THE LANCET

Supplementary appendix

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Supplement to: Chen Z, Peto R, Zhou M, et al, for the China Kadoorie Biobank (CKB) collaborative group. Contrasting male and female trends in tobacco-attributed mortality in China: evidence from successive nationwide prospective cohort studies. *Lancet* 2015; **386:** 1447–56.

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Webtable 1: ICD codes for specific causes of death _

| Category | ICD-9 | ICD-10 |
|---|-------------------------------|-------------------------------|
| Lung cancer (including trachea) | 162 | C33-C34 |
| Liver cancer | 155 | C22 |
| Stomach cancer | 151 | C16 |
| Desophagus cancer | 150 | C15 |
| Five minor sites (Lip/oral cavity, larynx, pharynx, pancreas and bladder) | 140-149, 157, 161, 188 | C00-C14, C25, C32, C67 |
| Cancer, other or unknown | Rest of 140-208 | Rest of C00-C97 |
| All cancer | 140-208 | C00-C97 |
| schaemic heart disease | 410-414 | 120-125 |
| schaemic stroke | 433-435 | 163 |
| Haemorrhagic stroke | 430-432 | I61 |
| Other or unknown type of stroke | 436-438 | Rest of 160-169 |
| Other cardiovascular | Rest of 410-438, exc. 415-417 | Rest of 100-199, exc. 126-127 |
| All vascular | 410-438, exc. 415-417 | 100-199 exc. 126-127 |
| COPD | 490-496, 415-417 | J40-J44, I26-I27 |
| Other respiratory | Rest of 460-519 | Rest of J00-J99 |
| All respiratory | 460-519, 415-417 | J00-J99, I26-I27 |
| All other medical | Rest of 001-799 | Rest of A00-R99 |
| All non-medical | 800-999, E800-E999, V01-V82 | V01-Y98 |

Webtable 2: Number of deaths at age 40-79, mortality RRs and proportion of deaths attributed to smoking among men in ~1995 (CPSS, 1991-99) and men and women in ~2010 (CKB, 2006-14)

| Cause of deaths | | nen only y, 1991-99) | | 3 men udy, 2006-14) | CKB women (Second study, 2006-14) | | | | |
|---------------------|-------------------------------------|--------------------------------|-------------------------|-------------------------------------|--------------------------------------|-------------------------|-------------------------------------|-----------------------------|-------------------------|
| | No of deaths Smokers/Non-smokers | RR [†] (95% CI) | PAF [‡] (%) | No of deaths Smokers/Non-smokers | RR [†] (95% CI) | PAF [‡] (%) | No of deaths Smokers/Non-smokers | RR [†] (95% CI) | PAF [‡] (%) |
| Lung cancer | 1024/232 | 1.95 (1.68 - 2.26) | 39 | 1073/161 | 2.58 (2.17 - 3.05) | 51 | 101/618 | 2.56 (2.02 - 3.26) | 9 |
| Liver cancer | 977/374 | 1.06 (0.93 - 1.20) | 3 | 644/176 | 1.26 (1.06 - 1.50) | 16 | 31/382 | 1.40 (0.93 - 2.12) | 2 |
| Stomach cancer | 870/273 | 1.22 (1.06 - 1.41) | 14 | 602/170 | 1.25 (1.05 - 1.49) | 14 | 26/334 | 1.43 (0.92 - 2.23) | 2 |
| Oesophagus cancer | 662/233 | 1.18 (1.01 - 1.38) | 11 | 567/124 | 1.58 (1.30 - 1.93) | 28 | 13/236 | 1.05 (0.55 - 2.01) | 0 |
| Five minor sites | 215/78 | 1.18 (0.90 - 1.54) | 11 | 282/62 | 1.53 (1.15 - 2.04) | 27 | 14/210 | 0.98 (0.55 - 1.75) | -1 |
| All cancer | 4201/1352 | 1.28 (1.20 - 1.36) | 16 | 3811/913 | 1.51 (1.40 - 1.63) | 26 | 272/3037 | 1.58 (1.37 - 1.81) | 3 |
| IHD | 1172/406 | 1.26 (1.12 - 1.42) | 16 | 1261/426 | 1.38 (1.23 - 1.54) | 19 | 130/1212 | 1.74 (1.42 - 2.12) | 5 |
| Ischaemic stroke | 817/333 | 1.11 (0.97 - 1.27) | 7 | 395/132 | 1.41 (1.15 - 1.73) | 19 | 25/353 | 1.15 (0.74 - 1.78) | 1 |
| Haemorrhagic stroke | 2437/816 | 1.06 (0.98 - 1.16) | 5 | 1251/397 | 1.03 (0.92 - 1.16) | 1 | 88/1262 | 1.09 (0.86 - 1.39) | 1 |
| All cardiovascular | 6277/2203 | 1.12 (1.07 - 1.18) | 8 | 3578/1193 | 1.24 (1.16 - 1.33) | 13 | 319/3530 | 1.44 (1.27 - 1.63) | 3 |
| All respiratory | 5193/1504 | 1.27 (1.20 - 1.35) | 16 | 1068/244 | 1.64 (1.42 - 1.90) | 31 | 159/710 | 1.78 (1.46 - 2.17) | 8 |
| All other medical | 2055/752 | 1.07 (0.98 - 1.17) | 5 | 1062/360 | 1.20 (1.06 - 1.36) | 12 | 110/1133 | 1.41 (1.13 - 1.76) | 3 |
| All causes | 18875/6236 | 1.17 (1.14 - 1.21) | 11 | 10326/2955 | 1.33 (1.28 - 1.39) | 18 | 897/9037 | 1.51 (1.40 - 1.63) | 3 |

^{*} Excludes 4306 and 15,271 (14,080 male) ever regular smokers who stopped smoking by choice in CPSS and CKB respectively.

[†] Rate ratios (RRs) were all adjusted for area and age at risk (strata), education and alcohol. Additional adjustment for SBP ((1.18 [1.16-1.20], 1.35 [1.32-1.37], 1.56 [1.46-1.67] for, respectively CPSS men, CKB men and CKB women), for BMI ((1.15 [1.13-1.17], 1.30 [1.27-1.32], 1.48 [1.39-1.58] for, respectively CPSS men, CKB men and CKB women) and for SBP and BMI ((1.15 [1.13-1.17], 1.30 [1.27-1.33], 1.53 [1.43-1.63] for, respectively CPSS men, CKB men and CKB women) hardly changed the RRs for overall mortality.

[‡] In estimating Population Attributable Fraction (PAF) for ever regular smoking, the RR used is for all ever regular smokers including those who stopped by choice.

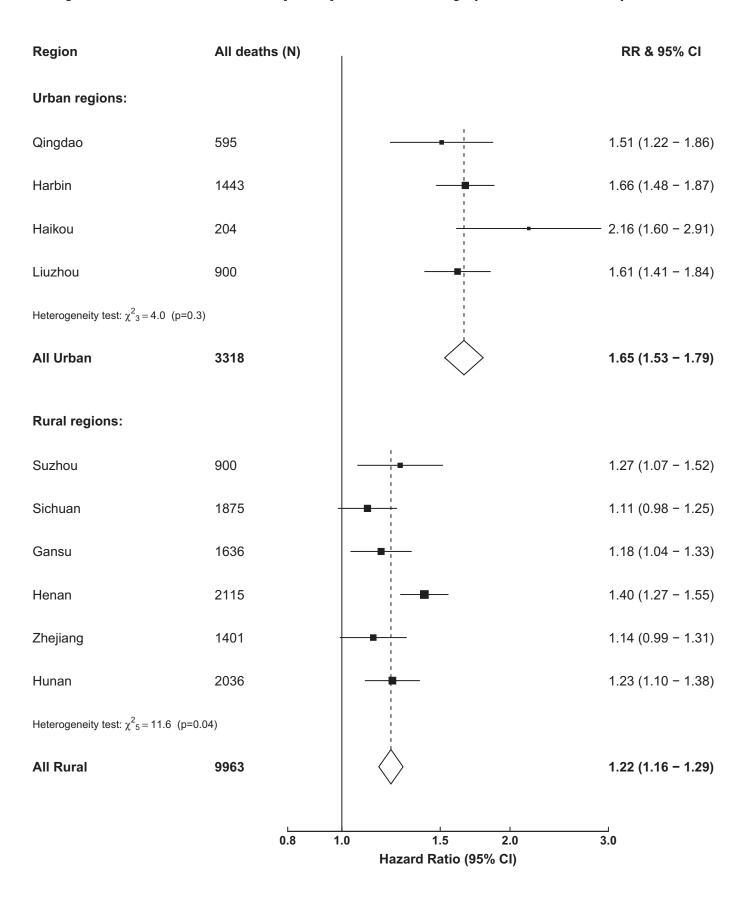
Webtable 3: Baseline characteristics of men at age 40-79 in the first (CPSS, 1991) versus second study (CKB, 2004-08)

| Characteristics | CPSS (n=219,893) | CKB * (n=180,658) |
|--|-------------------------|--------------------------|
| % urban | 27.1 | 43.7 |
| Year of baseline survey (median) | 1991 | 2006 |
| Year of birth (median) | 1938 | 1952 |
| Age at baseline (%) | | |
| 40-49 | 37.8 | 32.8 |
| 50-59 | 31.6 | 35.3 |
| 60-69 | 21.4 | 22.9 |
| 70-79 | 9.2 | 9.1 |
| Mean age (years) | 54.8 | 55.4 |
| Highest education (%) | | |
| None | 22.7 | 10.0 |
| Primary | 44.2 | 35.5 |
| Secondary | 27.8 | 47.4 |
| Tertiary | 5.3 | 7.0 |
| Current alcohol drinking in most weeks (%) | 33.4 | 33.2 |
| Smoking category (%) | | |
| Never smoked regularly | 26.8 | 24.7 |
| Ex-smoker (stopped by choice) | 2.0 | 7.2 |
| [Ex-smoker (stopped by choice) as % of ever-smoker] | [2.7] | [9.6] |
| Smoker (current smoker or stopped because ill) | 71.2 | 68.1 |
| – Current smoker | 67.4 | 60.8 |
| – Stopped because ill | 3.8 | 7.3 |
| Patterns among smokers # | | |
| Age started (%) | | |
| <20 | 30.2 | 32.2 |
| 20-24 | 41.6 | 37.2 |
| ≥25 | 28.2 | 30.6 |
| First started on cigarettes, by birth year (%) | | |
| <1925 | 41.5 | _ |
| 1925-1934 | 52.0 | 56.8 |
| 1935-1944 | 60.1 | 64.0 |
| 1945-1954 | 69.1 | 75.2 |
| ≥1955 | _ | 89.2 |
| Last smoked cigarettes only, by age (%) | | |
| 40-49 | 67.8 | 92.9 |
| 50-59 | 54.9 | 83.0 |
| 60-69 | 43.2 | 73.2 |
| 70-79 | 31.5 | 71.4 |
| Any age | 55.4 | 83.4 |
| Mean cigarettes/day/pure cigarette smoker of any age | 16.9 | 18.1 |

^{*} Excluded 29,564 men aged <40 at entry

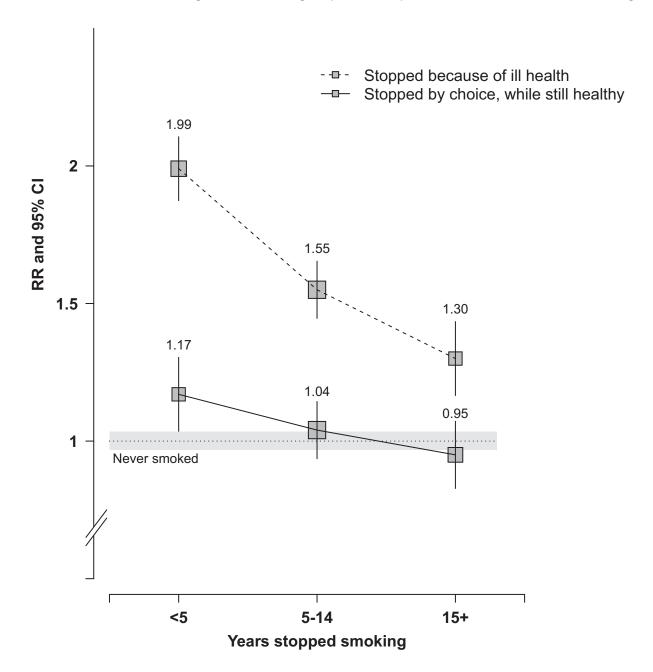
[#] Current smoker or stopped because ill

Webfigure 1: Adjusted rate ratios for all-cause mortality in men, by area, in the second prospective study (CKB, 2006-14)



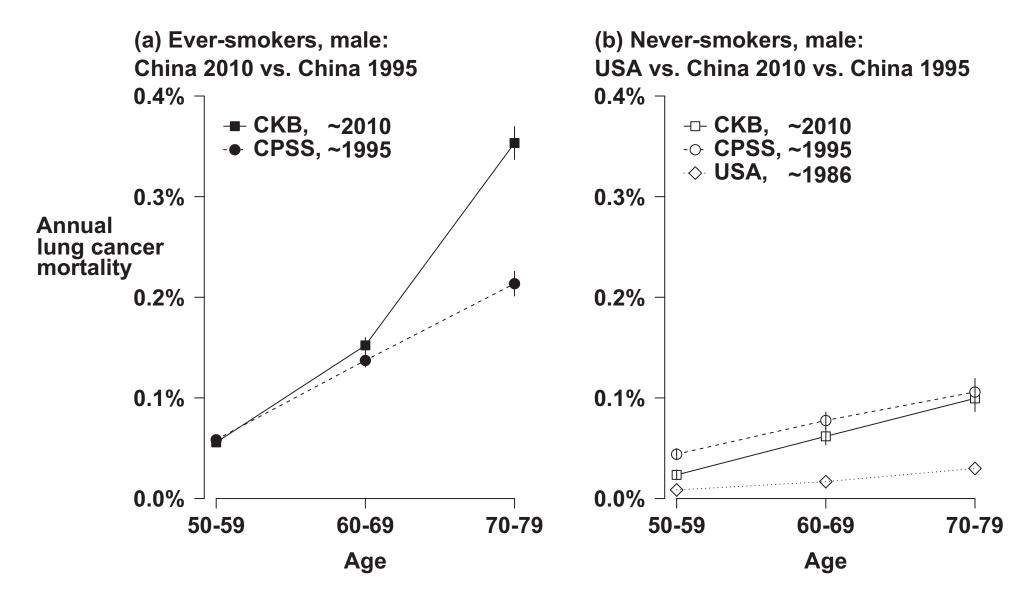
Webfigure 2: Ex- versus never-smoker all-cause mortality rate ratio, by years stopped smoking and reason stopped; men in both studies

Each group-specific CI (including that for never-smokers, given by the width of the shaded strip) reflects the variance of the log risk in that 1 group, so comparisons use variances from >1 group.



Webfigure 3: Changing Chinese male lung cancer mortality, 1995-2010, among ever-smokers and never-smokers in two nationally representative prospective studies about 15 years apart

NB: Rates in later middle age are increasing in Chinese smokers, decreasing in Chinese never-smokers, and roughly constant in US never-smokers. CKB, Chinese Kadoorie Biobank, follow-up ~2006-2014; CPSS, Chinese Prospective Smoking Study, follow-up ~1991-1999; USA, ACS CPS-II, follow-up 1984-1988 [3]. Non-smoker rates: webtables 4-5.



Webtable 4: Chinese never-smoker lung cancer death rates by age and sex in two successive prospective studies 15 years apart (1991-99 and 2006-14)

| | | le, 1991 I-year 1 | | Male, 2006-14 (mid-year 2010) | | | Female, 2006-14 (mid-year 2010) | | | |
|------------------------|-----------------|----------------------|------------------|----------------------------------|-------------------|------------------|------------------------------------|-------------------|------------------|--|
| Age at risk (in years) | Person years | Lung ca deaths | Rate / 100000 | Person years | Lung ca deaths | Rate / 100000 | Person years | Lung ca deaths | Rate / 100000 | |
| 40-44 | 32391 | 5 | 10 | 56656 | 6 | 10 | 321656 | 10 | 8 | |
| 45-49 | 76630 | 16 | 18 | 52234 | 5 | | 341274 | 45 | | |
| 50-54 | 71812 | 31 | 4.4 | 46190 | 7 | 26 | 328126 | 51 | 21 | |
| 55-59 | 68713 | 31 | 44 | 51852 | 19 | | 350710 | 91 | | |
| 60-64 | 62924 | 46 | 78 | 46243 | 22 | 66 | 259817 | 99 | 54 | |
| 65-69 | 49204 | 41 | 76 | 39430 | 33 | 66 | 176555 | 123 | 54 | |
| 70-74 | 35752 | 34 | 109 | 36527 | 37 | 125 | 131322 | 116 | 107 | |
| 75-79 | 22810 | 28 | | 21490 | 32 | 125 | 65702 | 83 | | |
| Total, by summation | 420235 | 232 | | 350621 | 161 | | 1975162 | 618 | | |

Never-smokers include those who reported at baseline having smoked occasionally, but never regularly.

The crude rate in each age group is $100,000 \times$ the number of lung cancer deaths / person-years.

The rate in each 10-year age group is the mean of the two crude rates in the component 5-year age groups.

Webtable 5: US never-smoker lung cancer mortality rates by age and sex in the American Cancer Society's Second Cancer Prevention Study, CPS-II, during years 3-6 of the prospective follow-up (approximately 1984-88)

| | | lale, 1984- iid-year 19 | | Female, 1984-88 (mid-year 1986) | | | |
|---------------------------|-----------------|----------------------------|------------------------|------------------------------------|--|----------|--|
| Age at risk (in years) | Person years | | ng cancer / 100,000 | Person years | Annual lung cancer mortality / 100,000 | | |
| (,) | 700 | Crude rate | Smoothed | | Crude rate | Smoothed | |
| 35-39 | 14560 | 7 | 2 | 30810 | 3 | 2 | |
| 40-44 | 13930 | 0 | 3 | 45760 | 0 | 3 | |
| 45-49 | 32110 | 9 | 5 | 116870 | 4 | 4 | |
| 50-54 | 81410 | 5 | 7 | 196520 | 5 | 7 | |
| 55-59 | 87920 | 3 | 10 | 224110 | 8 | 10 | |
| 60-64 | 80570 | 11 | 14 | 227400 | 14 | 14 | |
| 65-69 | 69670 | 24 | 20 | 192560 | 18 | 19 | |
| 70-74 | 50530 | 36 | 27 | 143280 | 28 | 26 | |
| 75-79 | 28880 | 38 | 35 | 95900 | 42 | 34 | |

Source: Appendix to Reference (3). Never-smoker means replied "never smoked regularly" at baseline survey.

The crude rate in each age group is 100,000 × the number of lung cancer deaths / person-years.

Smoothing the lung cancer mortality rates in the 5-year age groups with lower limits 35, 40, ..., 75 involved obtaining the best-fitting line of slope 4 on a logarithmic graph of the crude rates vs these lower age limits; then, in Webfigure 3, the US rates at ages 50-59, 60-69 and 70-79 are each the mean of two smoothed rates.