

SPECIAL ARTICLE

21st-Century Hazards of Smoking and Benefits of Cessation in the United States

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ABSTRACT

BACKGROUND

Extrapolation from studies in the 1980s suggests that smoking causes 25% of deaths among women and men 35 to 69 years of age in the United States. Nationally representative measurements of the current risks of smoking and the benefits of cessation at various ages are unavailable.

METHODS

We obtained smoking and smoking-cessation histories from 113,752 women and 88,496 men 25 years of age or older who were interviewed between 1997 and 2004 in the U.S. National Health Interview Survey and related these data to the causes of deaths that occurred by December 31, 2006 (8236 deaths in women and 7479 in men). Hazard ratios for death among current smokers, as compared with those who had never smoked, were adjusted for age, educational level, adiposity, and alcohol consumption.

RESULTS

For participants who were 25 to 79 years of age, the rate of death from any cause among current smokers was about three times that among those who had never smoked (hazard ratio for women, 3.0; 99% confidence interval [CI], 2.7 to 3.3; hazard ratio for men, 2.8; 99% CI, 2.4 to 3.1). Most of the excess mortality among smokers was due to neoplastic, vascular, respiratory, and other diseases that can be caused by smoking. The probability of surviving from 25 to 79 years of age was about twice as great in those who had never smoked as in current smokers (70% vs. 38% among women and 61% vs. 26% among men). Life expectancy was shortened by more than 10 years among the current smokers, as compared with those who had never smoked. Adults who had quit smoking at 25 to 34, 35 to 44, or 45 to 54 years of age gained about 10, 9, and 6 years of life, respectively, as compared with those who continued to smoke.

CONCLUSIONS

Smokers lose at least one decade of life expectancy, as compared with those who have never smoked. Cessation before the age of 40 years reduces the risk of death associated with continued smoking by about 90%.

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SMOKING IS A MAJOR CAUSE OF PREMATURE death worldwide.¹⁻³ Despite substantial declines in the prevalence of smoking by adults, estimates based on extrapolation from studies in the 1980s suggest that for those between 35 and 69 years of age, smoking currently accounts for almost 200,000 deaths annually in the United States, or about one fourth of all deaths in this age group.⁴⁻⁶ The prevalence of smoking peaked around 1960 among U.S. men and about two decades later among U.S. women.^{7,8} Rates of death from vascular disease have decreased substantially since the 1980s owing to reductions in smoking and in other risk factors and to improved treatment.⁹ Therefore, extrapolation from previous studies is increasingly uncertain, and direct measurement of the contemporary hazards of smoking in the United States is needed. Moreover, a substantial number of adults have quit smoking in recent decades,⁸ affording an opportunity to study the effects of cessation at various ages on current mortality.

We report the results of a large, prospective, nationally representative study of smoking and its relation to mortality to provide estimates of the 21st-century hazards of smoking and of the benefits of smoking cessation at various ages for male and female smokers.

METHODS

STUDY DESIGN

We examined data from a cohort of 216,917 adults in the U.S. National Health Interview Survey (NHIS) between 1997 and 2004¹⁰⁻¹² that were linked to the National Death Index.^{13,14} The NHIS is a nationally representative cross-sectional health survey of the civilian, noninstitutionalized population of the United States. The survey uses a stratified, multistage sample design that permits representative sampling of households. One adult (≥ 18 years of age) is randomly selected from each selected household for a detailed interview on health and other behaviors. The NHIS sample is drawn from each state and the District of Columbia. Each year, approximately 35,000 households and 87,500 persons are newly enrolled in the survey. Black and Hispanic persons are deliberately oversampled, but the sample weights ensure that the final totals conform to national ethnic proportions. The NHIS sampling frame excludes only about 7 million adults (chiefly patients in long-

term care facilities, prisoners, and active-duty military personnel) from the total U.S. domestic population of 226 million adults in 2004.¹²

Mortality among survey participants through the end of 2006 was assessed by means of periodic matching of their records to the National Death Index,^{13,14} which includes death-certificate information for all deaths in the United States since 1986. Matching was performed for a combination of name, Social Security number, and date of birth, with a success rate exceeding 95%.

Enrollment rates for women exceeded those for men. A total of 122,810 women and 94,107 men 25 years of age or older participated in the NHIS between 1997 and 2004. Of these participants, 9058 women (761 of whom died) and 5611 men (544 of whom died) were excluded because of missing variables (e.g., educational level, smoking status, drinking status, or cause of death).

STATISTICAL ANALYSIS

Life-threatening illness can cause smokers to quit, which distorts the rates of death among current smokers and among those who have quit smoking recently in opposite ways. Unlike previous analyses of NHIS results,^{10,11} our analyses classified former smokers who had quit within 5 years before death as current smokers. Participants were classified as former smokers if they had smoked at least 100 cigarettes in their lifetime but had not smoked within the previous 5 years. Participants were classified as never having smoked if they had smoked fewer than 100 cigarettes in their lifetime.

We calculated hazard ratios for current and former smokers with the use of an age-stratified Cox proportional-hazards model,¹⁵ adjusted for educational level (less than high school, high school, or more than high school), alcohol consumption (nondrinker, former drinker, light drinker [1 to 2 drinks per day for women and 1 to 3 drinks per day for men] or moderate-to-heavy drinker [3 or more drinks per day for women and 4 or more drinks per day for men]), and adiposity (a body-mass index [the weight in kilograms divided by the square of the height in meters] of < 25 , 25.0 to 29.9, or ≥ 30). Further adjustment for race (black, Hispanic, or white) did not materially affect these estimates, and the results of these adjustments are not presented. All estimates were weighted according to the NHIS sample weights.¹²

Since the NHIS death rates are slightly lower than the U.S. national death rates,¹³ we scaled

our estimates of absolute death rates upward to match the U.S. national rates for 2004 by combining smoking prevalence, hazard ratios for smokers versus those who had never smoked, and total deaths, as described previously.¹⁶ From the age-specific death rates among current smokers and those who had never smoked, we calculated the cumulative probabilities of survival from 25 to 79 years of age, with adjustment for differences in age, educational level, alcohol consumption, and adiposity. The absolute differences in mortality between current smokers and those who had never smoked were, however, not materially altered by scaling the death rates (data not shown).

RESULTS

CHARACTERISTICS OF THE STUDY PARTICIPANTS

Current smokers more commonly consumed alcohol, had lower educational levels, and had a lower body-mass index than did former smokers or those who had never smoked (Table 1). About two thirds of current smokers and of former smokers had started smoking before 20 years of age. The proportion of former smokers increased steeply with age (Fig. 1), so that they were, on average, older than current smokers and those who had never smoked. Cessation of smoking was less common among women than among men; the ratio of former smokers to current smokers at 65 to 69 years of age was 2:1 for women but 4:1 for men.

MORTALITY ACCORDING TO SMOKING STATUS

Among 113,752 women and 88,496 men 25 years of age or older at yearly recruitment, who were followed for a mean of 7 years (1.3 million person-years), a total of 15,715 deaths (8236 deaths in women and 7479 in men) were recorded (see Table S1 in the Supplementary Appendix, available with the full text of this article at NEJM.org); 10,743 deaths (5122 in women and 5621 in men) occurred at 25 to 79 years of age. The hazard ratios for overall mortality at 25 to 79 years of age among current smokers versus those who had never smoked were 3.0 for women (99% confidence interval [CI], 2.7 to 3.3) and 2.8 for men (99% CI, 2.4 to 3.1) (Table 2). Adjustment for educational level, alcohol consumption, and adiposity had little effect on these age-stratified hazard ratios (data not shown). These age-specific hazard ratios, together with the prevalence of current smoking, were combined with 2004 U.S. death rates to es-

timate survival from 25 to 79 years of age for current smokers versus those who had never smoked, after adjustment for differences in age, educational level, and adiposity (body-mass index). This analysis showed that a person who had never smoked was about twice as likely as a current smoker to reach 80 years of age (Fig. 2). Among women, the estimated probability of survival to the age of 80 years was 70% (99% CI, 64 to 76) for those who had never smoked but only 38% (99% CI, 30 to 45) for current smokers (Fig. 2A). For men, these probabilities were 61% (99% CI, 55 to 67) for those who had never smoked but only 26% (99% CI, 18 to 33) for current smokers (Fig. 2B). Among current smokers, survival was shorter by about 11 years for women and by about 12 years for men, as compared with participants who had never smoked.

Table 2 shows that much, though not all, of this difference in survival was due to neoplastic disease, vascular disease, or other diseases that previous studies have shown can be caused by smoking.²⁻⁶ Notably high hazard ratios were observed for deaths from lung cancer and from ischemic heart disease in both women and men and from stroke in women. However, about 7% of the excess mortality among smokers at 25 to 79 years of age was due to accidents and injuries. In Table 2, only the excess mortality from disease (not from accidents and injuries) is attributed to smoking. At 25 to 79 years of age, about 62% (range, 59 to 65) of all deaths among female smokers and 60% (range, 55 to 63) of all deaths among male smokers would have been avoided if the rates of death from diseases among smokers had been the same as the rates among those who had never smoked and after adjustments for any differences in age, educational level, alcohol consumption, and adiposity.

BENEFITS OF SMOKING CESSATION

Hazard ratios for deaths from all causes among former smokers, as compared with those who had never smoked, were similar for women and men and were combined for statistical stability (Table S2 in the Supplementary Appendix). Lung cancer was associated with the highest hazard ratio for death among former smokers (Table S3 in the Supplementary Appendix). The effect of smoking cessation at younger ages was especially favorable (Fig. 3). For smokers who quit at 25 to 34 years of age (median, 29), survival curves were nearly

Table 1. Baseline Characteristics and Deaths in the National Health Interview Survey (NHIS) Cohort, According to Smoking Status.*

Variable	Women			Men		
	Current Smoker	Former Smoker	Never Smoked	Current Smoker	Former Smoker	Never Smoked
No. of participants surveyed	23,839	22,389	67,524	23,348	25,859	39,289
No. of deaths, according to cause†						
All causes	1798	2070	4368	2174	3335	1970
Lung cancer	289	163	83	362	274	52
All cancers	572	518	875	696	901	440
Vascular diseases	571	742	1978	698	1304	840
Respiratory diseases	248	276	272	222	385	105
No. of all-cause deaths at ages 25 to 79 yr	1579	1353	2190	2030	2308	1283
Mean age — yr	45.8	55.3	50.1	45.0	56.9	45.7
Educational level — no. of participants (%)‡						
<High-school graduate	5,438 (31.3)	3,968 (13.8)	14,675 (55.0)	5,835 (41.2)	5,643 (23.1)	6,145 (35.8)
High-school graduate	8,532 (29.2)	6,625 (18.2)	18,670 (52.6)	8,157 (34.4)	7,389 (26.1)	9,710 (39.5)
>High-school graduate	9,869 (17.0)	11,796 (21.5)	34,179 (61.5)	9,356 (19.8)	12,827 (26.3)	23,434 (53.9)
Alcohol consumption — no. of participants (%)‡						
Nondrinker	3,388 (11.5)	3,158 (7.4)	27,303 (81.0)	1,522 (13.9)	1,894 (11.7)	8,323 (74.5)
Former drinker	4,328 (25.8)	5,367 (25.3)	8,877 (48.9)	3,788 (27.8)	6,925 (33.3)	4,977 (38.9)
Light drinker§	10,410 (20.7)	11,768 (23.2)	27,364 (56.1)	11,496 (24.5)	14,272 (27.3)	21,486 (48.2)
Moderate-to-heavy drinker¶	5,713 (46.8)	2,096 (22.7)	3,980 (30.5)	6,542 (47.0)	2,768 (23.8)	4,503 (29.2)
Body-mass index — no. of participants (%)‡						
<24	12,343 (24.4)	9,796 (18.7)	30,965 (56.9)	9,137 (34.4)	6,988 (20.9)	12,023 (44.7)
25–29	6,549 (22.0)	6,727 (18.7)	20,158 (59.3)	9,744 (25.0)	12,194 (26.8)	18,236 (48.2)
≥30	4,947 (19.6)	5,866 (20.7)	16,401 (59.7)	4,467 (22.9)	6,677 (29.3)	9,030 (47.8)
Race — no. of participants (%)‡**						
White	17,075 (24.5)	17,647 (21.8)	40,972 (53.7)	15,972 (27.6)	19,952 (27.2)	25,498 (45.2)
Nonwhite	6,764 (17.3)	4,742 (12.3)	26,552 (70.4)	7,376 (26.7)	5,907 (21.2)	13,791 (52.2)
Age when participant started smoking — no. of participants (%)††						
<20 yr	15,726 (67.8)	14,330 (66.4)		16,925 (73.5)	19,431 (75.5)	
20–24 yr	4,592 (18.9)	4,370 (18.8)		4,092 (17.2)	4,042 (15.5)	
≥25 yr	3,521 (13.2)	3,689 (14.8)		2,331 (9.3)	2,386 (9.0)	
Cigarettes smoked per day — no. of participants (%)††						
<10	6,865 (26.1)			5,592 (22.0)		
10–19	7,223 (30.9)			5,502 (23.6)		
≥20	9,393 (41.6)			11,831 (52.7)		
Missing data	358 (1.4)			423 (1.7)		

* Women and men who were 25 years of age or older were included in the study cohort. This analysis excluded 5611 men (5%) and 9058 women (7%) for whom data on educational level, alcohol consumption, body-mass index, race, or smoking status were missing.

† These constituted deaths that occurred up to December 31, 2006. The *International Classification of Diseases, 10th revision*, codes were C33–34 for lung cancer, C00–97 for all cancers, I00–99 for vascular diseases, and J00–99 for respiratory diseases.

‡ Prevalence (expressed as percentages in parentheses) was weighted by the NHIS survey weights and standardized to the age distribution of current smokers for men and for women.

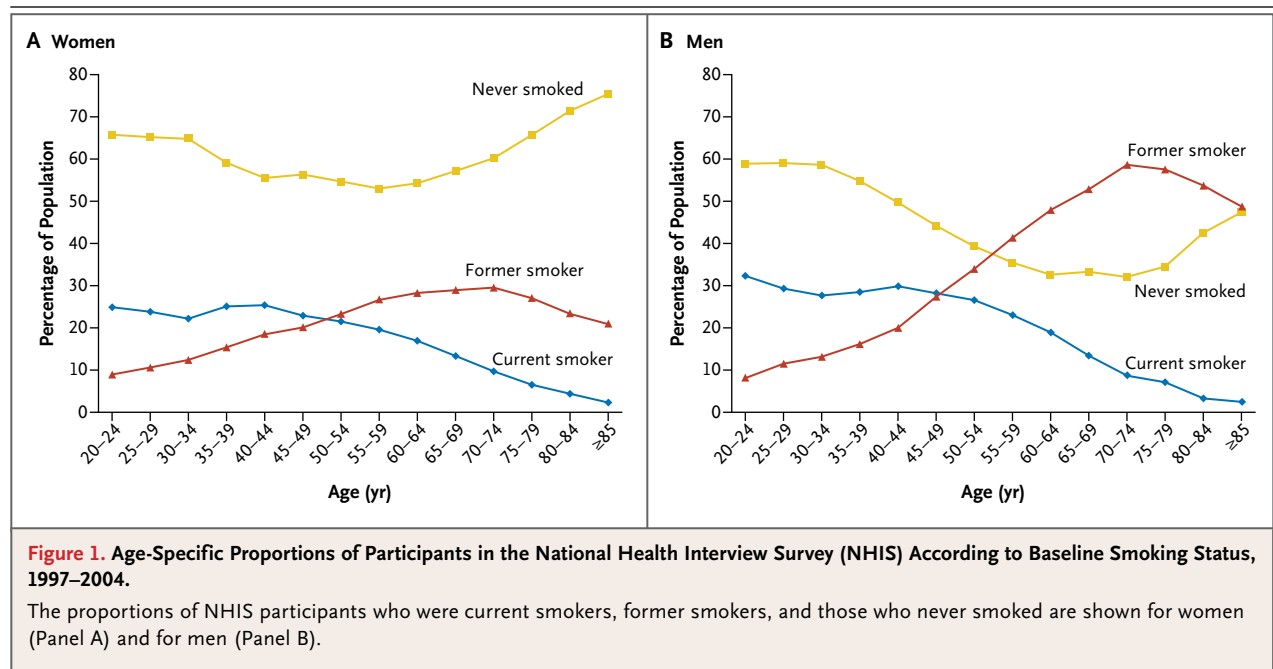
§ Light drinking was defined as 1 to 2 drinks per day for women and 1 to 3 drinks per day for men.

¶ Moderate-to-heavy drinking was defined as 3 or more drinks per day for women and 4 or more drinks per day for men.

|| The body-mass index is the weight in kilograms divided by the square of the height in meters.

** Race was self-reported.

†† Column proportions were weighted by the NHIS survey weights.



identical to those for participants who had never smoked, meaning that those who quit smoking gained about 10 years of life, as compared with those who continued to smoke. Survival curves were somewhat worse for smokers who had quit at 35 to 44 years of age (median, 39) than for those who had never smoked; still, smokers who quit smoking could expect to gain about 9 years of life, as compared with those who continued to smoke. Thus, cessation at about 39 years of age reduced the excess risk of death from any cause by about 90%. Nevertheless, smokers who had quit by about 39 years of age still had a 20% excess risk (hazard ratio, 1.2), as compared with those who had never smoked. Although this hazard is substantial, it is much smaller than the 200% excess risk (hazard ratio, 3.0) among those who continued to smoke (Fig. 4).

Smokers who stopped smoking at 45 to 54 years of age and those who stopped at 55 to 64 years of age (median, 49 and 59 years, respectively) gained about 6 and 4 years of life, respectively. Even cessation at the age of 45 to 54 years reduced the excess risk of death by about two thirds. Since some of the smokers who died might have quit smoking as a result of the disease that eventually killed them, the hazard ratios for former smokers might be biased upward (and the benefits of cessation might be even greater than shown). Exclusion of the first few years of follow-up would

help limit this “reverse causality,” but when we excluded the first 2 years of follow-up from the analysis, the results were similar (data not shown).

DISCUSSION

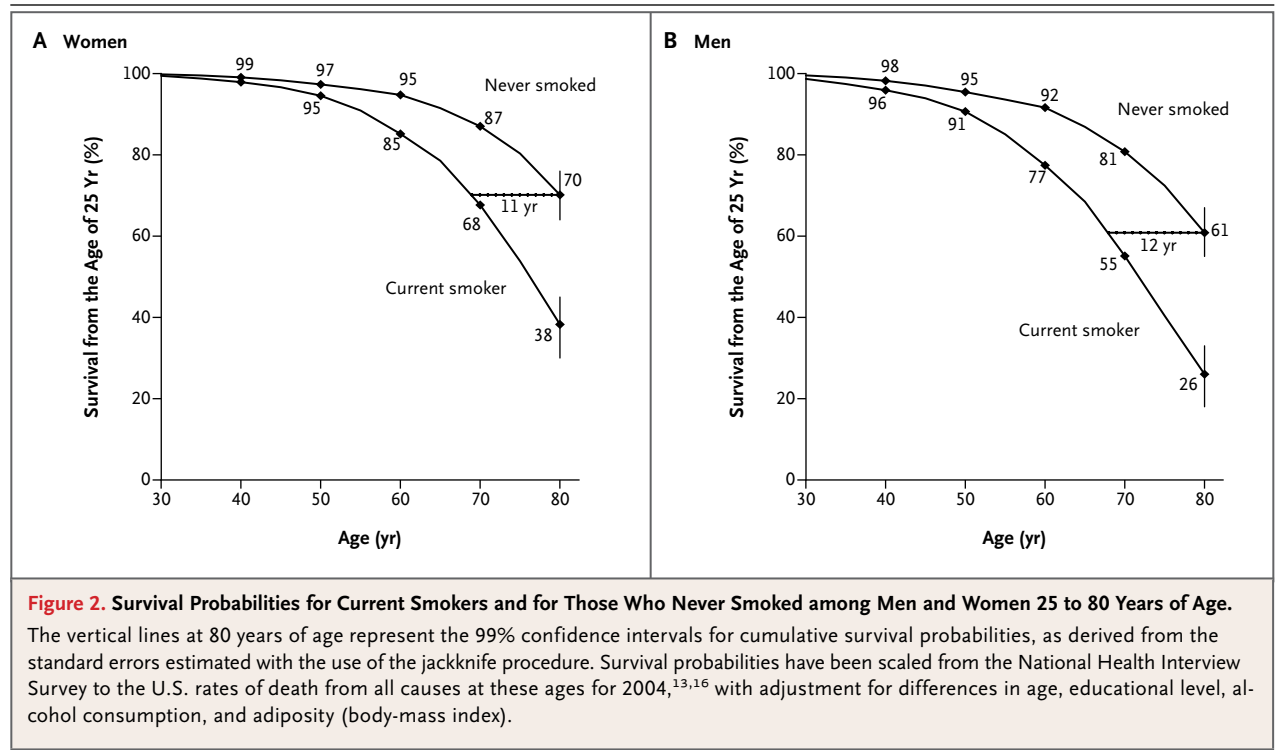
The overall mortality among smokers of both sexes in the United States is about three times as high as that among otherwise similar persons who never smoked, and the smokers lose, on average, at least a decade of life. The women in this cohort represent the first generation of women in the United States in which those who smoked began early in life and smoked for decades, and the risks of death for these women are about 50% greater than the risks reported in the 1980s studies.^{4,5} For both female and male smokers, the tripling of the relative risk of death and the reduction in survival by at least a decade are similar to the risks in four other studies: a study of male British doctors born between 1900 and 1930,^{17,18} the large U.K. study of women born between 1930 and 1950,¹⁹ a meta-analysis of several other U.S. cohort studies,²⁰ and a study in Japan of people born between 1920 and 1945.²¹ Although the relative risks were similarly tripled across the studies, the absolute death rates (for both current smokers and those who had never smoked) were much higher in our study than in other U.S. studies,²⁰ since the NHIS is more representative of

Table 2. Adjusted Hazard Ratios for Various Causes of Death among Current Smokers, as Compared with Those Who Never Smoked, among Women and Men 25 to 79 Years of Age.*

Cause of Death	Women				Men			
	Never Smoked	Current Smoker	Adjusted Hazard Ratio (99% CI)	Deaths Attributable to Smoking among Smokers	Never Smoked	Current Smokers	Adjusted Hazard Ratio (99% CI)	Deaths Attributable to Smoking among Smokers
	no. of deaths			no. (%)	no. of deaths		no. (%)	
Lung cancer	61	267	17.8 (11.4–27.8)	252 (94)	44	348	14.6 (9.1–23.4)	324 (93)
Cancers other than lung cancer	544	258	1.7 (1.4–2.1)	106 (41)	280	317	2.2 (1.7–2.8)	173 (55)
All cancers	605	525	3.2 (2.6–3.9)	360 (69)	324	665	3.8 (3.1–4.8)	491 (74)
Ischemic heart disease	382	251	3.5 (2.7–4.6)	179 (72)	285	416	3.2 (2.5–4.1)	288 (69)
Stroke	150	88	3.2 (2.2–4.7)	60 (69)	74	66	1.7 (1.0–2.8)	27 (40)
Other vascular disease	252	137	3.1 (2.2–4.4)	93 (68)	141	161	2.1 (1.5–3.0)	84 (52)
All vascular diseases	784	476	3.2 (2.7–3.9)	328 (69)	500	643	2.6 (2.1–3.2)	395 (61)
Respiratory diseases	119	206	8.5 (6.1–11.8)	182 (88)	45	188	9.0 (5.6–14.4)	167 (89)
Other medical disorders not shown above	581	277	2.2 (1.7–2.8)	151 (55)	295	370	2.2 (1.7–2.9)	205 (55)
All medical disorders	2089	1484	3.0 (2.7–3.3)	986 (66)	1164	1866	2.9 (2.5–3.2)	1211 (65)
Accidents and injuries	101	95	3.9 (2.4–6.2)	0	119	164	2.1 (1.4–3.0)	0
All causes†	2190	1579	3.0 (2.7–3.3)	986 (62)	1283	2030	2.8 (2.4–3.1)	1211 (60)

* Hazard ratios were adjusted for age, educational level, alcohol consumption, and body-mass index.

† Deaths attributable to smoking were determined with the use of the hazard ratios for all medical causes of death. With the exclusion of the 199 women and 222 men who had quit smoking less than 5 years before their deaths and the exclusion of the 1795 women and 2184 men who reported a history of coronary heart disease, stroke, or cancer, the hazard ratios for all-cause mortality were 3.1 for women and 2.8 for men.



the general U.S. population. Thus, in the NHIS, the large absolute differences in risk between current smokers and those who never smoked, as well as the gains in years of life for those who quit smoking, are likely to be true for the U.S. population as a whole. The hazards associated with smoking are substantial, even though in recent decades, most smokers in the United States have smoked cigarettes with relatively low levels of tar, as measured by machine testing, as compared with the cigarettes smoked in earlier decades.²²

Twenty-first-century measurement of the hazards of tobacco use must take into account the substantial reductions in mortality from vascular diseases and in overall mortality since 1970.^{5,9,23} As death rates among those who never smoked have fallen, the absolute differences in survival to the age of 80 years between those who continue to smoke and those who have never smoked have widened and now exceed 30% for both sexes. Moreover, the hazards among women who continue to smoke now approximate those among male smokers,^{10,11,20,21} at least in middle age.

Because the absolute risks of continuing to smoke are large, the absolute benefits of cessation will also be large, particularly as death rates

among those who have never smoked continue to fall.²⁴ Cessation at around 40 years of age results in approximately a 90% reduction in the excess risk of death associated with continued smoking in later middle age and old age. That is not to say, however, that it is safe to smoke until 40 years of age and then stop, for the remaining excess risk of about 20% (hazard ratio, 1.2) is substantial; it means that about one in six of these former smokers who dies before the age of 80 years would not have died if their death rates had been similar to those for persons who had never smoked who were similar in educational levels, adiposity, and alcohol use.

From 1965 to 2010, the prevalence of cigarette smoking among adults in the United States decreased from 42% to 19%, owing in large part to increased rates of cessation.⁸ Although former smokers still far outnumber current smokers at older ages (and especially among men) (Fig. 1), the United States has about 35 million current or future smokers under 35 years of age,²⁵ and the prevalence of smoking changed little from 2004 to 2010.⁸

Our study has some limitations. First, there may be confounding factors other than the few variables recorded in the NHIS. However, most

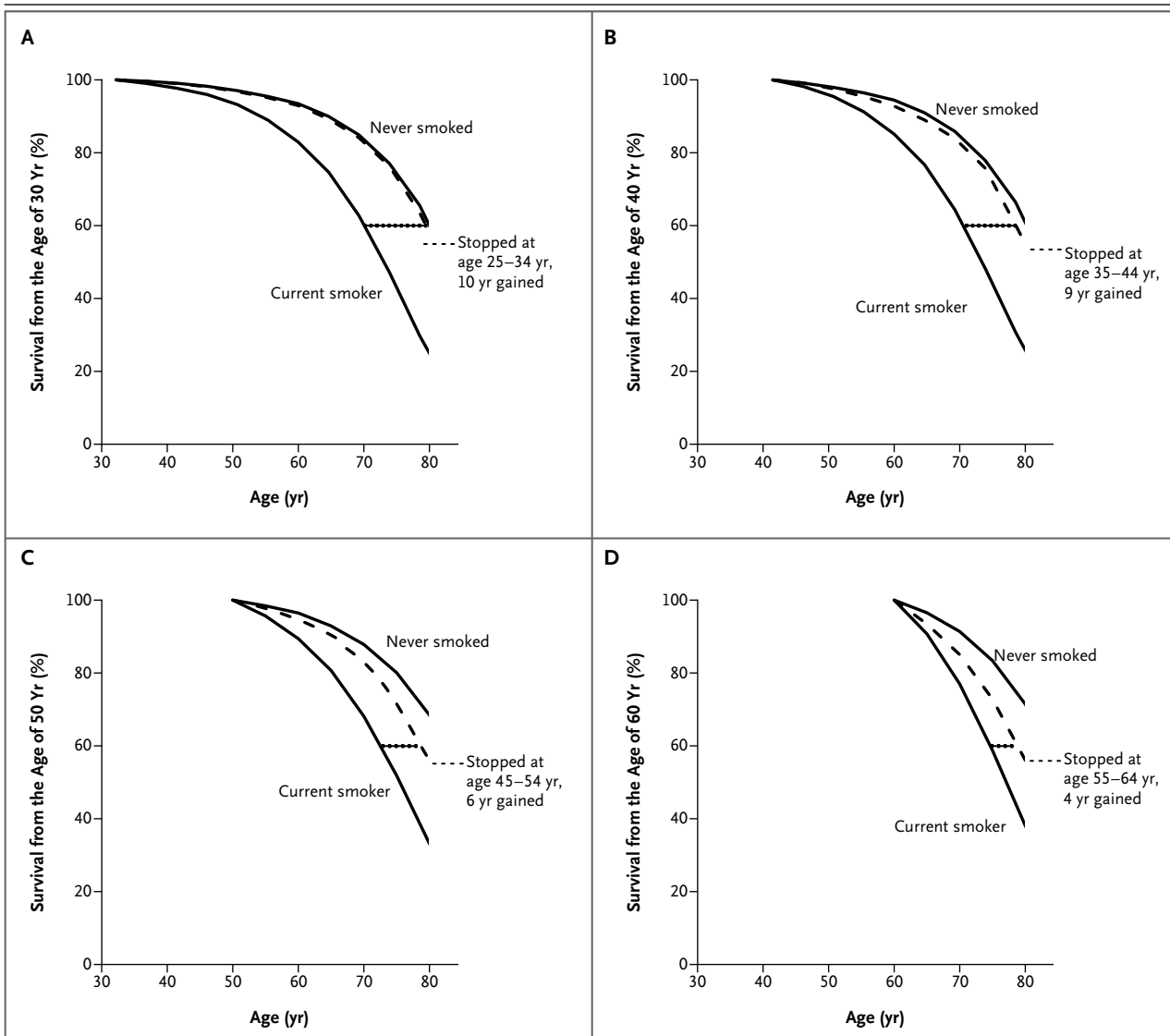


Figure 3. Effect of Smoking Cessation on Survival to 80 Years of Age, According to Age at the Time of Quitting.

Life expectancy was increased from 4 to 10 years among smokers who quit, depending on their age at the time of smoking cessation. Panel A shows the effect of quitting at 25 to 34 years of age (effect shown from the age of 30), Panel B the effect of quitting at 35 to 44 years of age (effect shown from the age of 40), Panel C the effect of quitting at 45 to 54 years of age (effect shown from the age of 50), and Panel D the effect of quitting at 55 to 64 years of age (effect shown from the age of 60). Survival probabilities have been scaled from the NHIS to the U.S. rates of death from all causes at these ages for 2004,^{13,16} with adjustment for differences in age, educational level, alcohol consumption, and adiposity (body-mass index). The horizontal dots represent years of life gained.

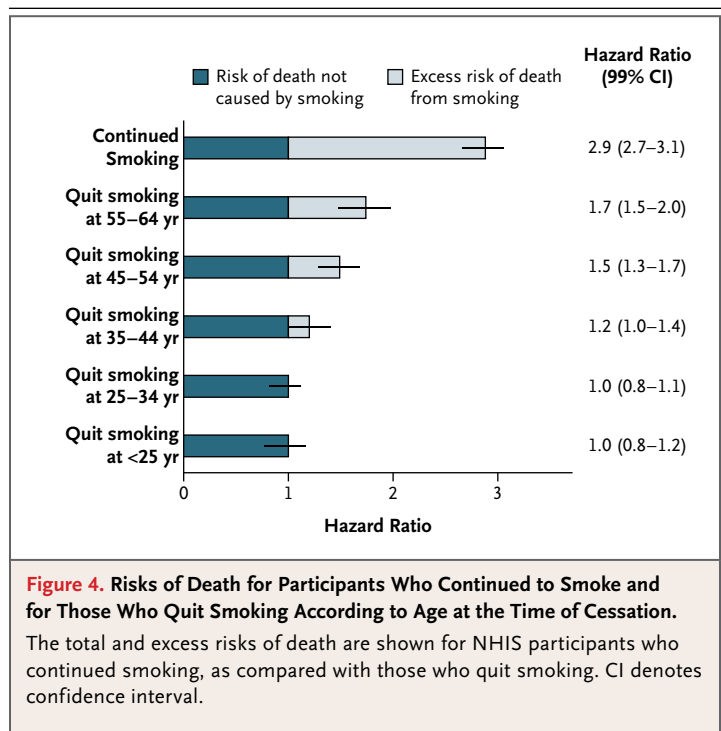
of the excess mortality among smokers was from neoplastic, vascular, or respiratory diseases that other studies have shown can be caused by smoking,²⁻⁶ and adjustment for educational level, alcohol use, and adiposity did little to alter the hazard ratios, suggesting that, to a large degree, the association between smoking and mortality is causal. Second, the NHIS excludes incarcerated

adults (who tend to have an increased prevalence of smoking²⁶). This should not, however, materially affect the observed differences between current smokers and those who never smoked, among the adults surveyed in the NHIS. Third, the number of deaths in the NHIS was lower than that noted in recent studies,^{19,20} but this is counterbalanced by the fact that the sur-

veyed population and death rates in our study are generally representative of the U.S. population and death rates.¹³ Fourth, misclassification of the causes of death, particularly at older ages, might affect the observed hazards for some causes of death (lowering the risks for some causes but raising the risks for others), but this would not affect our analyses of all-cause mortality. Fifth, the NHIS is a cross-sectional survey, and data on smoking status were collected only at baseline. Some of the surveyed smokers would have quit subsequently, thereby somewhat reducing their risk,⁸ but with 7 years of follow-up, any distortion of the hazard ratios should be slight. Similarly, the excess mortality among former smokers might be overestimated, since some deaths may well reflect deaths among smokers who quit because they became ill. In this case, the true gain of life-years from the time of cessation would be somewhat greater than we estimated. Finally, although the smokers who quit smoking might have been more likely than those who had never smoked to try to improve their health, we found little difference between these two groups with respect to alcohol use, adiposity, and other health-related variables.

Our findings, as well as those from other recent prospective studies,¹⁹⁻²¹ highlight the worldwide importance of tobacco control. About 40 million Americans smoke, but most of the world's estimated 1.3 billion smokers live in low- and middle-income countries.^{3,25} Worldwide, about 30 million young adults begin smoking each year (about 50% of young men and about 10% of young women), and current patterns of behavior suggest that most will not stop.²⁵ Our findings are consistent with emerging evidence from China²⁷ and India¹⁶ that cigarette smokers worldwide who continue to smoke can expect to lose about a decade of life.³⁻⁵ In most high-income countries, there are now more former smokers than current smokers; however, cessation remains uncommon in low- and middle-income countries.^{3,25} Thus, on the basis of current rates of smoking initiation and cessation, smoking, which killed about 100 million people in the 20th century, will kill about 1 billion in the 21st century.^{1,3,5,24}

Options to help increase cessation rates and decrease initiation rates worldwide include higher prices for cigarettes through an increased ex-



cise tax, restrictions on smoking in public places, bans on tobacco advertising and promotion, public education about the hazards of smoking and the benefits of cessation, and easy access to cessation efforts.^{3,28} In the United States, the recent increase of 62 cents in the federal excise tax on each pack of cigarettes,²⁹ a recent federal mass-media campaign,³⁰ and the extension of insurance coverage for cessation as part of the Affordable Care Act are likely to raise cessation rates. A focus on cessation of smoking is justified, since quitting smoking before the age of 40 years, and preferably much earlier, will reduce by about 90% the decade of life that is lost from continued smoking. Smoking is associated with a decade of lost life, and cessation reduces that loss by about 90%.

The opinions expressed in this article are those of the authors and do not necessarily represent the opinions of the agencies at which the authors are employed.

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