

# Breastfeeding and the Risk of Maternal Cardiovascular Disease: A Prospective Study of 300 000 Chinese Women

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**Background**—Breastfeeding confers substantial benefits to child health and has also been associated with lower risk of maternal cardiovascular diseases (CVDs) in later life. However, the evidence on the effects of CVD is still inconsistent, especially in East Asians, in whom the frequency and duration of breastfeeding significantly differ from those in the West.

**Methods and Results**—In 2004–2008, the nationwide China Kadoorie Biobank recruited 0.5 million individuals aged 30 to 79 years from 10 diverse regions across China. During 8 years of follow-up, 16 671 incident cases of coronary heart disease and 23 983 cases of stroke were recorded among 289 573 women without prior CVD at baseline. Cox regression yielded adjusted hazard ratios (HRs) and 95% CIs for incident CVD by breastfeeding. Overall,  $\approx 99\%$  of women had given birth, among whom 97% reported a history of breastfeeding, with a median duration of 12 months per child. Compared with parous women who had never breastfed, ever breastfeeding was associated with a significantly lower risk of CVD, with adjusted HRs of 0.91 (95% CI, 0.84–0.99) for coronary heart disease and 0.92 (95% CI, 0.85–0.99) for stroke. Women who had breastfed for  $\geq 24$  months had an 18% (HR, 0.82; 0.77–0.87) lower risk of coronary heart disease and a 17% (HR, 0.83; 0.79–0.87) lower risk of stroke compared with women who had never breastfed. Among women who ever breastfed, each additional 6 months of breastfeeding per child was associated with an adjusted HR of 0.96 (95% CI, 0.94–0.98) for coronary heart disease and 0.97 (95% CI, 0.96–0.98) for stroke.

**Conclusions**—Among Chinese women, a history of breastfeeding was associated with an  $\approx 10\%$  lower risk of CVD in later life and the magnitude of the inverse association was stronger among those with a longer duration of breastfeeding. (*J Am Heart Assoc.* 2017;6:e006081. DOI: 10.1161/JAHA.117.006081.)

**Key Words:** breastfeeding • cardiovascular disease • China • epidemiology • risk factor • women

Pregnancy causes substantial changes to the maternal cardiometabolic system, including weight gain, increased insulin resistance, and higher levels of circulating lipids.<sup>1,2</sup> Such pregnancy-related cardiometabolic changes may reverse more quickly and more completely with breastfeeding, with

several studies reporting that women who breastfeed have more favorable cardiometabolic profiles compared with women who do not breastfeed.<sup>3–5</sup> Longer breastfeeding duration has also been associated with a lower risk of metabolic syndrome,<sup>6</sup> hypertension,<sup>7,8</sup> and diabetes mellitus

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Accompanying Tables S1 through S3 and Figures S1 and S2 are available at <http://jaha.ahajournals.org/content/6/6/e006081/DC1/embed/inline-supplementalry-material-1.pdf>

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## Clinical Perspective

### What Is New?

- While breastfeeding confers substantial benefits to child health, its effects on maternal cardiovascular health remain uncertain.
- This large prospective cohort study among nearly 300 000 women in China showed that, among parous women, a history of breastfeeding was associated with an  $\approx 10\%$  lower risk of several major cardiovascular diseases in later life.
- Among women who had ever breastfed, each additional 6 months of breastfeeding was associated with a further  $\approx 3\%$  to  $4\%$  lower cardiovascular disease risk.

### What Are the Clinical Implications?

- If causal, these findings suggest that interventions to increase the likelihood and duration of breastfeeding could have persistent benefits to maternal cardiovascular health.

in later life,<sup>9</sup> which may lead to long-term protective benefits for cardiometabolic diseases.

A few previous studies of mostly Western populations have assessed the association between breastfeeding duration and maternal risk of cardiovascular diseases (CVDs),<sup>10–14</sup> but with inconclusive findings about the direction, magnitude, and shape of any putative relationship. Some of these discrepancies may be explained by methodological differences between studies, such as variation in study design, small sample sizes, different exposure definitions, or varying levels of covariate adjustment. Moreover, some studies were restricted to fatal CVD end points,<sup>10,11,14</sup> whereas others only examined specific types of CVD,<sup>12,13</sup> typically coronary heart disease (CHD), without evaluating other major types of incident CVD, such as ischemic and hemorrhagic stroke.

Major differences in breastfeeding practices, between and within populations, may also explain some of the discordance in study findings.<sup>15</sup> Breastfeeding initiation rates are typically considerably higher and breastfeeding duration is typically longer in women from low- and middle-income countries, compared with those in high-income countries. In China, breastfeeding is almost universal and many women continue to breastfeed their infants for long durations, particularly in older generations. While breastfeeding practices in China have changed considerably over recent decades, with declines in both the likelihood and duration of breastfeeding, they remain significantly different from those in the West.<sup>16</sup> There is limited evidence about the long-term relevance of breastfeeding on the risk of CVD in a contemporary Chinese population, overall or in different population subgroups. We examined the relationship between breastfeeding and the risk of several major CVDs among 300 000 women from the prospective China Kadoorie Biobank study.<sup>17</sup>

## Methods

### Study Population

Detailed information about the study design and procedures of China Kadoorie Biobank has been previously reported.<sup>17</sup> Briefly, between 2004 and 2008, 302 669 women and 210 222 men aged 35 to 79 years were recruited from 5 urban and 5 rural areas of China. At the baseline study assessment clinics, trained health workers administered a laptop-based questionnaire that covered demographic and socioeconomic status, lifestyle factors, and medical history. Detailed information on women's reproductive factors was also collected, including age at menarche; the number of pregnancies, abortions, and live births; and the total duration of any breastfeeding for each live birth. From this, the mean duration of breastfeeding per child and cumulative lifetime duration of any breastfeeding for all children were calculated for parous women. A series of physical measurements were taken using standard methods and a blood sample was collected for long-term storage. Central ethical approvals were obtained from Oxford University and the China National Center for Disease Control and Prevention (CDC). Approvals were also obtained from institutional research boards at the local CDCs in the 10 areas. All study participants provided written informed consent.

### Follow-Up for Morbidity and Mortality

Study participants were followed for cause-specific morbidity and mortality through linkage with regional disease and death registers and with the national health insurance system. Causes of death were derived from official death certificates and were, where necessary, supplemented by reviews of medical records. Data linkage with health insurance agencies was performed every 6 months in each region to retrieve all hospitalized events occurring in that period for study participants. Active follow-up was performed annually to minimize attrition. The end points for this study were incident CVD (I00–I99); CHD (I20–I25); cerebrovascular disease (henceforth, “stroke”) (I60–I69), including hemorrhagic stroke (I61) and ischemic stroke (I63); major CVDs (ie, I21–I23, I60–I61, I63–I64 [any] and I00–I20, I24–I25, I27–I59, I62, I65–I88, I95–I99 [only where fatal]); and fatal CVDs (I00–I99). Women with a self-reported history of CHD or stroke at baseline were excluded ( $n=13\ 096$ ), leaving 289 573 women for further analyses.

### Statistical Analyses

Baseline characteristics are presented as means (SDs) for continuous variables and as percentages for categorical variables. Cox proportional hazards models were used to estimate hazard ratios (HRs) and 95% CIs for the study end points associated with breastfeeding history, comparing

parous women who had never breastfed with nulliparous women and with parous women who ever breastfed. The Cox proportional hazards assumption was checked using log cumulative hazard plots and appeared to be reasonable. For comparisons involving more than 2 groups, CIs were estimated using floating absolute risks, enabling valid comparisons between any 2 groups, even if neither is the baseline group.<sup>18</sup> In analyses restricted to parous women, we obtained the HRs and CIs associated with mean duration of breastfeeding per child, comparing those who had never breastfed with those who breastfed each child for >0 to 6, 6 to 12, 12 to 18, 18 to 24, or >24 months. Among women who had ever breastfed, we estimated the HRs for each additional 6 months of breastfeeding per child. Analyses were stratified by age at risk (5-year age groups) and area of residence (10 areas) and adjusted for highest level of education attained (none, primary, secondary, and tertiary or above), household income (<5000, 5000–19 999, and ≥20 000 yuan), smoking (current, former, and never), alcohol use (weekly, occasionally, and never), physical activity, systolic blood pressure, history of hypertension, history of diabetes mellitus, and body mass index.

Several sensitivity analyses were conducted. First, in analyses among women who had ever breastfed, we additionally adjusted for other reproductive factors, ie, age at menarche, age at first birth, and total number of miscarriages, induced abortions, and stillbirths. Second, among parous women, we obtained the HRs and CIs associated with cumulative lifetime duration of breastfeeding, with additional adjustment for the number of live births. Third, we restricted the analyses on breastfeeding duration to women who had given birth to only 1 child. Subgroup analyses were conducted to obtain the HRs associated with mean duration of breastfeeding per child, separately by study region and by birth cohort. Further subgroup analyses by study region, birth cohort, highest level of attained education, body mass index, smoking status, and history of hypertension were conducted to obtain the HRs comparing parous women who had ever breastfed with parous women who had never breastfed. Among those who had ever breastfed, we conducted similar subgroup analyses to obtain the HRs associated with each additional 6 months of breastfeeding per child. Analyses were performed using SAS version 9.3 and R version 3.1.2.

## Results

Of the 289 573 women included, the mean baseline age was 51 years and 99% reported having at least 1 live birth, among whom, 97% had ever breastfed and 91% had breastfed each child for at least 6 months. The median duration of breastfeeding per child was 12 months. Women who had breastfed

for longer durations were older and were more likely to come from rural areas compared with women who had never breastfed or had breastfed for shorter durations (Table 1).

During a median of 8.1 years (quartile 1: 7.2; quartile 3: 9.1) of follow-up, 49 377 incident cases of CVD were recorded, including 16 671 cases of CHD and 23 983 of stroke (14 290 ischemic and 2998 hemorrhagic stroke).

## History of Breastfeeding and CVD Risk

There were no significant differences in the risk of CVD between nulliparous women and parous women who had never breastfed (Table 2). In contrast, parous women who had ever breastfed were at a significantly lower risk of major CVD and its main components (ie, CHD and stroke) than parous women who had never breastfed; the adjusted HRs were 0.88 (95% CI, 0.80–0.97) for major CVD, 0.91 (95% CI, 0.84–0.99) for CHD, 0.92 (95% CI, 0.85–0.99) for stroke, and 0.88 (95% CI, 0.79–0.97) for ischemic stroke. The HRs did not vary materially by study areas, birth cohorts, or other population subgroups (Figure 1).

## Breastfeeding Duration and CVD Risk

Among women who ever breastfed, there was an inverse log-linear association between the duration of breastfeeding per child and the risk of several major CVDs, except hemorrhagic stroke (Figure 2 and Tables S1 and S2). Compared with women who had never breastfed, women who had breastfed between 0 to 6 months, 6 to 12 months, 12 to 18 months, 18 to 24 months, or over 24 months had a 1%, 7%, 11%, 13%, and 18% lower risk of CHD, respectively, with each additional 6 months of breastfeeding per child associated with 4% (2–5%) lower CHD risk (*P* for trend <0.001). For stroke, the association was similar, with each additional 6 months of breastfeeding per child associated with an adjusted HR of 0.97 (95% CI, 0.96–0.98) for stroke (Table 1), with limited evidence of substantial differences across population subgroups (Figure 3). Additional adjustment for women's reproductive health factors had little impact on the results (Table S1). Moreover, analyses by total lifetime duration of breastfeeding, or restricted to women with only 1 child, yielded similar associations (Tables S3 and S4). The results were broadly similar between women in urban and rural areas and did not vary by different birth cohort (Figures S1 and S2).

## Discussion

This large prospective cohort study among nearly 300 000 women in China provides the most comprehensive assessment yet of the effects of breastfeeding on the risk of major

**Table 1.** Baseline Characteristics of Study Participants by Breastfeeding Duration

	Total	Nulliparous	Parous Women, by Breastfeeding Duration Per Child					
			Never	>0 to 6 mo	6 to 12 mo	12 to 18 mo	18 to 24 mo	>24 mo
No. (% rural)	289 573 (44)	3970 (37)	7785 (29)	17 780 (37)	131 546 (44)	65 685 (68)	42 728 (79)	20 079 (83)
Age, y	50.5 (10.3)	49.0 (11.7)	47.3 (9.2)	47.0 (9.3)	50.1 (10.2)	50.7 (10.4)	52.2 (10.8)	53.8 (9.7)
Education level, %								
Primary or below	56.8	39.6	26.2	34.2	51.8	59.1	74.2	79.5
Secondary or above	43.3	60.4	73.8	65.7	48.2	40.9	25.7	20.4
Household income, %								
Low	10.2	11.1	4.6	5.0	5.6	11.8	18.8	23.2
Middle	49.1	51.8	40.9	39.9	41.4	54.9	63.3	61.5
High	40.7	37.1	54.4	55.1	53.1	33.3	17.9	15.3
Current smoking, %	2.3	3.8	2.3	1.8	1.6	3.0	2.7	3.9
Regular alcohol use, %	2.1	3.2	2.9	2.6	1.7	2.5	1.9	2.4
Physical activity, MET h/d	17.2 (11.0–28.7)	15.0 (9.3–25.0)	16.6 (10.5–26.7)	18.0 (11.2–29.1)	17.3 (10.8–28.7)	17.0 (11.2–28.0)	18.4 (11.0–30.3)	16.5 (10.7–28.0)
Systolic blood pressure, mm Hg	129.4 (21.8)	126.3 (23.0)	123.3 (19.8)	123.1 (19.8)	128.2 (21.2)	130.2 (21.6)	133.3 (22.8)	135.2 (23.0)
Diastolic blood pressure, mm Hg	76.7 (10.8)	75.5 (11.2)	75.4 (10.7)	74.9 (10.5)	76.3 (10.7)	77.0 (10.8)	77.7 (11.2)	78.0 (11.3)
Body mass index, kg/m <sup>2</sup>	23.8 (3.4)	23.3 (3.7)	23.7 (3.4)	23.4 (3.3)	23.7 (3.4)	23.9 (3.5)	23.9 (3.5)	24.0 (3.6)
History of hypertension, %	10.2	8.8	8.6	7.7	10.7	10.4	10.0	10.4
History of diabetes mellitus, %	2.9	2.5	3.3	2.5	3.0	2.9	2.6	2.7
No. of livebirths, %								
1	35.2	0	77.3	60.6	42.8	29.8	15.0	14.4
2	31.9	0	14.4	26.1	31.7	33.9	35.3	38.1
≥3	31.5	0	8.4	13.3	25.6	36.3	49.7	47.5
Breastfeeding duration, mo								
Lifetime duration	24.0 (12.0–48.0)	NA	NA	6.0 (3.0–9.0)	20.0 (12.0–24.0)	34.0 (18.0–48.0)	48.0 (42.0–72.0)	78.0 (60.0–108.0)
Mean duration per child	12.0 (11.0–18.0)	NA	NA	5.0 (3.0–6.0)	12.0 (10.0–12.0)	15.7 (14.0–18.0)	24.0 (21.0–24.0)	30.0 (27.4–36.0)

Continued

Table 1. Continued

	Total	Nulliparous	Parous Women, by Breastfeeding Duration Per Child					
			Never	>0 to 6 mo	6 to 12 mo	12 to 18 mo	18 to 24 mo	>24 mo
Age at menarche, y	15.4 (2.0)	15.2 (2.6)	14.8 (1.9)	14.9 (1.9)	15.3 (1.9)	15.5 (1.9)	15.7 (2.0)	16.0 (2.0)
Age at first birth, y	23.4 (3.2)	NA	25.8 (3.8)	24.7 (3.4)	23.6 (3.2)	23.0 (3.0)	22.5 (2.9)	22.6 (2.8)
History of miscarriage, %	8.4	79.2	6.9	6.5	7.3	9.8	11.6	12.8
History of induced abortion, %	51.4	90.8	64.0	66.1	59.3	49.6	34.8	35.1
History of stillbirth, %	5.1	75.6	4.8	3.9	5.6	6.0	5.5	5.3

MET indicates metabolic equivalent; NA, not available. Values are percentages for categorical variables, and means and SD or median and 25th and 75th percentile for continuous variables.

CVDs in later life. Parous women who had ever breastfed were at an  $\approx 10\%$  lower risk of major CVD subtypes compared with parous women who had never breastfed. Moreover, among those who had ever breastfed, each additional 6 months of breastfeeding was associated with a further  $\approx 3\%$  to  $4\%$  lower CVD risk. These results persisted after adjustment for a range of potential confounders and were broadly consistent across major population subgroups.

Previous studies of mostly Western populations have reported conflicting results on the relationship between breastfeeding and the risk of CVD.<sup>10–14</sup> In the US Nurses' Health Study of 90 000 parous women with 2500 cases of CHD, there was a threshold in the association between breastfeeding and CHD; only women with a lifetime duration of breastfeeding of  $\geq 2$  years had a significantly lower risk of CHD than those who never breastfed.<sup>13</sup> The HUNT trial (Nord-Trøndelag Health Study)<sup>14</sup> of 20 000 Norwegian women reported that the inverse association between a history of breastfeeding and CVD mortality was modified by age. Among women younger than 65 years, there was some evidence of a U-shaped relationship between lifetime duration of breastfeeding and the risk of CVD death, with women who reported lifetime breastfeeding duration of 7 to 12 months having half the risk than those who had breastfed for  $\geq 2$  years. In contrast, the EPIC (European Prospective Investigation into Cancer and Nutrition)-CVD case-cohort study, among 15 000 women with 5000 incident cases of CHD, found an inverse association between breastfeeding and the risk of CHD at all ages.<sup>12</sup> The risk of CHD was the lowest among those who had breastfed for the longest durations, suggesting that prolonged durations of breastfeeding confer additional coronary benefits. An additional study among 200 000 parous women in the EPIC trial reported that a history of breastfeeding was associated with a 20% and 31% lower risk of mortality from CVD and CHD, respectively, but did not find any significant effect of breastfeeding on stroke.<sup>11</sup> In China, only 1 prospective study has previously examined the association between breastfeeding and CVD risk. In that study among 250 000 textile workers in Shanghai, born between 1925 and 1958,<sup>10</sup> there were no consistent associations between breastfeeding duration and the risk of fatal CHD, ischemic stroke, or hemorrhagic stroke, which may be attributable to the relatively small number of deaths included (420 CHD, 575 ischemic stroke, and 1583 hemorrhagic stroke). In contrast, the present analysis included a substantially larger number of fatal and nonfatal CVD cases and provides the largest and most comprehensive analyses to date on the relationship between breastfeeding initiation and duration and the risk of several major CVDs among urban and rural Chinese women.

The observational nature of the present study does not allow us to determine whether the inverse relationship between breastfeeding and CVD risk is causal. However,



**Table 2.** CVD End Points Associated With Breastfeeding\*

	Events, No.	Nulliparous	Parous Women <sup>†</sup>		Each Additional 6 mo Per Child <sup>‡</sup>
			Never Breastfed	Breastfed	
All CVD	49 377	1.02 (0.94–1.10)	1.00 (0.95–1.05)	0.96 (0.95–0.97)	0.98 (0.97–0.99)
Major CVD	19 227	0.99 (0.89–1.11)	1.00 (0.92–1.09)	0.88 (0.87–0.90)	0.97 (0.96–0.99)
Fatal CVD	3825	1.17 (0.92–1.48)	1.00 (0.77–1.29)	0.90 (0.87–0.94)	0.98 (0.95–1.01)
Coronary heart disease	16 671	1.01 (0.90–1.14)	1.00 (0.92–1.09)	0.91 (0.89–0.93)	0.96 (0.95–0.98)
Stroke	23 983	0.95 (0.86–1.06)	1.00 (0.93–1.08)	0.92 (0.90–0.93)	0.97 (0.96–0.98)
Hemorrhagic stroke	2998	0.92 (0.68–1.25)	1.00 (0.77–1.31)	0.84 (0.81–0.88)	0.99 (0.96–1.03)
Ischemic stroke	14 290	0.96 (0.85–1.10)	1.00 (0.91–1.09)	0.88 (0.86–0.90)	0.97 (0.95–0.98)

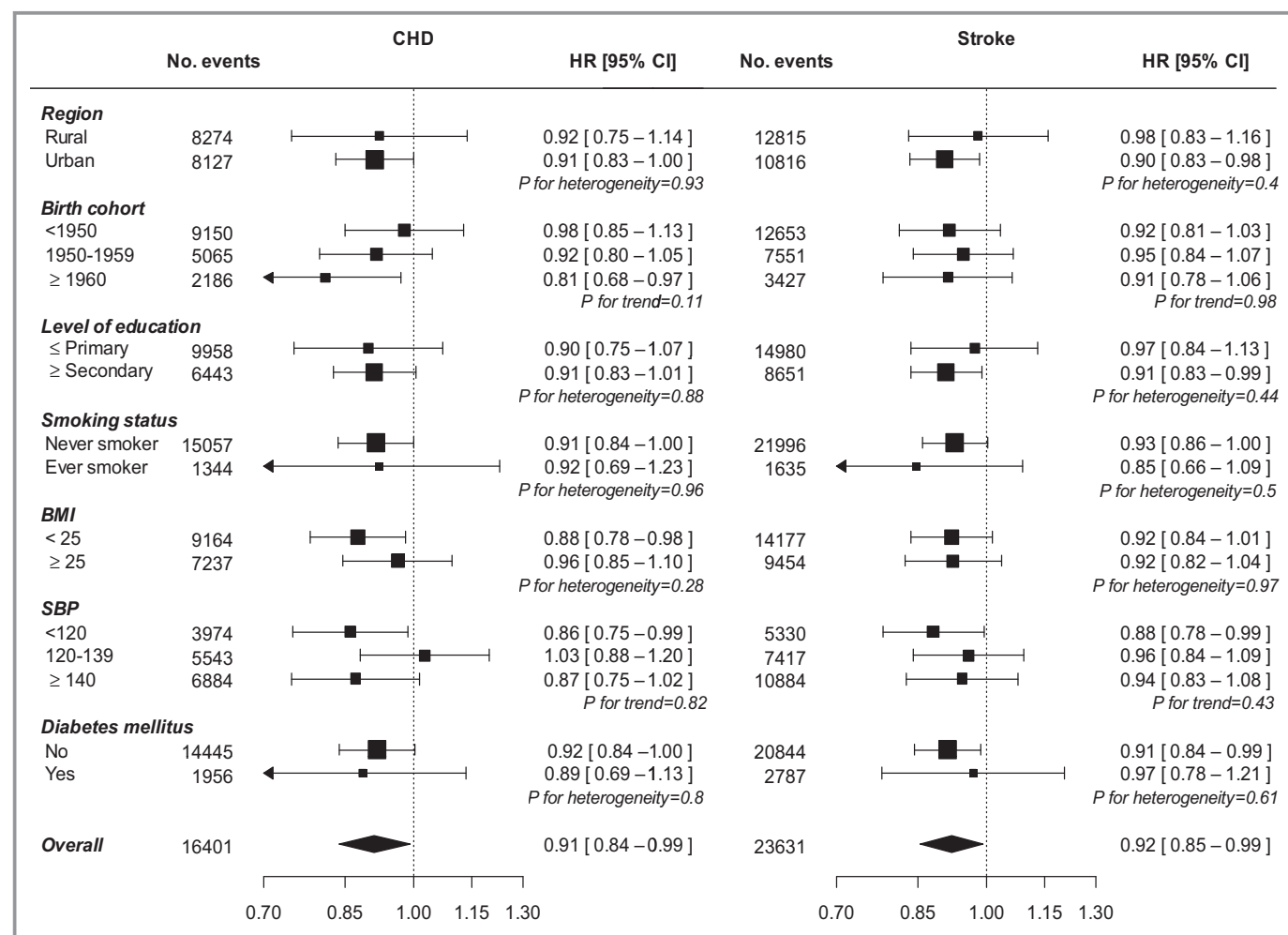
Values are adjusted hazard ratios (95% CIs).

CVD indicates cardiovascular disease.

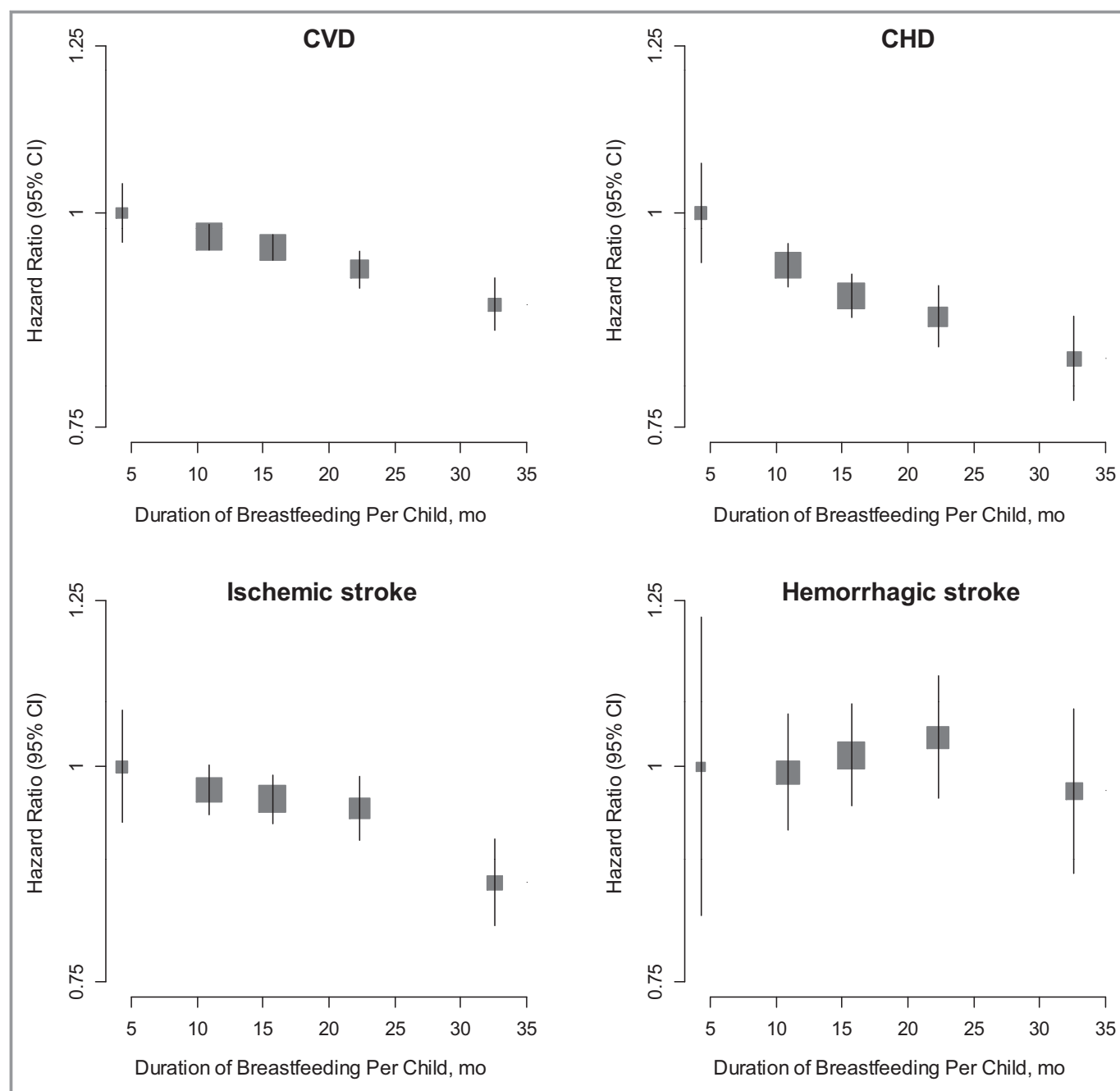
\*Analyses are stratified by age at risk and study area and adjusted for level of attained education, household income, smoking status, alcohol use, systolic blood pressure, history of hypertension, physical activity, body mass index, and history of diabetes mellitus.

<sup>†</sup>SEs were obtained using floating absolute risks.

<sup>‡</sup>Among parous women who ever breastfed only.



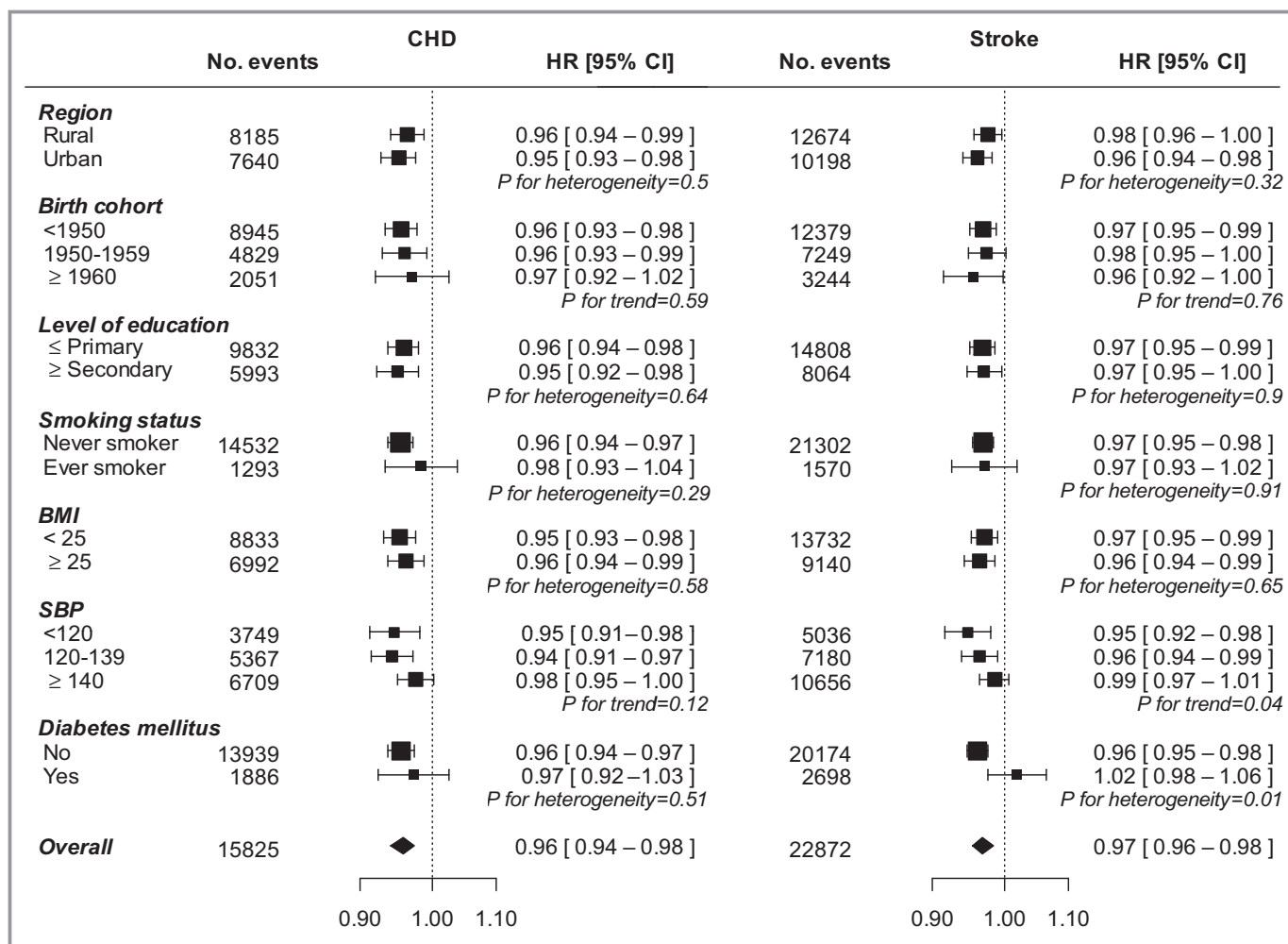
**Figure 1.** Adjusted\* hazard ratios (HRs) and 95% CIs for incident coronary heart disease (CHD) and stroke comparing parous women who ever breastfed with parous women who never breastfed, by baseline characteristics. \*Analyses are stratified by age at risk and study area, and, where appropriate, adjusted for level of attained education, household income, smoking status, alcohol use, systolic blood pressure (SBP), history of hypertension, physical activity, body mass index (BMI), and history of diabetes mellitus. Each square represents the HR. Horizontal lines indicate the corresponding 95% CIs. The diamond indicates the overall estimate and its 95% CI. Nulliparous women are excluded.



**Figure 2.** Adjusted\* hazard ratios and 95% CIs for incident cardiovascular disease (CVD), coronary heart disease (CHD), ischemic stroke, and hemorrhagic stroke associated with duration of breastfeeding per child among parous women who ever breastfed. \*Analyses are stratified by age at risk and study area, and adjusted for level of attained education, household income, smoking status, alcohol use, systolic blood pressure, history of hypertension, physical activity, body mass index, and history of diabetes mellitus. The hazard ratios are plotted on a floating absolute scale. Each square has an area inversely proportional to the SE of the log risk. Vertical lines indicate the corresponding 95% CIs.

several mechanisms have been postulated to underpin the association between breastfeeding and CVD risk in later life.<sup>19</sup> Pregnancy is characterized by major changes in the maternal metabolic system to support fetal growth and in anticipation of breastfeeding, including accumulation of visceral fat, increased insulin resistance, and higher circulating lipid levels.<sup>1,2</sup> The “reset hypothesis” posits that breastfeeding

plays a central role in mobilizing these accumulated fat stores and in resetting the maternal metabolism after birth.<sup>3</sup> The longer a woman breastfeeds, the more completely these accumulated stores may be deposited, potentially with lasting benefits for maternal cardiometabolic health. Prolonged breastfeeding has been associated with improved  $\beta$ -cell function,<sup>20</sup> more favorable cardiovascular risk factor



**Figure 3.** Adjusted\* hazard ratios (HRs) and 95% CIs for incident coronary heart disease (CHD) and stroke associated with each additional 6 months of breastfeeding per child, by baseline characteristics. Analyses are restricted to parous women who ever breastfed. \*Analyses are stratified by age at risk and study area, and, where appropriate, adjusted for level of attained education, household income, smoking status, alcohol use, systolic blood pressure, history of hypertension, physical activity, body mass index, and history of diabetes mellitus. Each square represents the HR. Horizontal lines indicate the corresponding 95% CIs. The diamond indicates the overall estimate and its 95% CI. Nulliparous women are excluded.

profiles,<sup>4,21</sup> and lower risks of incident hypertension and type 2 diabetes mellitus.<sup>8,9</sup> In contrast, studies on the potential benefits of breastfeeding on gestational weight retention and body adiposity in later life have reported mixed results.<sup>22–24</sup> In a previous analysis of this study, we did not find a relationship between breastfeeding duration and general or central adiposity.<sup>22</sup> Therefore, breastfeeding may also be a marker of health-promoting behaviors, rather than a causal determinant of improved cardiometabolic outcomes itself. Several studies in Western populations have demonstrated that women who breastfeed are generally wealthier and more likely to engage in other beneficial health behaviors than women who do not breastfeed.<sup>15,25,26</sup> However, this pattern is reversed in the present study, in which poorer women from rural areas of China were more likely to breastfeed for a

longer duration than their wealthier urban counterparts. Moreover, adjustment for various sociodemographic and lifestyle factors had little impact on our findings.

The strengths of this study include its large sample size, the prospective follow-up for the occurrence of several major CVDs, and the availability of detailed information on a range of reproductive, sociodemographic, and lifestyle-related factors. The generalizability of the study findings was enhanced by the inclusion of women from 10 diverse localities across China. The applicability of our findings to other countries and ethnicities needs to be addressed in further research. Our study did not collect data on CVD risk factors either before or during pregnancy that might determine breastfeeding as well as future CVD risk. Therefore, we cannot preclude the possibility that the findings were subject to unmeasured or residual confounding.



Moreover, our results may be liable to reverse causality as certain metabolic conditions, such as pregestational diabetes mellitus or high maternal body mass index, may affect the ability to initiate and sustain breastfeeding and may also confer a higher risk of cardiometabolic conditions.<sup>27–29</sup> Furthermore, as in most previous studies, information on breastfeeding was self-reported and solicited several years after childbirth and breastfeeding. This could have led to measurement error, which, if at random, would have underestimated the true strengths of the observed associations. Information about the duration of exclusive breastfeeding and the provision of complementary infant food was also not available, which limits interpretations of the effects seen.

Breastfeeding is one of the few beneficial health behaviors that are more common in low- and middle-income countries than in high-income countries, and within low- and middle-income countries is more common among poorer than richer women.<sup>15</sup> This socioeconomic gradient in breastfeeding is particularly relevant to China, a country where breastfeeding durations have declined dramatically over the past several decades and where breastfeeding practices continue to vary greatly across different parts of the country.<sup>16</sup> In 2008, fewer than 16% of women in urban and  $\approx 30\%$  of women in rural areas exclusively breastfed their babies throughout the World Health Organization's recommended period of 6 months.<sup>30</sup> Common reasons for prematurely ceasing exclusive breastfeeding are the perceived insufficiency of breast milk, traditions regarding the introduction of complementary food, and maternal or child illness. Intensive industry promotion of breast milk substitutes, limited duration of maternity leave, and inadequate facilities to continue to breastfeed at the workplace also contribute to shorter durations of breastfeeding, particularly in urban China.<sup>31</sup> Although there is increasing recognition of the importance of exclusive breastfeeding, genuine commitment from policy makers is needed to implement strategies in the healthcare system, communities and families, and the work environment that promote and support every woman to breastfeed.<sup>32</sup> If effective and sustained, such efforts are likely to confer major benefits to the health of children and women, along with substantial cost savings.<sup>15,32</sup>

## Conclusions

This large prospective study showed that breastfeeding is associated with an  $\approx 10\%$  lower risk of several major CVDs in later life among Chinese women and the magnitude of the inverse association was stronger among those with a longer duration of breastfeeding. If causal, these findings suggest that interventions to increase the likelihood and duration of breastfeeding could have persistent benefits to maternal cardiovascular health.

## Appendix

### Members of the China Kadoorie Biobank Collaborative Group

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## Disclosures

None.

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# Supplemental Material

**Table S1. Hazard ratios (95% confidence intervals) for cardiovascular disease associated with duration of breastfeeding among parous women who ever breastfed across different levels of adjustment**

	N events	Duration of breastfeeding per child, months				
		>0-6 months	6-12 months	12-18 months	18-24 months	>24 months
Model 1						
All cardiovascular disease	47301	1.00 (0.96; 1.04)	0.98 (0.96; 1.00)	0.97 (0.96; 0.99)	0.95 (0.92; 0.97)	0.91 (0.88; 0.94)
Major cardiovascular disease	18348	1.00 (0.94; 1.07)	1.00 (0.97; 1.03)	1.00 (0.97; 1.02)	0.99 (0.96; 1.03)	0.94 (0.89; 0.98)
Fatal cardiovascular disease	3695	1.00 (0.84; 1.20)	0.96 (0.89; 1.03)	0.99 (0.94; 1.06)	0.95 (0.89; 1.02)	0.99 (0.91; 1.09)
Coronary heart disease	15825	1.00 (0.94; 1.07)	0.94 (0.91; 0.97)	0.90 (0.88; 0.93)	0.87 (0.83; 0.91)	0.82 (0.78; 0.87)
Stroke	22872	1.00 (0.94; 1.06)	0.98 (0.96; 1.01)	0.98 (0.95; 1.00)	0.97 (0.93; 1.00)	0.88 (0.84; 0.92)
Haemorrhagic stroke	2900	1.00 (0.82; 1.22)	1.04 (0.96; 1.12)	1.07 (1.00; 1.15)	1.11 (1.02; 1.20)	1.06 (0.95; 1.18)
Ischaemic stroke	13580	1.00 (0.93; 1.08)	1.01 (0.97; 1.04)	1.00 (0.97; 1.03)	0.99 (0.95; 1.04)	0.91 (0.86; 0.96)
Model 2						
All cardiovascular disease	47301	1.00 (0.96; 1.04)	0.97 (0.96; 0.99)	0.97 (0.96; 0.99)	0.94 (0.92; 0.96)	0.90 (0.87; 0.93)
Major cardiovascular disease	18348	1.00 (0.94; 1.07)	0.98 (0.95; 1.01)	0.97 (0.94; 1.00)	0.96 (0.93; 0.99)	0.90 (0.86; 0.95)
Fatal cardiovascular disease	3695	1.00 (0.84; 1.20)	0.89 (0.83; 0.96)	0.92 (0.87; 0.98)	0.87 (0.81; 0.94)	0.89 (0.82; 0.98)
Coronary heart disease	15825	1.00 (0.94; 1.07)	0.94 (0.91; 0.97)	0.92 (0.89; 0.94)	0.89 (0.85; 0.92)	0.84 (0.79; 0.89)
Stroke	22872	1.00 (0.94; 1.06)	0.97 (0.95; 1.00)	0.97 (0.95; 0.99)	0.96 (0.93; 0.99)	0.87 (0.83; 0.92)
Haemorrhagic stroke	2900	1.00 (0.82; 1.22)	1.02 (0.94; 1.10)	1.05 (0.98; 1.12)	1.08 (0.99; 1.17)	1.02 (0.91; 1.14)
Ischaemic stroke	13580	1.00 (0.93; 1.08)	0.99 (0.96; 1.03)	0.99 (0.96; 1.02)	0.98 (0.94; 1.02)	0.89 (0.84; 0.94)

<b>Model 3</b>						
All cardiovascular disease	47301	1.00 (0.96; 1.04)	0.97 (0.95; 0.99)	0.95 (0.94; 0.97)	0.93 (0.90; 0.95)	0.88 (0.85; 0.92)
Major cardiovascular disease	18348	1.00 (0.94; 1.07)	0.96 (0.93; 0.99)	0.95 (0.92; 0.97)	0.94 (0.91; 0.97)	0.88 (0.84; 0.92)
Fatal cardiovascular disease	3695	1.00 (0.84; 1.20)	0.88 (0.82; 0.95)	0.91 (0.86; 0.97)	0.86 (0.80; 0.92)	0.88 (0.80; 0.96)
Coronary heart disease	15825	1.00 (0.94; 1.07)	0.93 (0.90; 0.96)	0.89 (0.87; 0.92)	0.87 (0.84; 0.91)	0.82 (0.78; 0.87)
Stroke	22872	1.00 (0.94; 1.06)	0.96 (0.94; 0.98)	0.95 (0.92; 0.97)	0.94 (0.91; 0.97)	0.85 (0.81; 0.89)
Haemorrhagic stroke	2900	1.00 (0.82; 1.22)	0.99 (0.92; 1.07)	1.02 (0.95; 1.09)	1.04 (0.96; 1.13)	0.97 (0.87; 1.08)
Ischaemic stroke	13580	1.00 (0.93; 1.08)	0.97 (0.94; 1.00)	0.96 (0.93; 0.99)	0.95 (0.91; 0.99)	0.86 (0.81; 0.91)
<b>Model 4</b>						
All cardiovascular disease	47301	1.00 (0.96; 1.04)	0.96 (0.95; 0.98)	0.95 (0.93; 0.96)	0.92 (0.89; 0.94)	0.88 (0.85; 0.91)
Major cardiovascular disease	18348	1.00 (0.93; 1.07)	0.96 (0.93; 0.99)	0.94 (0.92; 0.97)	0.94 (0.90; 0.97)	0.88 (0.83; 0.92)
Fatal cardiovascular disease	3695	1.00 (0.84; 1.20)	0.88 (0.82; 0.94)	0.90 (0.85; 0.96)	0.85 (0.80; 0.91)	0.87 (0.79; 0.95)
Coronary heart disease	15825	1.00 (0.94; 1.07)	0.93 (0.90; 0.95)	0.89 (0.86; 0.91)	0.86 (0.83; 0.90)	0.81 (0.77; 0.86)
Stroke	22872	1.00 (0.94; 1.06)	0.96 (0.93; 0.98)	0.94 (0.92; 0.97)	0.93 (0.90; 0.96)	0.84 (0.80; 0.88)
Haemorrhagic stroke	2900	1.00 (0.82; 1.22)	0.99 (0.91; 1.07)	1.01 (0.94; 1.08)	1.03 (0.95; 1.12)	0.96 (0.86; 1.07)
Ischaemic stroke	13580	1.00 (0.93; 1.08)	0.97 (0.94; 1.00)	0.96 (0.93; 0.99)	0.94 (0.91; 0.99)	0.86 (0.81; 0.91)

Model 1: stratified by age at risk and study area; Model 2: Model 1 + adjustment for level of attained education, household income, smoking status, and alcohol use; Model 3: Model 2 + adjustment for systolic blood pressure, history of hypertension, physical activity, body mass index, and history of diabetes. Model 4 + adjustment for age at menarche, age at first birth, and total number of miscarriages, induced abortions, and stillbirths.



**Table S2. Hazard ratios (95% confidence intervals) for cardiovascular disease associated with duration of breastfeeding per child among parous women across different levels of adjustment, using those who never breastfed as the reference group**

	Duration of breastfeeding per child, months						
	N events	Never	>0-6 months	6-12 months	12-18 months	18-24 months	>24 months
<b>Model 1</b>							
All cardiovascular disease	48692	1.00 (0.95; 1.06)	0.99 (0.95; 1.03)	0.97 (0.95; 0.99)	0.96 (0.95; 0.98)	0.93 (0.91; 0.96)	0.90 (0.87; 0.93)
Major cardiovascular disease	18923	1.00 (0.92; 1.09)	0.90 (0.85; 0.97)	0.91 (0.88; 0.93)	0.90 (0.88; 0.93)	0.90 (0.86; 0.93)	0.85 (0.81; 0.89)
Fatal cardiovascular disease	3755	1.00 (0.77; 1.29)	1.00 (0.84; 1.20)	0.96 (0.90; 1.03)	1.00 (0.94; 1.06)	0.95 (0.89; 1.02)	1.00 (0.91; 1.09)
Coronary heart disease	16401	1.00 (0.92; 1.09)	0.97 (0.91; 1.04)	0.91 (0.89; 0.94)	0.88 (0.85; 0.91)	0.85 (0.81; 0.88)	0.80 (0.76; 0.85)
Stroke	23631	1.00 (0.93; 1.08)	0.95 (0.90; 1.00)	0.93 (0.91; 0.96)	0.93 (0.90; 0.95)	0.92 (0.89; 0.95)	0.84 (0.80; 0.88)
Haemorrhagic stroke	2956	1.00 (0.77; 1.31)	0.84 (0.69; 1.02)	0.87 (0.81; 0.94)	0.90 (0.84; 0.97)	0.93 (0.86; 1.01)	0.89 (0.79; 0.99)
Ischaemic stroke	14066	1.00 (0.91; 1.10)	0.89 (0.83; 0.96)	0.90 (0.87; 0.93)	0.89 (0.86; 0.92)	0.88 (0.85; 0.92)	0.81 (0.76; 0.85)
<b>Model 2</b>							
All cardiovascular disease	48692	1.00 (0.95; 1.06)	0.99 (0.95; 1.03)	0.97 (0.95; 0.98)	0.96 (0.95; 0.98)	0.93 (0.91; 0.96)	0.89 (0.86; 0.93)
Major cardiovascular disease	18923	1.00 (0.92; 1.09)	0.91 (0.85; 0.97)	0.90 (0.87; 0.92)	0.89 (0.86; 0.91)	0.88 (0.85; 0.91)	0.83 (0.79; 0.87)
Fatal cardiovascular disease	3755	1.00 (0.77; 1.29)	1.00 (0.84; 1.20)	0.89 (0.83; 0.95)	0.92 (0.86; 0.97)	0.87 (0.81; 0.93)	0.88 (0.80; 0.96)
Coronary heart disease	16401	1.00 (0.92; 1.09)	0.98 (0.92; 1.05)	0.92 (0.90; 0.95)	0.90 (0.87; 0.93)	0.87 (0.84; 0.91)	0.82 (0.78; 0.87)
Stroke	23631	1.00 (0.93; 1.08)	0.97 (0.91; 1.02)	0.93 (0.90; 0.95)	0.92 (0.89; 0.94)	0.91 (0.88; 0.94)	0.82 (0.78; 0.86)
Haemorrhagic stroke	2956	1.00 (0.77; 1.31)	0.84 (0.69; 1.02)	0.85 (0.79; 0.92)	0.87 (0.81; 0.93)	0.89 (0.82; 0.97)	0.84 (0.76; 0.94)
Ischaemic stroke	14066	1.00 (0.91; 1.10)	0.90 (0.83; 0.96)	0.89 (0.86; 0.92)	0.88 (0.85; 0.91)	0.87 (0.83; 0.91)	0.79 (0.75; 0.84)
<b>Model 3</b>							
All cardiovascular disease	48692	1.00 (0.95; 1.06)	1.00 (0.96; 1.04)	0.97 (0.96; 0.99)	0.96 (0.94; 0.97)	0.93 (0.91; 0.95)	0.89 (0.86; 0.92)
Major cardiovascular disease	18923	1.00 (0.92; 1.09)	0.93 (0.87; 0.99)	0.89 (0.87; 0.92)	0.88 (0.85; 0.90)	0.87 (0.84; 0.90)	0.81 (0.78; 0.85)
Fatal cardiovascular disease	3755	1.00 (0.77; 1.29)	1.01 (0.84; 1.20)	0.90 (0.84; 0.96)	0.91 (0.85; 0.96)	0.85 (0.80; 0.92)	0.89 (0.81; 0.97)
Coronary heart disease	16401	1.00 (0.92; 1.09)	0.99 (0.93; 1.06)	0.93 (0.90; 0.96)	0.89 (0.86; 0.92)	0.87 (0.83; 0.90)	0.82 (0.77; 0.87)
Stroke	23631	1.00 (0.93; 1.08)	0.95 (0.90; 1.01)	0.93 (0.90; 0.95)	0.92 (0.90; 0.94)	0.91 (0.88; 0.94)	0.83 (0.79; 0.87)
Haemorrhagic stroke	2956	1.00 (0.76; 1.31)	0.84 (0.69; 1.03)	0.84 (0.78; 0.90)	0.86 (0.80; 0.92)	0.88 (0.81; 0.95)	0.82 (0.73; 0.91)
Ischaemic stroke	14066	1.00 (0.91; 1.10)	0.92 (0.85; 0.99)	0.89 (0.86; 0.92)	0.88 (0.85; 0.91)	0.87 (0.83; 0.90)	0.78 (0.74; 0.83)

Adjustment as in Table S1

**Table S3. Adjusted\* hazard ratios (95% confidence intervals) for cardiovascular disease associated with total lifetime duration of breastfeeding among parous women**

	N events	Lifetime duration of breastfeeding, months					
		Never	>0-12 months	12-24 months	24-36 months	36-48 months	>48 months
All cardiovascular disease	48692	1.00 (0.95; 1.06)	0.96 (0.93; 0.99)	0.97 (0.95; 0.99)	0.96 (0.94; 0.98)	0.92 (0.89; 0.94)	0.91 (0.88; 0.93)
Major cardiovascular disease	18923	1.00 (0.92; 1.09)	0.90 (0.86; 0.95)	0.91 (0.87; 0.94)	0.87 (0.84; 0.90)	0.80 (0.77; 0.83)	0.82 (0.78; 0.85)
Fatal cardiovascular disease	3755	1.00 (0.77; 1.29)	0.88 (0.74; 1.04)	0.98 (0.87; 1.09)	0.93 (0.85; 1.02)	0.81 (0.74; 0.89)	0.86 (0.79; 0.92)
Coronary heart disease	16401	1.00 (0.92; 1.09)	0.93 (0.89; 0.99)	0.92 (0.89; 0.96)	0.86 (0.83; 0.89)	0.86 (0.82; 0.90)	0.82 (0.79; 0.86)
Stroke	23631	1.00 (0.93; 1.08)	0.93 (0.89; 0.97)	0.93 (0.90; 0.96)	0.91 (0.88; 0.94)	0.86 (0.82; 0.89)	0.85 (0.82; 0.89)
Haemorrhagic stroke	2956	1.00 (0.76; 1.31)	0.84 (0.72; 0.99)	0.87 (0.78; 0.97)	0.81 (0.74; 0.89)	0.78 (0.70; 0.86)	0.84 (0.77; 0.93)
Ischaemic stroke	14066	1.00 (0.91; 1.10)	0.91 (0.85; 0.96)	0.89 (0.85; 0.93)	0.86 (0.83; 0.90)	0.80 (0.76; 0.84)	0.81 (0.77; 0.85)

\*Analyses are stratified by age at risk and study area, and adjusted for level of attained education, household income, smoking status, alcohol use, systolic blood pressure, history of hypertension, physical activity, body mass index, history of diabetes, and number of livebirths.

**Table S4. Adjusted\* hazard ratios (95% confidence intervals) for cardiovascular disease associated with duration of breastfeeding among women with one livebirth, using those who never breastfed as the reference group**

	N events	Lifetime duration of breastfeeding, months						
		Ever vs. never	Never	>0-6 months	6-12 months	12-18 months	18-24 months	>24 months
All cardiovascular disease	10760	0.95 (0.88; 1.01)	1.00 (0.95; 1.06)	0.96 (0.93; 0.99)	0.97 (0.95; 0.99)	0.96 (0.94; 0.98)	0.92 (0.89; 0.94)	0.91 (0.88; 0.93)
Major cardiovascular disease	3274	0.89 (0.80; 1.00)	1.00 (0.92; 1.09)	0.90 (0.86; 0.95)	0.91 (0.87; 0.94)	0.87 (0.84; 0.90)	0.80 (0.77; 0.83)	0.82 (0.78; 0.85)
Fatal cardiovascular disease	303	0.94 (0.62; 1.45)	1.00 (0.77; 1.29)	0.88 (0.74; 1.04)	0.98 (0.87; 1.09)	0.93 (0.85; 1.02)	0.81 (0.74; 0.89)	0.86 (0.79; 0.92)
Coronary heart disease	3597	0.89 (0.80; 0.99)	1.00 (0.92; 1.09)	0.93 (0.89; 0.99)	0.92 (0.89; 0.96)	0.86 (0.83; 0.89)	0.86 (0.82; 0.90)	0.82 (0.79; 0.86)
Stroke	4676	0.93 (0.84; 1.02)	1.00 (0.93; 1.08)	0.93 (0.89; 0.97)	0.93 (0.90; 0.96)	0.91 (0.88; 0.94)	0.86 (0.82; 0.89)	0.85 (0.82; 0.89)
Haemorrhagic stroke	350	0.92 (0.62; 1.37)	1.00 (0.76; 1.31)	0.84 (0.72; 0.99)	0.87 (0.78; 0.97)	0.81 (0.74; 0.89)	0.78 (0.70; 0.86)	0.84 (0.77; 0.93)
Ischaemic stroke	2689	0.90 (0.80; 1.02)	1.00 (0.91; 1.10)	0.91 (0.85; 0.96)	0.89 (0.85; 0.93)	0.86 (0.83; 0.90)	0.80 (0.76; 0.84)	0.81 (0.77; 0.85)

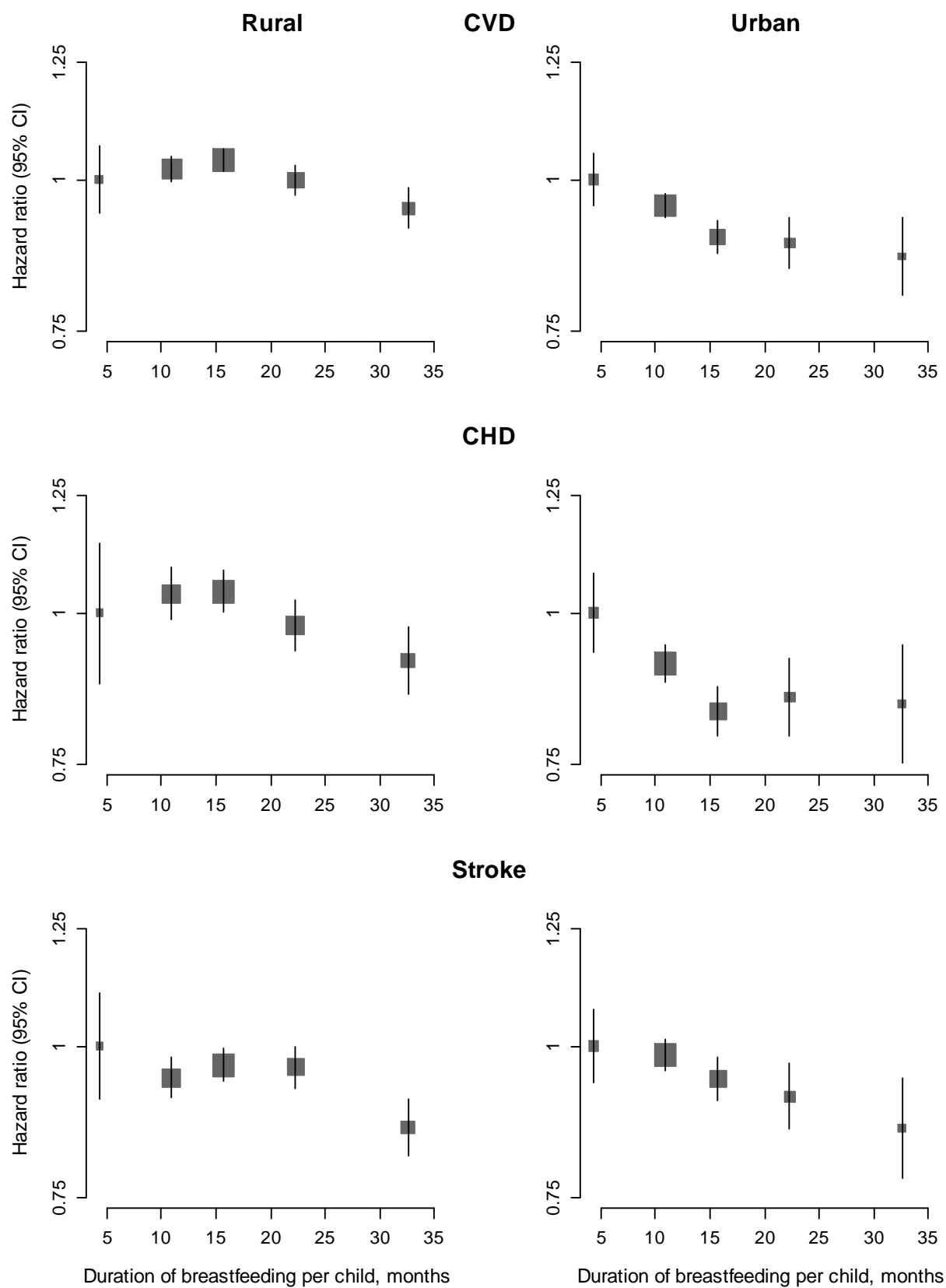
\*Analyses are stratified by age at risk and study area, and adjusted for level of attained education, household income, smoking status, alcohol use, systolic blood pressure, history of hypertension, physical activity, body mass index, and history of diabetes.

## Supplemental Figure Legends:

**Figure S1: Adjusted\* hazard ratios and 95% CIs for incident cardiovascular disease, coronary heart disease, and stroke associated with duration of breastfeeding per child among parous women who ever breastfed, by region.**\*Analyses are stratified by age at risk and study area, and adjusted for level of attained education, household income, smoking status, alcohol use, systolic blood pressure, history of hypertension, physical activity, body mass index, and history of diabetes. The hazard ratios (HRs) are plotted on a floating absolute scale. Each square has an area inversely proportional to the standard error of the log risk. Vertical lines indicate the corresponding 95% confidence intervals (CIs).

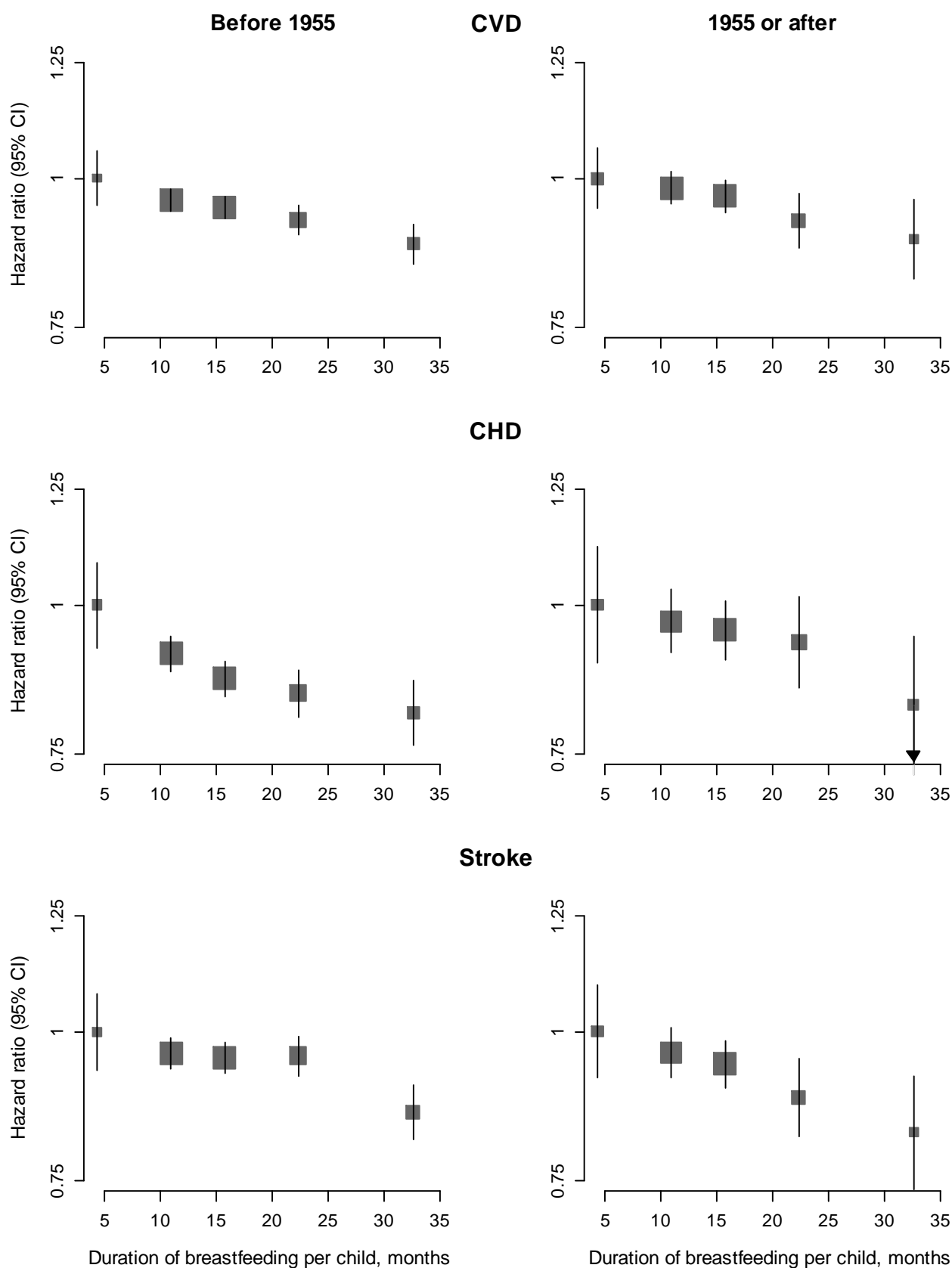
**Figure S2. Adjusted\* hazard ratios and 95% CIs for incident cardiovascular disease, coronary heart disease, and stroke associated with duration of breastfeeding per child among parous women who ever breastfed, by birth cohort.**\*Analyses are stratified by age at risk and study area, and adjusted for level of attained education, household income, smoking status, alcohol use, systolic blood pressure, history of hypertension, physical activity, body mass index, and history of diabetes. The hazard ratios (HRs) are plotted on a floating absolute scale. Each square has an area inversely proportional to the standard error of the log risk. Vertical lines indicate the corresponding 95% confidence intervals (CIs).

Figure S1. Adjusted\* hazard ratios for incident CVD, CHD, and stroke associated with duration of breastfeeding per child among parous women who ever breastfed, by region





**Figure S2. Adjusted\* hazard ratios for incident CVD, CHD and stroke associated with duration of breastfeeding per child among parous women who ever breastfed, by birth cohort**



## **Breastfeeding and the Risk of Maternal Cardiovascular Disease: A Prospective Study of 300 000 Chinese Women**

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