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(4) Supplemental Reference

1 Supplemental Methods

2 General overview of the Cardiovascular Disease Policy Model-China

3 The Cardiovascular Disease (CVD) Policy Model-China is a computer-simulation, state-4 transition (Markov cohort) mathematical model of coronary heart disease (CHD) and stroke incidence, prevalence, mortality, non-cardiovascular deaths, and costs of health care in Chinese 5 6 population aged 35-84 years old. This model has been used for CVD epidemiologic projections and effectiveness analysis of specific policy interventions.¹⁻⁴ Because air pollution was much 7 8 severer in urban areas and no reliable PM_{2.5} data was available for rural areas, we created an 9 urban China version of the model with updated levels of traditional cardiovascular risk factors for projections. The model start year is 2010 and the model cycle length is one year. Simulations 10 11 are at the sub-national population level. The standard model simulates a dynamic national population, adding waves of 35-year adults with each successive cycle. 12

The CVD Policy Model consists of three sub-models: the Demographic-Epidemiological model, 13 14 the Bridge model and the Disease History model. The Demographic-Epidemiological model 15 predicts CHD and stroke incidence and non-CVD mortality among subjects without CVD, stratified by age, sex, systolic blood pressure (SBP, <140, 140-159.9, ≥160 mmHg), body mass 16 index (BMI, <25, 25-29.9, \geq 30 kg/m²), low density lipoprotein (LDL) cholesterol (<100, 100-17 18 129.9, \geq 130 mg/dL) and high density lipoprotein (HDL) cholesterol levels (<40, 40-59.9, \geq 60 19 mg/dL), and status of smoking (active smoker, non-smoker with exposure to environmental tobacco smoke, non-smoker without environmental exposure), diabetes (yes or no) and PM_{2.5} 20 21 exposures (yes or no) in urban Chinese population in ten-year age categories among those aged 22 35-84 years. Means and proportions of CVD risk factors were estimated from the China Cardiovascular Health Study and the China Multicenter Collaborative Study of Cardiovascular 23

| 1 | Epidemiology (ChinaMUCA) for urban adults in 10-year age categories aged 35 to 84 years. ^{5, 6} |
|---|--|
| 2 | An annual population-weighted average $PM_{2.5}$ level during the period of 2014-2015 was |
| 3 | extracted in 190 cities with over 950 monitoring sites and was assumed as national $PM_{2.5}$ level in |
| 4 | 2017. ⁷ All individuals were assigned the mean $PM_{2.5}$ exposure for urban China. Multivariable |
| 5 | adjusted hazard ratios of SBP, LDL, HDL, BMI, smoking and diabetes for CHD, stroke, and |
| 6 | non-CVD (non-CHD, non-stroke) death by age and sex were estimated from the China Multi- |
| 7 | provincial Cohort Study (CMCS) ⁸ using a competing risk Cox proportional hazard model for |
| 8 | each outcome. |

9 For individuals in whom CVD develops, the Bridge Sub-model characterizes the initial CHD or stroke event (cardiac arrest, myocardial infarction, or angina) and its sequelae for 30 days. Then, 10 11 the Disease History Sub-model predicts subsequent CVD events, coronary revascularization procedures, CVD mortality, and non-CVD mortality among patients with CVD, stratified by age, 12 13 sex, and history of events. The general chronic CVD categories include CHD only, stroke only, and combined prior CHD and prior stroke. Each state and event has an annual probability of a 14 recurrent event and/or transition to a different CVD state. The model assumes survivors persist in 15 a chronic disease state without remission. 16

Stroke incidence^{9, 10}, mortality¹¹ and case-fatality⁹ were obtained from other studies. The main outcomes predicted were CHD events (nonfatal and nonfatal first-ever and repeat episodes of stable and unstable angina, myocardial infarction, or cardiac arrest) and stroke events (nonfatal and fatal ischemic and hemorrhagic strokes). The CVD Policy Model-China defined CHD as myocardial infarction (ICD-9 410, 412 or ICD-10 I21, I22), angina and other CHD (ICD-9 411, 413 and 414, or ICD-10 I20, I23-I25), and a fixed proportion of "ill-defined" CVD coded events and deaths (ICD-9 codes 427.1, 427.4, 427.5, 428, 429.0, 429.1, 429.2, 429.9, 440.9 or ICD-10
 I47.2, I49.0, I46, I50, I51.4, I51,5, I51.9, and I70.9).¹²

Stroke was defined by ICD-9 codes 430-438 (excluding transient ischemic attack) or ICD-10
I60-I69. Finally, starting with CHD and stroke case fatality obtained from the Beijing SinoMONICA study.⁹ The CVD Policy Model-China mortality projections were calibrated to fit with
age-specific and overall CHD and stroke mortality numbers for the years 2010-2011 estimated
by the China Center for Disease Control (CDC).¹³

8 Urban China population estimates

9 Estimates for the urban China population aged 35-84 years old by age and sex were based on the
10 6th China census conducted in 2010.¹⁴ The impact of aging and growth on population were
11 estimated based on by *World Population Prospects* by United Nation Population Division.
12 Population projections by age and sex started in 2010 were based on historical estimates of
13 population by age and sex using probabilistic projections up to 2100 of total fertility and life
14 expectancy at birth by sex.¹⁵

15 Urban-rural ratio was estimated by World Urbanization Prospects by United Nation Population Division¹⁶ using an established and robust extrapolation method. Last two empirical data points 16 from two censuses were used to calculate the urban-rural ratio. The average annual rate of 17 change in the urban-rural ratio between the last two data points was calculated and then 18 19 extrapolated, assuming that the proportion urban follows a logistic path. Then empirical urbanrural growth differences from 148 countries with 2 million or more inhabitants were combined in 20 21 a regression equation. The fitted regression line was used to calculate a hypothetical urban-rural growth difference for each level of an initial observed percentage urban. Starting from the most 22

recent urban-rural growth difference of a particular country, the hypothetical urban-rural growth
difference of all countries over a period of 25 years was converged. In China, urban was defined
as cities and towns, excluding villages according to China census protocol.¹⁴ The urban-rural
ratio of China was 49.2% in 2010 and projected to increase from 55.6% in 2015 to 68.7% in
2030. Then urban population for year 2017-2030 was estimated by multiplying the projected
total China population by urban-rural ratio (Supplemental Table S1).

7 Effects of traditional non-communicable disease (NCD) risk factors

For the standard CVD Policy Model-China, annual probability of first CVD events and non-8 9 CVD deaths conditioned on demographic and risk factors were estimated by analyzing the 10 CMCS. The CMCS was a cohort study of 30,121 male and female participants aged 35-64 years and with no CVD at baseline in 1992-1993. Details could be found elsewhere.⁸ These 11 participants were recruited from 16 centers in 11 Chinese provinces using a multistage sampling 12 method. Majority of participants (80.3%) were in urban areas and the remainder were in rural 13 14 areas. Overall baseline participation rate was 82%. Baseline measurement of risk factors followed a standard protocol (WHO-MONICA protocol) and blood samples were processed at a 15 16 central laboratory. Case-finding of new CHD and stroke events and non-cardiovascular deaths was first done by face-to-face interview. Events were ascertained by 1) detailed interview of 17 participants or family members, 2) review of hospital records. These events were later 18 adjudicated by investigators at the Beijing Institute for Heart, Lung, and Blood Vessel Diseases. 19 After 1996, six centers ceased follow up because of completion of that national research project, 20 21 but the remaining 10 centers (16,552 participants) were followed up through the end of 2002. 22 Follow up rate was 86% for the centers followed all of 1992-2002, and 65% of the original cohort of 16 centers. Multivariable Cox proportional hazard ratios for SBP, diabetes, LDL, HDL, 23

6

1 BMI, and active smoking were estimated from baseline measurements and ischemic and

2 hemorrhagic events occurring over 159,400 person-years of observation in CMCS participants

aged 35-74 years (**Supplemental Table S2**).⁸ Significant (P < 0.05) age*risk factor coefficient

4 interactions (higher risk at higher ages) were found for smoking in CMCS multivariable CHD

5 models, SBP, and smoking in total stroke models, and smoking and diabetes in non-

6 cardiovascular mortality models, so these were incorporated in age-specific risk coefficients.

7 Traditional NCD risk trend estimations (2017-2030)

8 Future traditional NCD risk factors trends for population aged 35-84 years were projected 9 forward from 2017 to 2030 based on recent temporal trends from 1990 to 2009. Temporal trend 10 estimations were based on repeated China Health and Nutrition Surveys (CHNS) from 1991 to 11 2009. The CHNS is repeated household survey which initiated in 1989 using a multistage, random cluster process to draw a sample of over 30,000 individuals in 15 provinces and 12 13 municipal cities across China. Follow-ups were conducted continuously every two to four years 14 to obtain repeated measures on health and nutrition, including traditional NCD risk factors. Data are available at http://www.cpc.unc.edu/projects/china. 15

After the participants have seated for at least 5 minutes, blood pressure (BP) was measured on
the right arm by trained research staff. BP was measured three times at each survey visit using a
standard mercury sphygmomanometer. Then SBP was calculated as the mean of the second two
measurements. Weight and height was measured at each survey year for BMI calculation.
Weight was measured to the nearest 0.01 kg with a balance-beam scale, and height to the nearest
0.10 cm using a stadiometer. BMI was calculated as weight in kilograms divided by the square of
height in meters. Active smoking was defined as self-report of current smoking cigarettes.

| 1 | Temporal SBP, BMI and active smoking trends were estimated using age-adjusted mixed linear |
|----|--|
| 2 | random effects model with 10-year age groups. Due to limited participants aged over 75 years in |
| 3 | CHNS, we combined the last two age groups together for trend estimates. Age-time interactions |
| 4 | observed in trends for SBP, BMI, or active smoking were incorporated into age-specific risk |
| 5 | factor trend projections. Both SBP and BMI were projected to increase over time except for SBP |
| 6 | in the oldest age group. While linear declining trends of active smoking prevalence were |
| 7 | observed for both male and female. Since the active smoking prevalence among female is |
| 8 | relatively low. It was decided a priori that we assumed zero active smoking prevalence among |
| 9 | female if the estimated coming active smoking prevalence would be lower than zero. |
| 10 | HDL and LDL trend analysis was not estimated from CHNS because serum lipid data were only |
| 11 | available for year 2009. We assumed HDL and LDL changes would be mediated by the BMI |
| 12 | trend. ^{1, 17} An increase of 1 kg/m ² in BMI was associated with 2.75 mg/dL increase in LDL and |
| 13 | 1.55 mg/dL decrease in HDL among male and 2.24 mg/dL increase in LDL and 0.77 mg/dL |
| 14 | decrease in HDL among female, respectively. In this model analyses, diabetes was defined as a |
| 15 | having a past diagnosis of diabetes, taking anti-diabetes medications, or a fasting glucose ≥ 126 |
| 16 | mg/dL. Since blood sample was collected ever since 2009 in CHNS, diabetes prevalence |
| 17 | recorded in the CHNS before 2009 might be underestimated without fasting glucose data. In |
| 18 | order to address this issue, we assumed diabetes awareness (the proportion of self-reported |
| 19 | diabetes among participants defined as diabetes) gradually increased over time. The number of |
| 20 | diabetes before 2009 was estimated using the following formula: the number of diabetes = self- |
| 21 | reported diabetes/diabetes awareness. Self-reported diabetes information was obtained from the |
| 22 | CHNS, while diabetes awareness data were from the China Cardiovascular Health Study and the |
| 23 | ChinaMUCA study, which defined diabetes in the same way as the CVD Policy Model. Then the |

1 prevalence of diabetes could be obtained as the proportion of the estimated number of diabetes 2 over the total number of subjects in CHNS. Based on the calculated diabetes prevalence, we projected diabetes trend accordingly. The age-adjusted prevalence of diabetes from the China 3 4 Cardiovascular Health study was 5.98% in 2000 and 8.33% from the China Cardiovascular Health Study and the ChinaMUCA in 2008. The awareness rate of diabetes grew from 36.1% to 5 59.8%. We assumed similar awareness change in the CHNS and then estimated diabetes 6 prevalence using linear regression. The diabetes prevalence was projected to increase yearly by 7 8 0.187% in male and 0.125% in female (Supplemental Table S3).

9 Effects of long term PM_{2.5} exposure

Reduction in PM_{2.5} air pollution levels was associated with decreased cardiovascular event 10 rates.¹⁸ However, no previous studies were conducted in China to explore the relationship 11 between long term PM_{2.5} exposure and health outcomes. Therefore, relative risks of CHD, stroke 12 and all-cause mortality associated with long term PM2.5 exposure were obtained from a meta-13 analysis of cohort studies.^{18, 19} Published studies addressing long term PM_{2.5} exposure with CHD, 14 stroke and all-cause mortality as outcomes were identified (Supplemental Table S4).²⁰⁻³⁶ If 15 multiple data derived from the same study, the study with the most incident cases was included. 16 Relative risks (RRs) or hazard ratios (HRs) and their 95% confidence intervals (CIs) were 17 extracted and uniformly standardized as 10 μ g/m³ increment of PM_{2.5}. The overall RRs and 95% 18 CIs were pooled using a random-effects model via the DerSimonian-Laird method. The RRs 19 (95% CIs) for a 10 μ g/m³ increase in long term PM_{2.5} exposure were 1.06 (1.03-1.08) for all-20 cause mortality, 1.19 (1.10-1.30) for CHD mortality and 1.07 (1.01-1.13) for stroke mortality 21 (Supplemental Figure S1-S3). These estimates were further incorporated into the model. 22 Though an integrated-exposure function³⁷ developed for Global Burden Disease Study showed a 23

1 non-linear $PM_{2.5}$ -CVD relationship by age, due to limitation of model's characteristics, we 2 assumed a uniform relative risk effect of $PM_{2.5}$ on all urban adults across age. It was likely to 3 over-estimate the effect sizes among those at the highest levels of air pollution exposure using 4 the linear function.

5 Epidemiologic input parameters and calibration

6 Prior to calibration (see below), CHD incidence in male and female aged 35-84 years with no prior CHD diagnosis was based on 10-year incidence rates from the China Hypertension 7 Epidemiology Follow Up Study (CHEFS)¹⁰ and calibrated to fit with CHD mortality and case-8 fatality assumptions. Incident stroke rates were also identified from the CHEFS.¹⁰ Main CVD 9 10 Policy Model-China 28-day case-fatality assumptions were estimated from pooled Beijing Sino-11 MONICA Study data from 1993-2004 (personal communication, Dong Zhao, MD, PhD, 2006) and the main age-specific CHD case-fatality rate assumptions were estimated from the overall 12 rates. Self-reported history of a physician-diagnosed myocardial infarction and/or stroke was 13 14 based on data from CHEFS. In CHEFS, each self-reported case of prevalent CVD was ascertained with chart review by study staff. Final epidemiologic parameter estimates are shown 15 in Supplemental Tables S5-6. 16

In order to evaluate the accuracy of CVD Policy Model predictions over time, China stroke and CHD mortality estimates for ages 35-84 years were obtained from the China CDC.¹³ In the calibration procedure, CHD and stroke parameters were calibrated separately. Starting with default incidence, case-fatality, and prevalence assumptions, the simulation model was run forward from year 2010 to 2016. Incidence and case-fatality inputs were iteratively calibrated primarily to match with age-specific mortality numbers in 2010 overall and within ten-year age

10

groups (Supplemental Tables S7-8). After CHD and stroke mortality were satisfactorily
 calibrated, age and sex specific non-cardiovascular death rates were also calibrated so that the
 totals of cardiovascular and non-cardiovascular deaths fitted within the envelope of all-cause
 mortality based on China CDC data.¹³

5 Monte Carlo Simulations

6 Markov Monte Carlo analyses were performed to estimate a range of uncertainty surrounding the results of projected air quality improvement and traditional risk factor intervention scenarios. We 7 assumed that the beta coefficient distributions of SBP, smoking and PM_{2.5} on CHD deaths and 8 stroke deaths were normally distributed. Standard deviations for the SBP and smoking beta 9 coefficients came from the CMCS study and the standard deviation for the PM_{2.5} beta coefficient 10 11 came from a meta-analysis of air pollution studies (Supplemental Table S2). The beta coefficient distributions for SBP, smoking, and PM_{2.5} were randomly and simultaneously 12 sampled 1,000 times in the Monte Carlo simulations. 13

14 Sensitivity Analysis

15 In the main analysis, no PM_{2.5} change in 2017-2030 was assumed as status quo case. However, the Global Burden of Disease - Major Air Pollution Sources (GBD MAPS) project has estimated 16 that PM_{2.5} in China would modestly reduce by 4 μ g/m³ from 2013 to 2030 under the business as 17 usual scenario (current legislation and implementation status as of end of 2012 and twelfth five-18 year plan for environmental protection).³⁸ Thus a sensitivity analysis was conducted assuming a 19 graded reduction trend over 2017-2030 as the base case. The starting level of PM_{2.5} remained 61 20 $\mu g/m^3$ in 2017, and it will slowly reduce to 57.9 $\mu g/m^3$ in 2030, with an average annual decrease 21 of 0.24 μ g/m³ estimated from GBD MAPS project. 22

| 1 | A linear PM _{2.5} -CVD morality relationship assumption might overestimate the health benefits, |
|---|--|
| 2 | thus additional sensitivity analysis was conducted to quantify the impact of attenuated relative |
| 3 | risk on health benefits. A 10% and 20% diminished beta-coefficient for the association between |
| 4 | PM _{2.5} and CHD and stroke death was recalculated. In recalculation for 10% diminished beta- |
| 5 | coefficient, the point estimate of $PM_{2.5}$ for CHD deaths changed from 0.0174 to 0.0157 and |
| 6 | stroke deaths changed from 0.0068 to 0.0061, and were further incorporated into the model. We |
| | |

7 re-run the CVD Policy Model-China (**Supplemental Table S10 and S11**).

| Year | Male | Female | Total |
|------|-------------|-------------|-------------|
| 2017 | 214,357,359 | 206,957,964 | 421,315,323 |
| 2018 | 221,214,027 | 213,805,560 | 435,019,587 |
| 2019 | 228,483,453 | 221,030,198 | 449,513,652 |
| 2020 | 236,269,514 | 228,703,794 | 464,973,308 |
| 2021 | 244,183,114 | 236,467,837 | 480,650,951 |
| 2022 | 252,549,397 | 244,626,274 | 497,175,671 |
| 2023 | 261,167,065 | 252,978,094 | 514,145,160 |
| 2024 | 269,637,173 | 261,153,768 | 530,790,940 |
| 2025 | 277,640,682 | 268,853,018 | 546,493,700 |
| 2026 | 284,503,701 | 275,452,872 | 559,956,573 |
| 2027 | 290,753,757 | 281,438,995 | 572,192,751 |
| 2028 | 296,427,057 | 286,839,505 | 583,266,562 |
| 2029 | 301,588,483 | 291,731,420 | 593,319,904 |
| 2030 | 306,302,815 | 296,167,648 | 602,470,463 |

Supplemental Table S1. Estimated China urban population aged 35-84 years old during 2017-2030 according to *World Population Prospects* by Population Division, United Nation.

| | SBP (1 mmHg) | | | Smoking (yes/no) | | | PM _{2.5} (1 μg/m ³) | | | | | |
|----------------|---------------------------|------------------------|---------------------------|------------------------|---------------------------|------------------------|--|------------------------|---------------------------|------------------------|---------------------------|------------------------|
| | СНІ |) death | Strok | e death | СНІ |) death | Strol | ke death | CHD death | | Stroke death | |
| | Main point estimate | Standard deviations | Main point estimate | Standard deviations | Main point estimate | Standard deviations | Main point estimate | Standard deviations | Main point estimate | Standard deviations | Main point estimate | Standard deviations |
| Males, years | | | | | | | | | | | | |
| 35-44 | .0338 | .0036 | .0472 | .0048 | .5640 | .1526 | .2550 | .1064 | .0174 | .0041 | .0068 | .0017 |
| 45-54 | .0302 | .0017 | .0422 | .0035 | .6940 | .1526 | .2550 | .1064 | .0174 | .0041 | .0068 | .0017 |
| 55-64 | .0271 | .0013 | .0368 | .0022 | .8240 | .1526 | .2550 | .1064 | .0174 | .0041 | .0068 | .0017 |
| 65-74 | .0221 | .0016 | .0283 | .0041 | .9540 | .1526 | .2550 | .1064 | .0174 | .0041 | .0068 | .0017 |
| 75-84 | .0161 | .0018 | .0162 | .0022 | 1.0840 | .1526 | .2550 | .1064 | .0174 | .0041 | .0068 | .0017 |
| Females, years | | | | | | | | | | | | |
| 35-44 | .0319 | .0036 | .0433 | .0048 | .5890 | .1526 | .4450 | .2707 | .0174 | .0041 | .0068 | .0017 |
| 45-54 | .0303 | .0015 | .0420 | .0032 | .8400 | .1526 | .4450 | .2707 | .0174 | .0041 | .0068 | .0017 |
| 55-64 | .0265 | .0013 | .0358 | .0022 | .9200 | .1526 | .4450 | .2707 | .0174 | .0041 | .0068 | .0017 |
| 65-74 | .0216 | .0015 | .0267 | .0021 | 1.1000 | .1526 | .4450 | .2707 | .0174 | .0041 | .0068 | .0017 |
| 75-84 | .0159 | .0017 | .0159 | .0021 | 1.1700 | .1526 | .4450 | .2707 | .0174 | .0041 | .0068 | .0017 |

Supplemental Table S2. Beta coefficients for CHD and stroke estimated from China Multi-provincial Cohort Study (CMCS) and standard deviations for SBP, smoking and PM_{2.5} for Monte Carlo simulation.

| | SBP (mmHg) | BMI (kg/m ²) | Smoking (%) | HDL (mg/dL) | LDL (mg/dL) | Diabetes (%) |
|---------------|---------------|-----------------------------|----------------|----------------|----------------|-----------------|
| Male, years | | | | | | +0.187 |
| 35-44 | +0.394 | +0.121 | -1.54 | -0.188 | +0.333 | |
| 45-54 | +0.381 | +0.117 | -0.63 | -0.181 | +0.322 | |
| 55-64 | +0.121 | +0.092 | -0.20 | -0.143 | +0.530 | |
| 65-84 | -0.026 | +0.088 | -0.09 | -0.136 | +0.242 | |
| Female, years | | | | | | +0.125 |
| 35-44 | +0.224 | +0.054 | -0.12 | -0.042 | +0.121 | |
| 45-54 | +0.263 | +0.087 | -0.19 | -0.067 | +0.195 | |
| 55-64 | +0.235 | +0.095 | -0.32 | -0.073 | +0.213 | |
| 65-84 | -0.096 | +0.111 | -0.12 | -0.085 | +0.249 | |

Supplemental Table S3. Annual future changes of traditional NCD risk factors were estimated based on China Health and Nutrition Survey.

| Study | Population | Time period | Average PM _{2.5} exposure level | RRs and 95% CIs with 10 µg/m ³ increment of PM _{2.5} | | | |
|--|---|-------------|---|--|------------------|------------------|--|
| | | - | (µg/m ³) | All-cause mortality | CHD mortality | Stroke mortality | |
| Harvard six cities ²⁰ , 2012 | Six cities in US | 1974-2009 | 16 | 1.14 (1.07-1.22) | | | |
| ACS, extended I ²¹ , 2004 | Adults in metropolitan areas in US | 1982-1998 | 17 | | | 1.02 (0.95-1.10) | |
| ACS, extended II ²² , 2009 | Adults in metropolitan areas in US | 1982-2000 | 14 | 1.06 (1.04-1.08) | 1.24 (1.20-1.29) | | |
| AHSMOG ²³ , 2000 | Nonsmoking, non-Hispanic whites | 1977-1992 | NA | 1.09 (0.97-1.21) | | | |
| VA study I ²⁴ , 2006 | Hypertensive male veterans in US | 1989-1996 | 19 | 1.15 (1.05-1.26) | | | |
| VA study II ²⁵ , 2006 | Hypertensive male veterans in US | 1997-2001 | 12 | 1.06 (0.94-1.22) | | | |
| CA CPS ²⁶ , 2005 | 11 counties in California, US | 1973-1982 | 23 | 1.04 (1.01-1.07) | | | |
| CA CPS ²⁶ , 2005 | 11 counties in California, US | 1983-2002 | 23 | 1.00 (0.98-1.02) | | | |
| WHI ²⁷ , 2007 | Postmenopausal women in 36 US metropolitan areas | 1994-2002 | 14 | | 2.21 (1.17-4.16) | 1.83 (1.11-3.00) | |
| HPFS ²⁸ , 2011 | Health professionals in US | 1989-2003 | 18 | 0.86 (0.72-1.02) | 0.98 (0.70-1.35) | | |
| NHS ²⁹ , 2009 | Registered nurses in US | 1992-2002 | 14 | 1.26 (1.02-1.54) | 2.02 (1.07-3.78) | | |
| NLCS ³⁰ , 2008 | Adults in 204 municipalities through Netherlands | 1987-1996 | 28 | 1.06 (0.97-1.16) | | | |
| California Teachers Study ³¹ , 2011 | Female public school teachers in California, US | 1997-2005 | 16 | | | 1.16 (0.92-1.46) | |
| California Teachers Study ³² , 2015 | Female public school teachers in California, US | 2001-2007 | 18 | 1.01 (0.97-1.05) | 1.19 (1.08-1.31) | | |
| US trucking industry cohort ³³ , 2011 | Male employed in US trucking industry | 1985-2000 | 14 | 1.10 (1.03-1.18) | | | |
| Canadian national cohort ³⁴ , 2012 | Nonimmigrant Canadian adults | 1991-2001 | 9 | 1.10 (1.05-1.15) | 1.30 (1.18-1.43) | 1.04 (0.93-1.16) | |
| Rome Longitudinal Study ³⁵ , 2013 | Italian population-based cohort | 2001-2010 | 23 | 1.04 (1.03-1.05) | 1.10 (1.06-1.13) | 1.08 (1.04-1.13) | |
| *ESCAPE study ³⁶ , 2014 | 22 cohorts across in Europe | 1985-2007 | 7~31 | | 0.98 (0.74-1.30) | 1.21 (0.87-1.69) | |

Supplemental Table S4. Prospective studies exploring risk of long term PM_{2.5} exposure and CHD, stroke and all-cause mortality.

US, the United States; ACS, American Cancer Society; AHSMOG, Adventist Health Study of Smog; VA, Veterans cohort; CA CPS, California Cancer Prevention Study; WHI, Women's Health Initiative; HPFS, Health Professionals Follow-up Study; NHS, Nurses' Health Study; NLCS, Netherlands Cohort Study on Diet and Cancer; ESCAPE, European Study of Cohorts for Air Pollution Effects.

*ESCAPE study includes 22 European cohorts using a standardized protocol for analysis.

| | CHD incidence rate per 100,000 | CHD 28 day case- fatality (proportion) | CHD mortality per 100,000 | Prevalence of prior myocardial infarction (proportion) |
|----------------|-----------------------------------|---|------------------------------|--|
| Males, years | | | | |
| 35-44 | 130 | 0.12 | 10 | 0.006 |
| 45-54 | 135 | 0.21 | 36 | 0.012 |
| 55-64 | 220 | 0.29 | 97 | 0.034 |
| 65-74 | 500 | 0.33 | 243 | 0.047 |
| 75-84 | 2,010 | 0.48* | 1,104 | 0.060* |
| Females, years | | | | |
| 35-44 | 19 | 0.18 | 1 | 0.004 |
| 45-54 | 49 | 0.23 | 20 | 0.013 |
| 55-64 | 141 | 0.27 | 43 | 0.031 |
| 65-74 | 310 | 0.43 | 160 | 0.040 |
| 75-84 | 1,900 | 0.51* | 1,028 | 0.060* |

Supplemental Table S5. Coronary Heart Disease (CHD) Inputs used for the CVD Policy Model-China

*Estimate not available from original source data and imputed using linear interpolation.

| | Total stroke incidence rate per 100,000 | Total stroke 28 day case-fatality (proportion) | Total stroke mortality per 100,000 | Prevalence of prior stroke (proportion) |
|----------------|--|--|---------------------------------------|--|
| Males, years | | | | |
| 35-44 | 24 | 0.25 | 20 | 0.013 |
| 45-54 | 145 | 0.18 | 62 | 0.032 |
| 55-64 | 670 | 0.12 | 151 | 0.088 |
| 65-74 | 1,250 | 0.20 | 502 | 0.142 |
| 75-84 | 2,510 | 0.45* | 1,708 | 0.150* |
| Females, years | | | | |
| 35-44 | 23 | 0.18 | 10 | 0.009 |
| 45-54 | 180 | 0.14 | 30 | 0.024 |
| 55-64 | 800 | 0.15 | 131 | 0.060 |
| 65-74 | 1,500 | 0.20 | 375 | 0.100 |
| 75-84 | 2,500 | 0.45* | 1.359 | 0.120* |

Supplemental Table S6. Stroke Inputs used for the CVD Policy Model-China

*Estimate not available from original source data and imputed using linear interpolation.

| Supplemental Table 87 | Dro collibration and | nost colibration | CUD incidence | and 28 day | and fotality | innuta |
|------------------------|----------------------|------------------|----------------|------------|--------------|--------|
| Supplemental Table 57. | 1 re-canor anon and | post-canor ation | CIID incluence | anu 20 uay | case-latanty | inputs |

| | CHD incidence rate per 100,000 CHD 28 day case-fatality (proportion) | | | | |
|----------------|--|---|---|---|--|
| | Pre-calibration (based on CHEFS, 1991-2000, ICD9 430-438) | Post-calibration (identical CVDPM definition) | Pre-calibration (based on Sino-Monica Beijing) | Post-calibration (identical CVDPM definition) | |
| Males, years | | | | | |
| 35-44 | 54 | 130 | 0.12 | 0.05 | |
| 45-54 | 112 | 135 | 0.21 | 0.15 | |
| 55-64 | 342 | 220 | 0.29 | 0.18 | |
| 65-74 | 540 | 500 | 0.33 | 0.25 | |
| 75-84 | 889 | 2,010 | 0.48* | 0.50 | |
| Females, years | | | | | |
| 35-44 | 23 | 19 | 0.18 | 0.14 | |
| 45-54 | 96 | 49 | 0.23 | 0.20 | |
| 55-64 | 188 | 141 | 0.27 | 0.20 | |
| 65-74 | 368 | 310 | 0.43 | 0.46 | |
| 75-84 | 752 | 1,900 | 0.51* | 0.47 | |

*Estimate not available from original source data and imputed using linear interpolation.

Supplemental Table S8. Pre-calibration and post-calibration stroke incidence and 28 day case-fatality inputs

| | Stroke incidence rate per 10 | 00,000 | Stroke 28 day case-fatality (proportion) | | |
|----------------|---|---|---|--|--|
| | Pre-calibration (based on CHEFS, 1991-2000, ICD9 410-414) | Post-calibration (identical CVDPM definition) | Pre-calibration (based on Sino-Monica Beijing) | Post-calibration (identical CVDPM definition) | |
| Males, years | | | | | |
| 35-44 | 91 | 24 | 0.25 | 0.19 | |
| 45-54 | 240 | 145 | 0.18 | 0.17 | |
| 55-64 | 711 | 670 | 0.12 | 0.12 | |
| 65-74 | 1,292 | 1,250 | 0.20 | 0.25 | |
| 75-84 | 1,904 | 2,510 | 0.45* | 0.48 | |
| Females, years | | | | | |
| 35-44 | 59 | 23 | 0.18 | 0.11 | |
| 45-54 | 176 | 180 | 0.14 | 0.11 | |
| 55-64 | 424 | 800 | 0.15 | 0.12 | |
| 65-74 | 848 | 1,500 | 0.20 | 0.22 | |
| 75-84 | 1,500 | 2,500 | 0.45* | 0.38 | |

*Estimate not available from original source data and imputed using linear interpolation.

Supplemental Table S9. Sensitivity analysis for projected CHD and stroke deaths averted with hypothetical air pollution controls in urban Chinese population aged 35-84 years over 2017-2030 with business as usual scenario as status quo case

| | CHD Deaths (thousands) | Averted CHD Deaths (thousands) | Stroke Deaths (thousands) | Averted Stroke Deaths (thousands) | Life years gained (thousands) |
|--|---------------------------|-----------------------------------|------------------------------|--------------------------------------|----------------------------------|
| Alternative Status quo case† | 7,604 (7,450-7,863) | - | 10,902 (10,355-11,528) | - | |
| PM _{2.5} improvement scenarios* | | | | | |
| Target 1: Beijing Olympic Games | 7,462 (7,172-7,726) | 143 (120-299) | 10,824 (10,233-11,451) | 78 (58-169) | 1,115 (1,281-1,950) |
| Target 2: China Class II standard limit | 6,216 (5,537-6,931) | 1,388 (886-1,982) | 10,080 (9,385-10,839) | 823 (449-1,246) | 11,620 (9,738-14,355) |
| Target 3: WHO recommended level | 5,031 (4,122-6,089) | 2,574 (1,701-3,403) | 9,240 (8,330-10,314) | 1,663 (914-2,399) | 23,313 (19,526-27,682) |
| Comparison scenarios* | | | | | |
| 25% reduction in uncontrolled systolic hypertension (to <140 mmHg) | 6,915 (6,823-7,107) | 689 (555-859) | 9,657 (9,227-10,112) | 1,245 (869-1,674) | 9,911 (8,750-11,327) |
| 30% reduction in tobacco use | 7,210 (6,984-7,519) | 394 (258-542) | 10,788 (10,146-11,505) | 115 (5-225) | 3,034 (2,429-3,708) |
| 25% reduction in uncontrolled systolic hypertension (to <140 mmHg) plus 30% reduction in tobacco use | 6,560 (6,369-6,813) | 1,044 (866-1,255) | 9,559 (9,029-10,110) | 1,343 (948-1,781) | 12,774 (11,435-14,275) |

[†] Alternative status quo case scenario (PM_{2.5} remained 61 μ g/m³ in 2017, and it will slowly reduce to 57.9 μ g/m³ in 2030)

*Each scenario is compared with the status quo case. Ninety-five percent uncertainty intervals were calculated from the results of 1,000 probabilistic simulations.

Supplemental Table S10. Sensitivity analysis for projected CHD and stroke deaths averted with hypothetical air pollution controls in urban Chinese population aged 35-84 years over 2017-2030 with 10% attenuated PM_{2.5}-CVD health effects

| | CHD Deaths (thousands) | Averted CHD Deaths (thousands) | Stroke Deaths (thousands) | Averted Stroke Deaths (thousands) | Life years gained (thousands) |
|--|---------------------------|-----------------------------------|------------------------------|--------------------------------------|-------------------------------|
| Status quo case (remain current PM _{2.5} level) | 7,910 (7,741-8,094) | - | 11,069 (10,219-11,611) | - | |
| PM _{2.5} improvement scenarios* | | | | | |
| Target 1: Beijing Olympic Games | 7,514 (7,291-7,693) | 397 (206-616) | 10,857 (10,049-11,406) | 212 (92-323) | 3,177 (2,629-3,822) |
| Target 2: China Class II standard limit | 6,370 (5,672-7,016) | 1,540 (845-2,258) | 10,187 (9,408-10,814) | 882 (392-1,317) | 13,114 (11,018-15,467) |
| Target 3: WHO recommended level | 5,253 (4,241-6,332) | 2,657 (1,546-3,664) | 9,422 (8,658-10,467) | 1,647 (758-2,397) | 24,280 (20,744-27,809) |
| Comparison scenarios* | | | | | |
| 25% reduction in uncontrolled systolic hypertension (to <140 mmHg) | 7,186 (7,117-7,235) | 724 (591-905) | 9,801 (9,159-10,129) | 1,268 (808-1,642) | 10,068 (8,929-11,237) |
| 30% reduction in tobacco use | 7,498 (7,257-7,704) | 412 (306-562) | 10,953 (9,955-11,563) | 116 (30-273) | 3,095 (2,623-3,831) |
| 25% reduction in uncontrolled systolic hypertension (to <140 mmHg) plus 30% reduction in tobacco use | 6,815 (6,629-6,899) | 1,095 (953-1,265) | 9,701 (8,924-10,102) | 1,368 (958-1,726) | 12,988 (12,036-14,320) |

*Each scenario is compared with the status quo case. Ninety-five percent uncertainty intervals were calculated from the results of 1,000 probabilistic simulations.

Supplemental Table S11. Sensitivity analysis for projected CHD and stroke deaths averted with hypothetical air pollution controls in urban Chinese population aged 35-84 years over 2017-2030 with 20% attenuated PM_{2.5}-CVD health effects

| | CHD Deaths (thousands) | Averted CHD Deaths (thousands) | Stroke Deaths (thousands) | Averted Stroke Deaths (thousands) | Life years gained (thousands) |
|--|---------------------------|-----------------------------------|---------------------------|--------------------------------------|----------------------------------|
| Status quo case (remain current PM _{2.5} level) | 7,922 (7,780-8,066) | - | 11,077 (10,547-11,305) | - | |
| PM _{2.5} improvement scenarios* | | | | | |
| Target 1: Beijing Olympic Games | 7,570 (7,265-7,933) | 352 (100-551) | 10,890 (10,348-11,132) | 187 (95-326) | 2,967 (2,128-3,803) |
| Target 2: China Class II standard limit | 6,539 (5,718-7,608) | 1,383 (422-2,050) | 10,295 (9,530-10,727) | 782 (409-1,326) | 12,303 (8,970-15,437) |
| Target 3: WHO recommended level | 5,504 (4,409-7,227) | 2,418 (800-3,377) | 9,608 (8,493-10,305) | 1,470 (789-2,411) | 22,892 (17,011-28,081) |
| Comparison scenarios* | | | | | |
| 25% reduction in uncontrolled systolic hypertension (to <140 mmHg) | 7,198 (7,119-7,307) | 724 (595-850) | 9,809 (9,291-10,135) | 1,269 (877-1,490) | 10,069 (8,907-10,925) |
| 30% reduction in tobacco use | 7,509 (7,323-7,719) | 413 (266-508) | 10,961 (10,339-11,277) | 116 (34-239) | 3,097 (2,584-3,666) |
| 25% reduction in uncontrolled systolic hypertension (to <140 mmHg) plus 30% reduction in tobacco use | 6,826 (6,683-6,996) | 1,096 (926-1,244) | 9,709 (9,088-10,106) | 1,368 (998-1,618) | 12,991 (11,636-13,907) |

*Each scenario is compared with the status quo case. Ninety-five percent uncertainty intervals were calculated from the results of 1,000 probabilistic simulations.

Figure Legends

Supplemental Figure S1. Relative risks for each 10 μ g/m³ increment in long term PM_{2.5} exposure and risk of all-cause mortality.

The horizontal lines represent 95% confidence interval and grey squares represent the weights of each study in random effect models.

Supplemental Figure S2. Relative risks for each 10 μ g/m³ increment in long term PM_{2.5} exposure and risk of coronary heart disease mortality.

The horizontal lines represent 95% confidence interval and grey squares represent the weights of each study in random effect models.

Supplemental Figure S3. Relative risks for each 10 μ g/m³ increment in long term PM_{2.5} exposure and risk of stroke mortality.

The horizontal lines represent 95% confidence interval and grey squares represent the weights of each study in random effect models.

Supplemental Figure S1. Relative risks for each 10 μ g/m³ increment in long term PM_{2.5} exposure and risk of all-cause mortality

| Study | | % |
|--|---------------------|--------|
| D | RR (95% CI) | Weight |
| Harvard Six-Cities | • 1.14 (1.07, 1.22) | 6.20 |
| ACS extended II | 1.06 (1.04, 1.08) | 13.50 |
| AHSMOG - | 1.09 (0.97, 1.21) | 2.97 |
| VA study I | • 1.15 (1.05, 1.26) | 4.00 |
| VA study II | 1.06 (0.94, 1.22) | 2.26 |
| CA CPS - | 1.04 (1.01, 1.07) | 11.82 |
| CA CPS 🔶 | 1.00 (0.98, 1.02) | 13.33 |
| HPFS | 0.86 (0.72, 1.02) | 1.36 |
| NHS | 1.26 (1.02, 1.54) | 1.00 |
| NLCS - | 1.06 (0.97, 1.16) | 4.11 |
| California Teachers Study | 1.01 (0.97, 1.05) | 9.91 |
| US trucking industy cohort | 1.10 (1.03, 1.18) | 5.94 |
| Canadian national cohort | 1.10 (1.05, 1.15) | 8.93 |
| Rome Longitudinal Study | 1.04 (1.03, 1.05) | 14.66 |
| Overall (I-squared = 74.1%, p = 0.000) | 1.06 (1.03, 1.08) | 100.00 |
| NOTE: Weights are from random effects analysis | | |
| 7 1 | 16 | |

Supplemental Figure S2. Relative risks for each 10 μ g/m³ increment in long term PM_{2.5} exposure and risk of coronary heart disease mortality



Supplemental Figure S3. Relative risks for each 10 μ g/m³ increment in long term PM_{2.5} exposure and risk of stroke mortality



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