cancer or
217 COPD can die not only from these diseases but also from comorbidities like asthma, liver
218 disease and previous metastatic cancer, we rescaled the probability of death as previously
219 described in our companion paper (12).

220

221 State transition probabilities of death due to other diseases

222 Similar to disease pathways, we calculated transition probabilities of death from other
223 diseases based on the most recent population life tables for Germany (39). Since these life
224 tables include deaths caused by the four diseases modelled here, we excluded such deaths
225 (Appendix 3). This was taken as mortality due to other diseases in the general population and
226 used to estimate the risk of mortality among never smokers by applying smoking-attributable
227 fractions based on the most relevant German study (21). Finally, since smoking increases the
228 risk of mortality due to other causes, we adjusted the risk of death from other causes for
229 smokers and ex-smokers by using relative risk estimations for mortality for each group (40,
230 41).

231

232 Health care costs for smoking-related diseases and mortality costs from other causes

233 For the MI and stroke pathways, we considered costs in the acute phase, as well as in the first
234 and subsequent years (year 1+) (Table 2). All estimates of costs were used from two recent
235 and most relevant German bottom-up COI studies (42, 43). In contrast to other German COI
236 studies, these studies provide detailed medical costs per patient after MI and stroke. For the
237 lung cancer pathway, both health care costs per patient and the costs of initial and terminal
238 care were used from the most recent German COI study (44) (Table 2). Since health care
239 costs of COPD vary depending on the severity of the disease, we included annual costs of

240 COPD and costs of death due to a severe exacerbation of COPD. Data were taken from two
241 German studies (45, 46), which include detailed in- and outpatient COPD costs per patient
242 depending on COPD severity grades. Lastly, although health care costs from other causes
243 were not considered, mortality costs from other causes were calculated using the most relevant
244 evaluation report by the German Scientific Advisory Committee (47), which is based on the
245 nationally representative health insurance data of 4.8 million individuals covered by statutory
246 health insurance in Germany. Since the data include mortality costs due to smoking-related
247 diseases, these mortality costs were excluded. All costs were indexed to year 2015 euros (€).

248

249 Sensitivity and scenario analyses

250 Probabilistic sensitivity analyses (Monte Carlo Simulations) were used to test the robustness
251 of our results. Specifically, we randomly sampled parameter values from statistical
252 distributions defined by standard error or confidence interval estimates in the literature. We
253 ran 10,000 simulations for the probabilistic sensitivity analysis (PSA) (48). Commonly used
254 statistical distributions were implemented for relative risks (log normal distribution) and costs
255 (gamma distribution) (49). A uniform distribution was used only when the parameter range
256 was available without standard errors. The 95% confidence intervals based on the PSA are
257 reported in the Results section (48).

258 For scenario analyses, we assumed that Germany strengthens its efforts to implement the
259 WHO FCTC policies that had the lowest ranking in TCS. Specifically, we evaluated an
260 increase of smoking cessation rate by i) 3% (strong health warnings), ii) 4% (comprehensive
261 marketing bans) and iii) 34% (cessation treatment such as pharmaco- and behavioural
262 therapies) (11). Since Germany implemented a telephone quitline in 1999, we followed Levy
263 et al. (11) and assumed a 34% increase in smoking cessation (instead of 39% if no cessation
264 treatment policy had been implemented).

265

266 **3. Results**

267 We estimated the magnitude of lifetime health care costs of four smoking-related diseases for
268 current smokers, never smokers, and ex-smokers using a cohort modelling approach. Our
269 analysis projected four clinical pathways and associated costs and then estimated potential
270 economic consequences of smoking cessation if Germany strengthens efforts in implementing
271 WHO FCTC policies that had the lowest ranking in TCS.

272

273 Lifetime health care costs for smoking-related diseases

274 Table 3 presents the lifetime health care costs due to MI, stroke, lung cancer and COPD for
275 current smokers, never smokers, and ex-smokers. Smokers have significantly higher lifetime
276 health care costs than never smokers and ex-smokers. Compared to never smokers, the
277 lifetime health care costs for both male and female smokers are twice as high. For the current
278 smoking population in Germany lifetime excess health care costs of smoking were €41.56
279 billion (2015, discounted at 3.5%). Moreover, compared to ex-smokers, the lifetime health
280 care costs for smokers are 20% higher for males and 30% higher for females (Table 3).

281 Figure 2 presents the distribution of cumulative health care costs incurred by smokers and ex-
282 smokers by age. It shows that the difference in incurred costs is much more pronounced after
283 the age of 65 years before it stabilises at the age of 80. This variation in costs is likely due to
284 the higher probability of using medical services as smoking-related diseases become clinically
285 relevant in older individuals. Indeed, Figure 3 shows that smoking-attributable deaths increase
286 with age, particularly between the ages of 55 and 85. For example, while 9% of all deaths are
287 attributable to smoking-related diseases in males aged 35-45, 28% of all deaths are
288 attributable to smoking-related diseases in males aged 70-74. Finally, our results confirm that
289 MI and stroke are the top contributors to the overall smoking burden over lifetime, which
290 indicates potential sizeable reductions in costs as MI and stroke events can be prevented (see
291 Appendix 4).

292

293 Potential economic consequences of implementing WHO FCTC policies

294 Table 3 shows the magnitude of the difference in costs between smokers and ex-smokers if
295 Germany implements strong health warnings, comprehensive marketing bans or cessation
296 treatment policies, which are in line with WHO FCTC guidelines. We found that a 3%
297 increase of smoking cessation (which is potentially achievable due to strong health warnings
298 (11)) would reduce the lifetime cost-difference between smokers and ex-smokers by €1.7
299 billion (discounted). Similarly, this potential difference would decline by €2.2 billion if
300 smoking cessation rates increase by 4% due to an implementation of comprehensive
301 marketing bans (11). Finally, if Germany implements further cessation treatment policies like
302 pharmaco- and behavioural therapies, we predict a reduction in cost-difference by €18.9
303 billion (discounted) over a lifetime.

304

305 **3. Discussion**

306 Twelve years after the introduction of the WHO FCTC, Germany still lacks comprehensive
307 tobacco control policies and has implemented only a minimal set of policies (3). By using
308 German EstSmoke, we projected that lifetime health care costs of smoking were twice as high
309 in smokers than in never smokers under Germany's tobacco control policies. Moreover, if it
310 does not strengthen its policies, Germany will impose substantial lifetime excess health care
311 costs of smoking on the population level (€41.56 billion, discounted). These costs underscore
312 the economic and public health relevance of smoking in Germany. More importantly, using
313 German EstSmoke we showed that small effect sizes of tobacco control policies (3%-4%)
314 result in sizeable cost-differences (€1.7-2.2 billion) compared to programme costs, which vary
315 between €0.008 billion (50) and €0.4 billion (51). Using German EstSmoke, we also
316 demonstrated that larger potential cost savings can be achieved with the help of pharmaco-
317 and behavioural therapies (€18.9 billion, discounted). Therefore we urge swift
318 implementation of stronger health warnings, more comprehensive advertising bans and new
319 cessation therapies, all covered by statutory health insurance, as major components of an
320 effective tobacco control policy in Germany.

321 A recent systematic literature review (52) provides strong evidence that only few international
322 studies have also estimated lifetime costs of smoking using a dynamic modelling framework.
323 For instance, Hodgson (15) estimated that compared to never smokers, male (female) smokers
324 have a 47% (41%) higher lifetime cost, which is in line with our results. However, it should
325 be noted that Hodgson's approach (15) to estimating lifetime medical expenditures was based
326 on smoking status (not on clinical pathways, as in our model). A similar approach was applied
327 by the European study on Quantifying Utility of Investment in Protection from Tobacco
328 (EQUIPT) (53), a recent multicentre study which evaluates the cross-context transferability of
329 economic evidence of tobacco control policies among European countries. For Germany,
330 annual health care costs of smoking-attributable diseases (MI, stroke, COPD, lung cancer)
331 were estimated at €15.12 billion (54) which are slightly higher than our estimates
332 (€13.26 billion). This cost difference is mainly due to methodological differences. For
333 instance, the prevalence of smoking-attributable diseases for smokers and ex-smokers was
334 calculated by using population-attributable fractions in the EQUIPT study (53, 55), compared
335 to modelling clinical pathways for each disease. Moreover, morbidity and mortality of ex-
336 smokers depending on the number of years since quitting was modelled in German EstSmoke

337 while an average risk for ex-smokers was assumed in the EQUIPT model. With respect to the
338 economic consequences of smoking cessation, our results are consistent with the results of
339 other COI studies (52) but cannot be compared with cost-effectiveness models like the
340 frequently used Benefits of Smoking Cessation on Outcomes (BENESCO) model, which
341 assesses the cost-effectiveness of smoking cessation pharmacotherapies in the US and
342 European countries. All COI studies, except that of Barengregt et al. (13), found that smoking
343 cessation will result in short- and/or long-term cost savings. However, Barengregt et al. (13)
344 concluded that smoking cessation reduces only short-term costs while increasing long-term
345 costs. This conclusion can be challenged for a number of reasons. Firstly, smoking increases
346 the risk of more expensive health conditions (such as lung cancer), which otherwise have
347 lower incidence rates. Secondly, smoking increases the risk of chronic non-fatal conditions
348 that incur ongoing medical costs (such as COPD and asthma). Indeed, our model showed that
349 between ages 50 and 80, there is a higher proportion of the smoker cohort with at least one of
350 the four diseases than the never-smoking cohort. For instance, at age 70, 16% of smokers are
351 alive with at least one smoking-related disease compared to 7% of never smokers. Thirdly,
352 Barengregt et al. (13) restricted their analysis to four conditions and did not include the
353 increased relative risk of other conditions, many of which are non-fatal. Finally, since
354 smoking increases the early onset risk of diseases (including long-term chronic conditions and
355 expensive fatal conditions such as lung cancer), costs incurred earlier in life are valued more
356 than costs incurred later in life when discounting is applied. This impact results in higher
357 discounted costs than those incurred later in life. However, Barengregt et al. (13) did not
358 apply discounting to their COI model. Similar arguments were raised by other authors in
359 response to this paper (see comments of Hodgson; Fries; Heaney; Leistikow and Miller, and
360 Sauter 1998 (16)).

361 Our study has a number of strengths. It provides the first model, German EstSmoke, which is
362 capable of estimating lifetime costs of smoking in Germany. Indeed, modelling clinical
363 pathways which depend on smoking status over the course of a lifetime helps clinicians and
364 health care specialists to make informed decisions on allocating scarce resources based on
365 both clinical and economic relevance. For instance, MI not only has a high clinical relevance
366 but also causes high lifetime health care costs. More importantly, since political decisions are
367 made within four-year electoral cycles, German EstSmoke is a useful economic tool in
368 decision-making processes: it predicts the economic consequences of tobacco control policies
369 not merely for a specific year but for longer timeframes. German EstSmoke can be thus used
370 to predict long-term economic consequences of a single or combined tobacco control policy at

371 federal or state level. Ranking these consequences provides an important basis for prioritising
372 tobacco control policies according to cost savings. However, we must bear in mind that the
373 implementation of these policies is under the purview of the 16 German states, which enact
374 the WHO FCTC guidelines differently. Thus our model may also help to compare both
375 performances among the states and the economic consequences of inaction at the federal and
376 state level. Another strength of our study is its incorporation of time-dependent transitions
377 within a comprehensive decision analytic framework. In contrast to previous studies (15) (13),
378 our model takes into account that smoking cessation benefits depend on the number of years
379 since quitting. This, in turn, allows a more accurate estimation of future gains from smoking
380 cessation.

381 Since the limitations of the modelling approach are extensively discussed in earlier work (12),
382 we briefly summarise them and focus here on their implications. Firstly, since smoking
383 increases the risk of chronic (non-)fatal conditions such as asthma, modelling only four
384 clinical conditions according to smoking status, although in line with earlier studies (56) (57)
385 (14) (58), underestimates the costs of smoking-related diseases and the benefits of smoking
386 cessation. Similarly, since good quality epidemiological data about the impact of smoking
387 relapse on the risk of morbidity and mortality of smoking-related conditions are not available,
388 our results are rather conservative. Finally, since German EstSmoke does not consider both
389 the costs of implementing tobacco control policies and the impact of these policies on clinical
390 outcomes (e.g. reduced prevalence of smoking-related diseases), it is a partial economic
391 evaluation. In contrast to the BENESCO model (52), explicit investment choices cannot be
392 derived by using German EstSmoke.

393 In summary, by using German EstSmoke, we showed that the consequences of Germany's
394 inaction in implementing stronger, more comprehensive and new tobacco control policies are
395 considerable. Without the implementation of more rigorous policies, adult smoking-related
396 costs will substantially increase. Given that Germany is ranked among the most inactive
397 European countries in terms of tobacco control policies (3, 4), it can expect higher benefits
398 from policies to prevent or stop smoking, compared to countries which already meet WHO
399 FCTC guidelines. Indeed, our analysis shows that even small effect sizes (3-4%) can
400 substantially reduce the health-related costs of smoking in Germany. With the help of models
401 such as German EstSmoke, policy planners can make informed decisions to instigate effective
402 tobacco control policies.

403

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415

416 **Table and Figure Legend**

417 **Table 1:** Prevalence of never, current and ex-smokers by age group and sex

418 Age- and gender-specific prevalences of never smokers, current smokers and ex-smokers.

419

420 **Table 2:** Parameter values for Markov model and distribution of Monte Carlo Simulation

421 Epidemiological and economic input data for Markov model.

422

423 **Table 3:** Lifetime costs of health care resource use due to MI, stroke, lung cancer, COPD and
424 economic consequences of implementing WHO FCTC policies (2015)

425 Lifetime health care costs due MI, stroke, lung cancer, COPD, and other causes of death for
426 never smokers, current smokers, and ex-smokers. Smokers have significantly higher lifetime
427 health care costs than never smokers and ex-smokers.

428

429 **Figure 1:** Markov structure for four clinical pathways related to smoking and quitting
430 smoking

431 German EstSmoke focuses on four smoking-related conditions, i.e. myocardial infarction
432 (MI), stroke, chronic obstructive pulmonary disease (COPD) and lung cancer. Three base
433 cohorts, one each for never smokers, current smokers and ex-smokers, were created.

434

435 **Figure 2:** Cumulative distribution of lifetime health care costs in smokers and ex-smokers

436 Predicted lifetime health care costs incurred by smokers and ex-smokers, stratified by sex.
437 The difference in incurred costs is much more pronounced beyond the age of 65 before it
438 stabilises at the age of 80.

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440 **Figure 3:** Predicted survival of smokers and ex-smokers

441 Predicted survival curves for smokers and ex-smokers. They show that smoking cessation is
442 likely to increase the probability of survival in men and women, particularly between the ages
443 of 55 and 85 years.

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