Percentage of deaths attributable to poor cardiovascular health lifestyle factors: Findings from the Aerobics Center Longitudinal Study

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Abstract

PURPOSE—We assessed the effects of the four newly defined American Heart Association (AHA) lifestyle factors on mortality by examining the associated population attributable fractions (PAFs) of these factors.

METHODS—Slightly modified AHA cardiovascular health factors (smoking, BMI, cardiorespiratory fitness, and diet) were measured among 11,240 (24% women) participants from the Aerobics Center Longitudinal Study between 1987 and 1999. The cohort was followed to December 31, 2003 or death. PAFs were calculated as the proportionate reduction in death attributable to identified risk factors.

RESULTS—During an average 12 years of follow-up, 268 deaths occurred. Low fitness had the highest PAFs at the 5th, 10th, and 15th year of follow-up, respectively: 6.6%, 6.4%, and 5.5%. Current smokers had the second highest PAFs at the 5th, 10th, and 15th year of follow-up, respectively: 5.4%, 5.2%, and 5.0%. Additional adjusting for other confounders in the model did not change the above associations. The PAFs for overweight or obesity and unhealthy diet were not significant in the current analyses.

CONCLUSIONS—Assuming a causal relationship between smoking, low fitness and mortality, avoidance of both would have prevented 13% of the deaths in the current population. Preventive interventions to increase physical activity and stop smoking would most likely promote longevity.

INTRODUCTION

Cardiovascular disease (CVD) continues to be the leading cause of death in the United States with an average of 1 death every 39 seconds and an estimated direct and indirect cost of $286.6 billion [1]. Recently, the American Heart Association (AHA) 2020 Impact Goal
defines a new construct of cardiovascular health behaviors for adults and children based on 4 lifestyle factors (smoking, body mass index (BMI), physical activity, and diet) and sets national goals for promoting cardiovascular health and reducing CVD burden [2]. Previous studies consistently show an inverse association between multiple low-risk lifestyle factors and all-cause mortality [3–7], however, widely varying definitions of the lifestyle factors are applied in these studies. To date, only two studies have used AHA cardiovascular health behavior concept to define the 4 lifestyle factors [8, 9]. Bambs and colleagues addressed the prevalence of the new AHA metrics that define cardiovascular lifestyle factors and reported 81% of all participants in the community-based Heart SCORE study presented ≤ 3 ideal lifestyle factors (nearly 60% presented zero or one ideal factor) [8]. In an ideal world with unlimited resources, targeting all the 4 modifiable factors could result in substantial improvement in overall cardiovascular health, however, with the continuing financial crisis around the globe, it may be more cost-effective to choose the one or two most important lifestyle factors for intervention.

The population attributable fraction (PAF) is an integrated measure that assesses the proportion of an outcome in a population that is attributable to exposure to 1 or more risk factors [10]. The aim of our study is to estimate the death burden that is attributable to non-ideal or poor cardiovascular lifestyle factors using a slightly modified construct defined by the AHA, and to identify one or two factors which contribute to most of the deaths while accounting for age, gender, family history and health status.

METHODS

Study Population

The Aerobics Center Longitudinal Study (ACLS) began in 1970 as an observational epidemiological study to investigate health outcomes associated with physical activity and cardiorespiratory fitness [11, 12]. Participants came to the Cooper Clinic, Dallas, Texas for a preventive medical examination and for consultation regarding their unhealthy lifestyle behaviors. They were unpaid volunteers, sent by their employers, healthcare providers, or self-referred and came from all 50 states. Participants were told the purpose of the study and provided their written informed consent to participate. The study protocol was approved annually by the Cooper Institute's institutional review board. All participants included in this study were 20 to 82 years old at entry (23.5% women), had an extensive baseline health examination, normal electrocardiograms (ECGs), a body mass index (BMI) ≥8.5 kg/m², and complete data on the 4 lifestyle factors. Those who reported having a history of myocardial infarction, stroke, or cancer, and those who had less than one year of follow-up were excluded. Based on these inclusion and exclusion criteria, the final analysis included 11240 individuals whose baseline examination took place between 1987 and 1999. Figure 1 shows the flow diagram of the study participants.

Data Collection

A medical history questionnaire at baseline provided age, sex, history of chronic diseases (myocardial infarction, stroke, cancer, hypertension, or diabetes), family history of CVD, cigarette smoking, and physical activity. Diet was assessed by a 3-day dietary record. Height and weight were measured. Blood samples were obtained after a fast of at least 12 hours and analyzed for lipids and glucose using automated bioassays in accordance with the Centers for Disease Control and Prevention Lipid Standardization Program. Diabetes was defined as glucose ≥7.0 mmol/L, a history of physician diagnosis, or use insulin. Hypercholesterolemia was defined as serum cholesterol ≥6.2 mmol/L or a history of physician diagnosis. Hypertension was defined as systolic blood pressure ≥140 mm Hg or diastolic blood pressure ≥90 mm Hg, or a history of physician diagnosis.
**Modified AHA Cardiovascular Lifestyle Factors**

In accordance with AHA definition of cardiovascular health [2], we classified each ideal lifestyle factors in the ACLS at baseline. Some factors from the definition of ideal cardiovascular health were slightly modified, according to the information available in the ACLS.

**Smoking habits** were determined from a standardized medical history questionnaire. Participants were classified as never, former, or current smokers, and ideal smoking behavior was defined as nonsmoker (never or former smoker).

**Body mass index (BMI)** was calculated as weight (kilograms)/the square of height (meters) which were measured using a stadiometer and balance beam scale, respectively. Ideal BMI was defined <25 kg/m².

**Physical activity and cardiorespiratory fitness**

Physical activity was measured through a self-report questionnaire over the previous 3 months. We do not have uniform collection of information on activity type, frequency, duration and intensity over the entire duration of the ACLS. Therefore, for the purpose of this study, we used cardiorespiratory fitness (hereafter referred to as ‘fitness’) as an objective marker of physical activity [13]. Fitness was assessed by a symptom-limited maximal exercise treadmill test using a modified Balke protocol [14]. Total treadmill endurance time (minutes) was used as an index of aerobic power, with time on treadmill in this protocol correlated highly (r ≥0.92) with maximal oxygen uptake (VO₂max) in both men [15] and women [16]. Participants were classified as ‘low fit’ based on the lowest 20% of the age- and sex-specific distribution of treadmill exercise duration in the overall ACLS population. These cut-points are from previous reports on the relation between fitness and all-cause mortality in the ACLS [12]. Ideal fitness was defined as 80% of the age- and sex-specific distribution of treadmill duration in the overall ACLS population (moderate and high levels of fitness) and was considered as an indictor of ideal physical activity status because previous ACLS data have shown that a brisk walk of approximately 30 minutes on most days of the week was associated with moderate to high levels of fitness [17].

**Diet assessment** consisted of a 3-day diet record that required participants to keep detailed records of everything they ate over 2 pre-assigned weekdays and 1 weekend day. Participants were provided written instructions on how to accurately describe foods and estimate portion sizes. Participants kept an on-going, real-time written record of foods consumed during and between meals, including assessing portion sizes in common household measures. Registered dieticians at the Cooper Clinic coded and analyzed the diet records using the Cooper Clinic Nutrition and Exercise Evaluation system [18]. This provided detailed dietary information on the overall diet such as the number of foods consumed from specific food groups and the volume of micronutrients (vitamins and minerals).

The ideal cardiovascular health’s definition of the dietary goals included fruits and vegetables (≥4.5 cups per day, approximated as ≥4.5 servings/day in the ACLS study); fish (≥2 servings/week, approximated as ≥2 servings/week of cooked lean meat equivalents from fish, selffish and other seafood); fiber-rich whole grains (≥1.1 g of fiber per 10 g of carbohydrate: three 1-oz-equivalent servings/day, approximated as ≥3 servings/day of whole grains); sodium (< 1500 mg per day) and sugar-sweetened beverages (≤450 kcal (36 oz) per week). All these dietary components were included in this study, except sugar-sweetened beverages, due to a very low number of participants with this...
information. Ideal diet behavior was defined as meeting at least 3 of the above 4 diet components.

**Mortality Surveillance**

We followed participants for mortality from the baseline examination through the date of death for decedents or December 31, 2003 for survivors using the National Death Index. We excluded participants with less than one year of follow-up to minimize potential bias due to serious underlying illness on mortality.

**Statistical Analyses**

The length of follow-up for each person was determined as the time from baseline to either death or censoring, whichever came first. Descriptive analyses summarized baseline characteristics of participants by survival status. The mean levels of continuous variables were compared using student t-test, while chi-square tests compared the distribution of categorical variables values. Univariate and multivariate Cox regressions were used to estimate the strength of the association (hazard ratios (HRs) and 95% confidence intervals (CIs)) between the selected potential nonmodifiable (age, gender, and family history of CVD) and modifiable lifestyle factors and all-cause mortality. With a score of 0 being allocated for each ideal lifestyle factor, non-ideal or poor lifestyle factors were coded as 1. The PAFs for smoking (current smoking), fitness (low fitness), BMI (≥ 25), and diet (meeting 0 or 1 of the 4 diet components) on all-cause mortality were calculated. The PAF for the non-ideal or poor lifestyle factor is defined as the proportionate reduction in death that would be achieved if the entire population had been unexposed to that risk factor or a combination of two lifestyle factors, compared with its current (actual) exposure pattern. Without considering confounders, the PAF can be defined by PAF(t)=1-D_0(t)/D(t), where D(t) is the probability of death during the time interval (0, t) in the overall population, and D_0(t) is the death probability in the subgroup unexposed to the studied risk factor. When considering confounders we used the definition in Chen et al., [19] which has been shown to have a causal interpretation. For the special case of a categorical confounder and a fixed time point, the adjusted PAF in Chen et al. reduces to that defined in Whittemore [20] which replaced D_0(t) in the above formula with the weighted average of confounder-stratum-specific unexposed death probability. The unadjusted/adjusted PAFs and 95% confidence intervals were estimated based on the univariate/multivariate Cox regression models and the methods developed by Chen et al.[19]. Data analyses were performed using SAS (version 9.2; SAS Institute, Cary, NC) software, C and R 2.12.2. All P values are 2-sided with an alpha level of 0.05 established for significance.

**RESULTS**

There were 268 deaths during 130,584 person-years of follow-up. Compared with survivors, decedents were older, had lower fitness and higher BMI, had higher prevalence of major CVD risk factors, and had lower prevalence of ideal modifiable health factors other than body weight (Table 1).

We found statistically significant differences in risk of death between categories of age, gender, family history of CVD, smoking, and low fitness (Table 2). Low fitness and smoking, in addition to age, showed the strongest associations with mortality. No statistically significant interactions between the variables in the model were found, therefore, we used a model with main effects only to study their effects on the risk of death. The fully adjusted model showed similar associations between age, gender, low fitness, and smoking and death as the unadjusted model.
Table 3 shows the unadjusted and adjusted PAFs and their CIs for the individual modifiable lifestyle factor and 6 possible combinations of two factors in each of the 5 year follow-up intervals. Of the 4 lifestyle factors, fitness had the strongest association with risk of death, reducing it by 7% if low fit individuals had become at least moderately fit (95% CI: 2.2, 12). A reduction in smoking would have led to a 6% reduction in mortality risk. However, reduction of BMI to the ideal level had the smallest effect, the PAF being −7% (95% CI: −14–8). The unadjusted PAF estimates of fitness, smoking, and diet decreased at year 10 and year 15 comparing with the estimates at year 5, respectively. However, this pattern of associations disappeared after multivariate adjustment (Figure 2).

Finally, the cumulative PAF estimate obtained from the full model including any of the 2 lifestyle factors (Table 3 and Figure 3). Of the 6 combination groups, smoking plus fitness had the greatest association with PAF, reducing it by 13% if all current smokers who also had low fitness had never started smoking and also became at least moderate fit (95% CI: 6, 19). Improving diet and reducing BMI had the least effect on PAF in this population of men and women.

**DISCUSSION**

**Summary of main findings**

Attributable fractions are commonly used to measure the effect of risk factors on disease outcomes in a population. In this study, we extended these static measures to functions of time because we are interested in knowing if the PAFs will change with the event time. We found that fitness and smoking were the two most important factors for reducing risk of death in our population. The fraction of deaths attributable to low fitness equal to 7% and to smoking equal to 6% after accounting other potential confounders. Considering the growing burden of CVD, this study provides some useful information for decision-makers by providing more information about identifying priority targets of modifiable health factors. Furthermore, by comparing the relative importance of these factors, we can focus policy debate concerning the opportunity of financing one intervention over another.

**Comparison with other studies**

Previous studies have examined the 4 individual modifiable lifestyle factor, their association with mortality, and the PAF in different populations. Smoking has long been identified as a behavior that has an inverse association with longevity. Among developed Western countries, earlier reports show that 20% of all deaths are attributed to tobacco [21] and this number has varied in recent reports due to different populations studied [22, 23]. In Asian countries there have been several reports on PAF due to smoking ranging from 25% in Japan [24] to 13% in China [25]. It is becoming more evident that physical activity and fitness have an important role on health outcomes. The Nurse's Health Study [26], a large cohort study of registered US nurses, assessed the PAFs of a wide range of risk factors, found that a total of 17% of deaths during follow-up could be attributed to lack of physical activity (<30 min of physical activity per day). Another study from the UK reported a higher PAF of 25% associated with physical inactivity among 10,059 middle-aged women under general practitioners' observation [23]. Poor diet is often commonly reported to be associated with a wide range of chronic conditions, and therefore contribute to substantial burden of disease [22]. Due to the complex nature of the diet exposure assessment, definitions of poor diet or unhealthy diet often vary significantly. The Nurse's Health Study developed a healthy eating score [26], those in the upper two fifths was defined as a low risk category. They found that a total 13% of deaths were attributed to a low diet quality. A meta-analysis of US studies shows a 17% PAF for combined poor diet and physical inactivity, but the investigators did not provide separate estimates for diet and inactivity [27]. Obesity has
consistently been associated with higher risk of mortality [28]. Being overweight or obese had a higher PAF (14.2%, 95% CI = 11.6 to 16.9) in the US cohort [26], but there was no significantly higher mortality risk associated with overweight or obesity in the UK study [23].

Although there is an increasing literature on combinations of lifestyle factors with mortality [4, 23, 26, 29], few studies have estimated PAF for the individual factors [23, 26], even fewer studies have reported the PAF for a combination of factors [23], therefore, information is sparse on exactly which combination of lifestyle factors might be best to target to reduce overall disease burden. Among studies estimating PAFs, most of them failed to identify which lifestyle factor(s) is the most important one(s) to contribute to the overall mortality [4, 26, 29]. An important public health question is how to identify which lifestyle factor interventions might yield the greatest benefits in the population. The only current study that tried to answer this question is the UK study in middle-aged women [23]. Iversen and colleagues reported among the four individual lifestyle factors (smoking, physical inactivity, BMI and alcohol) smoking remains the leading cause of mortality and physical inactivity follows. When the investigators evaluated combinations of two factors, they found that smoking and physical inactivity are responsible for almost half of the deaths. They also report that a combination of three factors of smoking, physical inactivity and alcohol consumption are responsible for about 60% of the deaths. Finally, avoidance of all four unhealthy lifestyle factors would also have prevented 60% of the deaths. These analyses indicate that avoidance of all four unhealthy lifestyle risk factors would only have prevented additional 10% of the deaths when compared with the combination of smoking and inactivity. However, this study did not assess the contribution of diet on mortality.

The changes observed in “diet” and “BMI” before and after the adjustment deserve further comment. We have compared overweight/obese participants with those who had normal BMI and found that overweight/obese participants had significantly higher prevalence of hypertension, diabetes, and hypercholesterolemia, but no differences in mean age. The PAF estimate changed direction which means modifying BMI from non-ideal to ideal level did not contribute significantly to mortality after adjusting these other risk factors. We postulate this might be due to the strong effects of these negative confounders and simply improving BMI without changing other risk factors is not sufficient to change the proportion of death in the population level. On the other hand, we found ideal diet participants were not significantly different from those with non-ideal diet regarding the above mentioned confounders except age. Those with non-ideal diet were significantly younger than those with ideal diet. Based on Table 2, we can see that age had a significant and large effect on all-cause mortality. Therefore improving diet after adjusting other confounders especially age might influence the PAF estimate significantly.

ACLS participants were mainly white and middle to upper socioeconomic status. Although participants are similar in many respects to other US cohorts that have provided important information on disease prevention [11], the prevalences of smoking and low fitness were low (11% and 8%, respectively). However, even with such low prevalences, they were still responsible for the largest number of deaths in this population. From this point of view, our results are in fact consistent with previous cohort study and suggest that smoking cession programs and physical activity promotion have the greatest potential for reducing the total number of deaths.

Limitations

As mention previously, the ACLS population was mainly white, and of middle to upper socioeconomic status. The prevalence of the four behavior factors was lower than the US general population. Thus, these factors might have a larger population effect if they were
studied in a representative sample. We used fitness instead of physical activity in the current study. Although fitness and physical activity are not interchangeable because physical activity is a behavior, whereas fitness is a functional attribute that can be influenced by other factors, physical activity is the primary determinant of fitness, and fitness is less prone to misclassification and may better reflect the adverse health consequences of a sedentary lifestyle than does self-reported physical activity exposure [13]. We were unable to evaluate the effect of changes in the factors over time on all-cause mortality because we only had baseline assessments. The PAF estimates depend on the prevalence of the studied factors and the magnitude of the association between them and the outcomes. During this long follow-up period, both the prevalence of the lifestyle factors and the observed associations between these factors and all-cause mortality are likely to change, therefore influence the PAF estimates. Future studies are needed with repeat assessment of the behaviors. In addition, the measurement of fitness and diet might not be feasible for clinical practice. Finally, the estimates presented here assume that changes in the behaviors would affect the number of deaths. Our ability to change lifestyle factors in large populations is largely untested. However, we do know that it is possible to produce lifestyle changes in clinical trials [30].

Conclusions

In conclusion, low fitness was responsible for the highest proportion of deaths in this sample of men and women, and smoking had the second highest PAF among the four cardiovascular lifestyle factors. Preventive targeted interventions to decrease the prevalence of low fitness and smoking would most likely promote reduced mortality rates. Therefore the major public health gain in the coming years will come from getting sedentary individuals to start moving as well as getting smokers to stop smoking.

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FIGURE 1.
Participants flow diagram.
FIGURE 2.
Estimates of the population attributable fraction (PAF) by assuming one of the four behavior factors is controlled at the ideal level. a). unadjusted model; b). multivariate-adjusted model (age, gender, hypertension, diabetes, hypercholesterolemia, and family history of CVD), the Aerobics Center Longitudinal Study, 1987–2003.
FIGURE 3.
Estimates of the population attributable fraction (PAF) by assuming two of the four behavior factors are controlled at the ideal level. a), unadjusted model; b), multivariate-adjusted model (age, gender, hypertension, diabetes, hypercholesterolemia, and family history of CVD), the Aerobics Center Longitudinal Study, 1987–2003.
|                                | All   | Survivor | Decedent | P-value |
|--------------------------------|-------|----------|----------|---------|
| N                              | 11,240| 10,972   | 268      |         |
| Age, years                     | 45.7 (9.7) | 45.5 (9.6) | 53.5 (11.1) | <0.0001 |
| Female, %                      | 23.5 | 23.7     | 14.6     | 0.0005  |
| Body mass index, kg/m²         | 25.8 (4.1) | 25.8 (4.0) | 26.4 (4.7) | 0.02    |
| Treadmill time, minutes        | 18.1 (5.2) | 18.1 (5.2) | 16.5 (5.7) | <0.0001 |
| Total cholesterol, mmol/L      | 5.4 (1.0)  | 5.4 (1.0)  | 5.6 (1.0)  | 0.0003  |
| Fasting blood glucose, mmol/L  | 5.5 (0.9)  | 5.5 (0.9)  | 5.7 (1.3)  | 0.01    |
| Blood pressure, mmHg           |       |          |          |         |
| Systolic                       | 119 (14)  | 119 (14)  | 125 (16)  | <0.0001 |
| Diastolic                      | 80 (10)   | 80 (10)   | 83 (10)   | <0.0001 |
| Hypertension<sup>a</sup>, %    | 26.4     | 26.1     | 41.0     | <0.0001 |
| Diabetes mellitus<sup>b</sup>, %| 2.6     | 2.5     | 5.6     | 0.002   |
| Hypercholesterolemia<sup>c</sup>, % | 29.2 | 29.0   | 36.9     | 0.005   |
| Family history of CVD, %       | 11.3    | 11.2    | 14.6    | 0.08    |
| Ideal modifiable lifestyle factors, % |       |          |          |         |
| Nonsmoker                      | 88.6    | 88.6    | 84.7    | 0.045   |
| 18.5 ≤BMI <25                  | 47.0    | 47.1    | 45.9    | 0.71    |
| Moderate and high fit          | 91.3    | 91.4    | 86.9    | 0.01    |
| Healthy diet (3–4 components)  | 37.6    | 37.5    | 43.3    | 0.05    |

Mean (Standard deviation) for continuous variables; Percentage for categorical variables. CVD=cardiovascular disease.

<sup>a</sup>Hypertension is defined as systolic blood pressure ≥140 mmHg, diastolic blood pressure ≥90 mmHg, or a history of physician diagnosis.

<sup>b</sup>Diabetes mellitus is defined as a fasting plasma glucose concentration ≥7.0 mmol/L, a history of physician diagnosis, or insulin use.

<sup>c</sup>Hypercholesterolemia is defined as total cholesterol ≥6.20 mmol/L, or a history of physician diagnosis.
### TABLE 2
Estimated hazard ratios of death in categories of potential risk factors, the Aerobics Center Longitudinal Study, 1987–2003.

| Nonmodifiable variables | No. of Deaths | Total No. of Participants | Unadjusted HR (95% CI) | Multivariate-adjusted HR (95% CI) |
|-------------------------|--------------|---------------------------|------------------------|----------------------------------|
| Age group, years        |              |                           |                        |                                  |
| 20–39                   | 30           | 2999                      | 1                      | 1                                |
| 40–49                   | 71           | 4453                      | 1.62 (1.05–2.48)       | 1.58 (1.03–2.42)                 |
| 50–59                   | 83           | 2874                      | 2.91 (1.92–4.42)       | 2.88 (1.88–4.41)                 |
| ≥60                     | 84           | 914                       | 8.48 (5.59–12.86)      | 8.53 (5.55–13.13)                |
| Gender                  |              |                           |                        |                                  |
| Male                    | 229          | 8598                      | 1                      | 1                                |
| Female                  | 39           | 2642                      | 0.59 (0.42–0.82)       | 0.66 (0.47–0.94)                 |
| Family history of CVD   |              |                           |                        |                                  |
| No                      | 229          | 9974                      | 1                      | 1                                |
| Yes                     | 39           | 1266                      | 1.37 (0.98–1.93)       | 1.46 (1.04–2.05)                 |
| Non-ideal modifiable lifestyle factors | |                           |                        |                                  |
| Current smoker          | 41           | 1287                      | 1.50 (1.07–2.09)       | 1.62 (1.16–2.28)                 |
| BMI ≥25                 | 145          | 5954                      | 1.81 (0.93–1.50)       | 0.88 (0.68–1.14)                 |
| Unhealthy diet (<3 components) | 152          | 7011                      | 0.87 (0.68–1.11)       | 1.01 (0.79–1.29)                 |
| Low fitness             | 35           | 981                       | 1.81 (1.27–2.59)       | 2.04 (1.40–2.99)                 |

CVD = cardiovascular disease.

a Adjusted for all the variables in the table plus hypertension, diabetes, and hypercholesterolemia.
### TABLE 3

Unadjusted and multivariable-adjusted PAF for each 5-year time interval using the Aerobics Center Longitudinal Study, 1987–2003.

| Individual factor | Unadjusted model | Multivariate-adjusted model |
|-------------------|------------------|-----------------------------|
|                   | PAF5 95% CI      | PAF10 95% CI                | PAF15 95% CI                |
| Smoking           | 5% 0.1%, 10%     | 5% 0.1%, 10%                | 5% 0.2%, 10%                |
| BMI               | 9% −5%, 20%      | 9% −5%, 20%                 | 8% −4%, 19%                 |
| Diet              | −9% −26%, 6%     | −9% −25%, 5%                | −8% −24%, 5%                |
| CRF               | 7% 2%, 11%       | 6% 2%, 11%                  | 6% 1%, 10%                  |
| Multiple factors  |                  |                             |                             |
| Smoking+BMI       | 13% −1%, 24%     | 12% −1%, 24%                | 12% −1%, 23%                |
| Smoking+Diet      | −4% −22%, 11%    | −4% −21%, 10%               | −4% −20%, 10%               |
| Smoking+CRF      | 11% 4%, 17%      | 10% 4%, 16%                 | 9% 3%, 15%                  |
| BMI+Diet          | 4% −21%, 18%     | 0.4% −21%, 18%              | 0.2% −20%, 17%              |
| BMI+CRF          | 11% −2%, 22%     | 11% −2%, 22%                | 10% −3%, 20%                |
| Diet+CRF         | −4% −21%, 11%    | −4% −20%, 11%               | −4% −20%, 10%               |

| Individual factor |                  |                             |                             |
|                   | PAF5 95% CI      | PAF10 95% CI                | PAF15 95% CI                |
| Smoking           | 6% 1%, 11%       | 6% 1%, 11%                  | 6% 1%, 11%                  |
| BMI               | −7% −14%, 8%     | −7% −24%, 8%                | −7% −23%, 8%                |
| Diet              | 3% −13%, 16%     | 3% −13%, 16%                | 3% −12%, 15%                |
| CRF               | 7% 2%, 12%       | 7% 2%, 12%                  | 7% 2%, 12%                  |
| Multiple factors  |                  |                             |                             |
| Smoking+BMI       | −0.3% −17%, 14%  | −0.3% −17%, 14%             | −0.3% −16%, 14%             |
| Smoking+Diet      | 9% −7%, 22%      | 9% −7%, 22%                 | 9% −7%, 21%                 |
| Smoking+CRF      | 13% 6%, 19%      | 13% 6%, 19%                 | 13% 6%, 18%                 |
| BMI+Diet          | −4% −28%, 15%    | −4% −28%, 16%               | −4% −27%, 14%               |
| BMI+CRF          | 1% −15%, 15%     | 0.9% −15%, 14%              | 0.8% −14%, 14%              |
| Diet+CRF         | 10% −5%, 23%     | 10% −5%, 23%                | 9% −5%, 22%                 |

PAF=population attributable fraction; CI=confidence interval; BMI=body mass index; CRF=cardiorespiratory fitness; CVD=cardiovascular disease.

\(^{a}\)PAF value at 5th, 10th, and 15th year was denoted by PAF5, PAF10, and PAF15, respectively.

\(^{b}\)The definition of Smoking, BMI, Diet, CRF is the same as in Table 2.

\(^{c}\)Adjusted for age, gender, hypertension, diabetes, hypercholesterolemia, and family history of CVD.