Maternal exposure to Wenchuan earthquake and prolonged risk of offspring birth outcomes: a natural experiment study

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Abstract

Background: The prolonged effects of disasters on reproductive outcomes among the survivors are less studied, and the findings are inconsistent. We examined the associations of maternal exposure to the 2008 Wenchuan earthquake years before conception with adverse birth outcomes.

Methods: We included 73,493 women who delivered in 96 hospitals in 24 provinces and autonomous regions from the 2015/16 China Labor and Delivery Survey. We weighted the multivariable logistic models based on the combination of coarsened exact matching (CEM) weight and survey weight, and performed sex-stratified analysis to test whether associations of maternal earthquake exposure with adverse birth outcomes (Stillbirth, preterm birth [PTB], low birthweight [LBW], and small for gestational age [SGA]) varied by sex.

Results: The bivariate models showed that the weighted incidence of each adverse birth outcome was higher in exposed group than unexposed group: stillbirth (2.00% vs. 1.33%), PTB (14.14% vs. 7.32%), LBW (10.82% vs. 5.76%), and SGA (11.32% vs. 9.52%). The multivariable models showed maternal earthquake exposure was only associated significantly with a higher risk of PTB in offspring among all births (adjusted risk ratio [aRR] (95%CI): 1.25(1.06–1.48), \( P = 0.010 \)). The sex-stratified analysis showed the association was significant among male births (aRR (95%CI): 1.40(1.12–1.75), \( P = 0.002 \)), but insignificant among female births. The sensitivity analysis reported similar findings.

Conclusions: The 2008 Wenchuan earthquake exposure has a long-term effect on PTB. Maternal acute exposure to disasters could be a major monitor for long-term reproductive outcomes. More attention should be paid to the underlying reasons for disaster-related adverse birth outcomes.

Keywords: Wenchuan earthquake, Adverse birth outcomes, Preterm birth

Background

Adverse birth outcomes, such as low birthweight (LBW), preterm birth (PTB) and small for gestational age (SGA), are associated with numerous risk factors, including not only maternal characteristics [1, 2] and environmental contaminants [3], but also external stressors [4]. There is growing evidence that maternal exposure to natural or manmade disasters, including major earthquakes, ice storms and terrorist attacks during pregnancy could impact the birth outcomes intensively [5–9]. However, most of the existing studies to date have focused on the women who were pregnant during or immediately after a disaster and leaving behind the exposed women who get pregnant later [10]. Hence, the long-term negative effects of maternal acute exposure to disasters on reproductive outcomes are less studied.
Moreover, the few existing studies of prolonged effects of disasters on reproductive outcomes among the survivors reported mixed results, with one showing negative effects among women exposed to the 2001 World Trade Center disaster in United States [10] and another showing no effects among the women experienced the 2011 Fukushima Daiichi nuclear disaster in Japan [11]. Thus, the long-term effects of disasters remain unclear and are worthy of further investigation.

The 8.0-magnitude Wenchuan earthquake hit Wenchuan, Sichuan province on May 12, 2008, leading to 69,197 deaths, 374,176 injuries and 18,222 missing persons [12]. The disaster was one of the biggest and deadliest earthquakes in Chinese history, it affected more than 15 million people and left 4.8 million people homeless [12]. Existing studies assessed the long-term effects of the earthquake on mental health [12] and puberty timing [13] among the adolescent survivors. No studies to date, however, have evaluated the long-term effects of the 2008 Wenchuan earthquake on birth outcomes, although a hospital-based study showed increased ratios of LBW and PTB within the first 12 months post-disaster in 2 counties affected by Wenchuan earthquake [8].

To bridge the gap, we evaluated the data from a national hospital-based cross-sectional study to investigate the long-term effects of the Wenchuan earthquake on birth outcomes. We hypothesized that the women exposed to the devastating earthquake and became pregnant years later would be associated with elevated risks of adverse birth outcomes in offspring. Given that male births are more vulnerable to maternal stress [14], we, therefore, hypothesized that there are sex differences in these associations.

Methods

Study design and participants

Data in the present study were from the China Labor and Delivery Survey (CLDS), a multi-center, hospital-based cross-sectional survey throughout China to estimate the national labour and delivery information of new births between March 1, 2015, and December 31, 2016. Using a stratified multistage cluster sampling design, the CLDS selected 112 hospitals with at least 1000 childbirths per year from 25 provinces and autonomous regions across China mainland. Details on the sampling strategy of CLDS have been described elsewhere [15]. The CLDS established a data coordination center (DCC) to retrieve, review and extract the data of new births and their mothers from hospital records in the selected hospitals. Within each calendar year, the DCC randomly selected 6 weeks for hospitals with at least 6000 childbirths per year, or 10 weeks for hospitals with less than 6000 childbirths per year for data collection. Within each selected week, all births born at 24 weeks gestation or more or weighing at least 500 g were eligible for inclusion.

Although the CLDS was a cross-sectional observational study, the Wenchuan earthquake generated a natural experimental setting to investigate the long-term effects of the earthquake for birth outcomes among the female survivors who became pregnancy years following the devastating earthquake [13, 16]. In this natural experiment, the earthquake event that divided the participants into exposed and unexposed groups was unexpected and not under the control [17]. In the current analysis, 96 hospitals with 70% or higher response rates in 24 provinces and autonomous regions, including Sichuan province, were involved.

Exposure

In this natural experiment, individual-level maternal exposure to the 2008 Sichuan earthquake was unmeasured in the original design and is difficult to define, as noted in the previous review [18], indicator of the earthquake exposure was obtained from the location of the delivery hospitals. We labeled the participants who delivered in 5 hospitals of Sichuan province as a natural indicator of earthquake exposure, and their counterparts in 91 hospitals of other provinces as an unexposed group.

Outcomes

The main outcomes of interest were the outcome of delivery, gestational age, birthweight, and birthweight for gestational age. The corresponding adverse birth outcomes included stillbirth, PTB, LBW, and SGA.

We defined stillbirth as the death of a baby before or during delivery after 23 weeks gestation or weighing at least 500 g at birth. We divided gestational age at birth as preterm (<37 weeks), term (37–41 weeks) and post-term (≥42 weeks) birth. We classified birthweight into low (<2500 g), normal (2500–3999 g) and high (≥4000 g) groups. We categorized sex-specific birth weight-for-gestational age as small (<10th percentile), appropriate (10th–90th percentile) and large (>90th percentile) categories.

Covariates

We divided maternal age into young (<20 years), reference (20–34 years) and advanced (≥35 years) class. We categorized maternal education attainment as low (junior school or below), middle (high school, technical
school, or junior college), and high (bachelor degree or above) group. We labeled the women as residents if they held the hukou in the province where they delivered, or migrants if not. According to the WHO classification of body mass index (BMI, defined as weight in kilograms divided by the square of height in meters), we categorized maternal pre-pregnancy BMI as underweight (< 18.5 kg/m²), healthy weight (18.5–24.9 kg/m²), and overweight or obesity (≥25 kg/m²) [19].

Maternal medical conditions in the present study included maternal diseases (hypertension, gestational hypertension, diabetes, gestational diabetes, cardiac disease, renal disease, autoimmune disease, thyroid disease, gestational thyroid disease, gestational anemia, asthma and gestational asthma), history of PTB, method of conception, parity (primipara or multipara), multiple pregnancy or singleton pregnancy, mode of delivery and sex of newborns.

Statistical analysis
Statistical analysis was performed from January 5, 2020, to March 12, 2020. Statistical analyses were performed using Stata/SE 15.1 (StataCorp LLC, College Station, TX, United States). To obtain a national estimation of labour and delivery information, survey weight for each newborn included in CLDS was calculated using the delivery data from 2016 China Statistical Yearbook in the previous study [15]. The `svy` and `subpop` option were used to account for complex survey design.

We first examined distributions of the exposure indicator, outcome variables, and covariates, and investigated the associations of exposure indicator with outcome variables and covariates using \( \chi^2 \) tests, and found the distribution of a few covariates was not balanced between the two groups (Table 1). To account for pregnant women who delivery in earthquake-affected areas being inherently different from those who do not, we matched the data first and weighted the analyses based on coarsened exact matching, the global imbalance was measured by overall L1 statistic, ranging from 0 to 1, with larger value indicating larger imbalance between the two groups [20]. Candidate variables for the coarsened exact matching (CEM)–weighted model included hospital level and the covariates measured in the study (Table 1). We obtained new weight using multiplying the CEM weights by the survey weight, then declared the data using `svyset` command with the new weight. We reported CEM–weighted adjusted risk ratio (aRR) and 95% confidence interval (CI) using `adjrr` command [21] after weighted logistic regression.

In sensitive analyses, we restricted our analysis to singleton births, given that multiple pregnancies carry significantly higher risks of birth outcomes, including preterm birth [22] and stillbirth [23]. We also performed sex-stratified analysis as we hypothesized sex differences in stress responses [14]. To minimize the effect of migration on earthquake exposure, we further excluded the migrants and assessed the associations among the local pregnant women only.

Results
We drew a total of 73,493 hospital records in our primary analysis, including 34,170 female births (47.30%), 1569 multiple births (1.78%) and 10,150 pregnant migrant women (12.85%), more details of the delivery information can be found in Table 1. After matching with CEM method, there are 37,059 matched cases (33,812 in unexposed group, 3247 in exposed group). The overall L1 statistic reduced from 0.55616578 before matching to 1.733e-14 after matching, suggesting that our matching significantly improved the imbalance in covariates between the two groups.

The weighted incidences of adverse birth outcomes were 1.35% for stillbirth, 7.55% for PTB, 5.93% for LBW, and 9.58% for SGA in our sample (Table 2). As showed in Table 2, the incidence of each adverse birth outcome was higher in exposed group: stillbirth (2.00% vs 1.33%, \( P = 0.018 \)), PTB (14.14% vs 7.32%, \( P < 0.001 \)), LBW (10.82% vs 5.76%, \( P < 0.001 \)), and SGA (11.32% vs 9.52%, \( P = 0.008 \)).

Maternal exposure to Wenchuan Earthquake years before pregnancy was associated with significantly higher of risk of PTB in offspring (CEM–weighted aRR, 1.25; 95% CI, 1.06–1.48; \( P = 0.010 \)) among all births, and similar results were found in other subpopulation, including singleton births, all livebirths, and singleton livebirths (Tables 3 and 4). However, the associations of earthquake exposure with stillbirth, LBW, and SGA were all non-significant in four samples (Table 3 and Table 4). Associations of maternal earthquake exposure with PTB varied by sex in all birth and singleton births, with significant associations observed in male births but not in female births (Table 3).

The analyses that focused on the local residents yielded similar results. Although maternal earthquake exposure was not associated with most of the adverse birth outcomes, it related to increased risk of PTB, and the associations were also sex differential (eTable 1 in the supplement).

Discussion
This, to our best knowledge, is the first study to investigate the long-effect of the Wenchuan earthquake on birth outcomes among the survivors that became pregnancy years later. The present study retrospectively compared the risks of 5 birth outcomes, including stillbirth, LBW, and SGA, from 2015 to 2016 at the national level between the Wenchuan earthquake-affected and
Table 1 Descriptive Statistics of potential covariables among Wenchuan exposure and non-exposure group

| Characteristic            | No. (Weighted %) | Exposure | Non-exposure | $\chi^2$ | P value |
|---------------------------|------------------|----------|--------------|---------|---------|
| **Maternal age**          |                  |          |              |         |         |
| < 20 years                | 1093 (2.39)      | 93 (2.70)| 1000 (2.38)  | 1.94    | 0.413   |
| 20–34 years               | 63,948 (86.41)   | 2933 (86.70)| 61,015 (86.40)| 9.83 | < 0.001 |
| > 34 years                | 8364 (11.20)     | 386 (10.60)| 7978 (11.22) |         |         |
| **Maternal education**    |                  |          |              |         |         |
| Low                       | 16,763 (36.38)   | 911 (35.34)| 15,852 (36.41)| 983  | < 0.001 |
| Middle                    | 29,302 (39.70)   | 1144 (44.18)| 28,158 (39.56)| 6.89 | 0.001   |
| High                      | 19,846 (23.93)   | 529 (20.48)| 19,317 (24.03)|         |         |
| **Hukou**                 |                  |          |              |         |         |
| Local                     | 62,833 (87.15)   | 3303 (96.73)| 59,530 (86.81)| 230.02| < 0.001 |
| Migrant                   | 10,150 (12.85)   | 101 (3.27)| 10,049 (13.19)|         |         |
| **Pre-pregnant BMI**      |                  |          |              |         |         |
| Underweight               | 6769 (12.95)     | 266 (15.18)| 6503 (12.90)  | 6.89    | 0.001   |
| Healthy weight            | 39,463 (72.49)   | 1208 (73.37)| 38,255 (72.47)|         |         |
| Overweight/Obese          | 8105 (14.56)     | 184 (11.45)| 7921 (14.64)  |         |         |
| **History of preterm**    |                  |          |              |         |         |
| Yes                       | 899 (1.47)       | 34 (1.04)| 865 (1.48)   | 3.01    | 0.083   |
| No                        | 72,388 (98.53)   | 3370 (98.96)| 69,018 (98.52)|         |         |
| **Artificial Conception** |                  |          |              |         |         |
| Yes                       | 4690 (6.47)      | 128 (3.33)| 4562 (6.58)  | 50.10   | < 0.001 |
| No                        | 68,491 (93.53)   | 3287 (96.67)| 65,204 (93.42)|         |         |
| **Parity**                |                  |          |              |         |         |
| Primiparous               | 41,207 (47.37)   | 1790 (52.69)| 39,417 (47.18)| 28.09  | < 0.001 |
| Multiparous               | 32,062 (52.63)   | 1613 (47.31)| 30,449 (52.82)|         |         |
| **Diabetes**              |                  |          |              |         |         |
| Yes                       | 770 (0.90)       | 38 (0.98)| 732 (0.90)   | 0.19    | 0.664   |
| No                        | 72,276 (99.10)   | 3368 (99.02)| 68,908 (99.10)|         |         |
| **Heart disease**         |                  |          |              |         |         |
| Yes                       | 278 (0.32)       | 21 (0.53)| 257 (0.32)   | 2.78    | 0.095   |
| No                        | 72,998 (99.68)   | 3393 (99.47)| 69,605 (99.68)|         |         |
| **Renal disease**         |                  |          |              |         |         |
| Yes                       | 121 (0.13)       | 4 (0.10)| 117 (0.13)   | 0.14    | 0.708   |
| No                        | 73,211 (99.87)   | 3410 (99.90)| 69,801 (99.87)|         |         |
| **Autoimmune disease**    |                  |          |              |         |         |
| Yes                       | 206 (0.16)       | 23 (0.69)| 183 (0.14)   | 52.33   | < 0.001 |
| No                        | 72,764 (99.84)   | 3391 (99.31)| 69,373 (99.86)|         |         |
| **Hyperthyroidism**       |                  |          |              |         |         |
| Yes                       | 270 (0.39)       | 15 (0.44)| 255 (0.38)   | 0.19    | 0.663   |
| No                        | 72,661 (99.61)   | 3396 (99.56)| 69,265 (99.62)|         |         |
| **Hypothyroidism**        |                  |          |              |         |         |
| Yes                       | 991 (0.88)       | 34 (1.13)| 957 (0.87)   | 1.72    | 0.190   |
| No                        | 71,922 (99.12)   | 3377 (98.87)| 68,545 (99.13)|         |         |
| **Asthma**                |                  |          |              |         |         |
unaffected areas. The results showed the incidence of 4 adverse birth outcomes were all higher in earthquake exposed group. After reducing the imbalance between the exposed and unexposed group based on CEM method, we observed the long-term effects of maternal earthquake exposure on PTB among male births only. The findings mentioned above still held within singleton births, all livebirth, and singleton livebirths, and remained unaffected by maternal migrant status.

Previous studies on disaster-related short-term adverse birth outcomes have produced mixed results between different earthquakes, with some studies, including one in Sichuan, China, showing increased risks of LBW, PTB, and SGA [7, 8, 24], while others studies suggesting nonsignificant changes in adverse birth outcomes [25, 26]. Furthermore, the findings on the same disasters, Hurricane Katrina for instance, have also been inconsistent [27, 28], indicating further research is likely needed to understand the effects on birth outcomes.

The few existing studies of natural and manmade disasters also turned up conflicting evidence of their long-term effects on adverse birth outcomes. One study on the 2001 World Trade Center disaster indicated that the effects of 11 September attacks on reproductive outcomes may last within the first 2 years, and reduce to zero in 3 years or later [10]. However, another study on the 2011 Fukushima Daiichi nuclear disaster showed no changes in birth outcomes within the 3 years following the Fukushima disaster [11]. Our findings suggested that Wenchuan earthquake increases the risks of PTB, but

| Table 1 Descriptive Statistics of potential covariables among Wenchuan exposure and non-exposure group (Continued) |
|---------------------------------------------------------------|
| Characteristic                                         | No. (Weighted %) | Exposure | Non-exposure | χ²  | P value |
|---------------------------------------------------------------|
| Yes                                                        | 57 (0.05)        | 2 (0.06) | 55 (0.05)    | 0.12 | 0.732   |
| No                                                         | 73,232 (99.95)   | 3411 (99.94) | 69,821 (99.95) |         |         |
| Gestational Diabetes                                       |                   |          |              |      |         |
| Yes                                                        | 7372 (11.43)     | 173 (5.99) | 7199 (11.63) | 64.48 | < 0.001 |
| No                                                         | 65,017 (88.57)   | 3234 (94.01) | 61,783 (88.37) |         |         |
| Gestational hypertension                                   |                   |          |              |      |         |
| Yes                                                        | 3415 (4.16)      | 149 (4.06) | 3266 (4.17)  | 0.06  | 0.802   |
| No                                                         | 69,004 (95.84)   | 3233 (95.94) | 65,771 (95.83) |         |         |
| Gestational anemia                                         |                   |          |              |      |         |
| Yes                                                        | 9293 (13.26)     | 147 (4.90) | 9146 (13.55) | 167.22 | < 0.001 |
| No                                                         | 64,029 (86.74)   | 3268 (95.10) | 60,761 (86.45) |         |         |
| Gestational hyperthyroidism                                |                   |          |              |      |         |
| Yes                                                        | 300 (0.32)       | 10 (0.27) | 290 (0.32)   | 0.20  | 0.658   |
| No                                                         | 72,636 (99.68)   | 3405 (99.73) | 69,231 (99.68) |         |         |
| Gestational hypothyroidism                                 |                   |          |              |      |         |
| Yes                                                        | 2024 (1.90)      | 58 (2.16) | 1966 (1.90)  | 0.73  | 0.394   |
| No                                                         | 70,910 (98.10)   | 3357 (97.84) | 67,553 (98.10) |         |         |
| Gestational asthma                                         |                   |          |              |      |         |
| Yes                                                        | 45 (0.05)        | 1 (0.02) | 44 (0.05)    | 0.69  | 0.407   |
| No                                                         | 73,350 (99.95)   | 3414 (99.98) | 69,936 (99.95) |         |         |
| Multiple pregnancy                                         |                   |          |              |      |         |
| Yes                                                        | 1569 (1.78)      | 105 (2.72) | 1464 (1.75)  | 10.76 | 0.001   |
| No                                                         | 71,921 (98.22)   | 3311 (97.28) | 68,610 (98.25) |         |         |
| Mode of delivery                                           |                   |          |              |      |         |
| Cesarean                                                   | 27,096 (36.77)   | 1383 (32.47) | 25,713 (36.34) | 332.93 | < 0.001 |
| Vaginal                                                    | 46,073 (63.23)   | 1794 (42.57) | 44,279 (63.66) |         |         |
| Infant sex                                                 |                   |          |              |      |         |
| Female                                                     | 34,170 (47.30)   | 1630 (47.97) | 32,534 (47.27) | 0.45  | 0.504   |
| Male                                                       | 39,323 (52.70)   | 1780 (52.03) | 37,543 (52.73) |         |         |
not all adverse birth outcomes, among the birth 7 years after the disaster, and female births seem to be immune to the earthquake exposure.

While it is difficult to compare the findings from different disasters directly, joint consideration of the commonalities may help to understand the mechanisms underlying disaster-related adverse birth outcomes [10]. Disaster-related adverse birth outcomes are associated with male vulnerability in utero perturbations, which could be possibly explained by genetic contributions [29]. The sex-specific X-linked genes expression in placenta may confer protection by fostering adaptability to survive [29, 30]. Disaster-related stress is another potential pathway from disaster occurrence to adverse birth outcomes [31]. Preterm birth, in evolutionary context, can be understood as a predictive adaptive response to the perceived deprived environment to increase the chances of survival.

### Table 2

| Birth outcome       | No. (Weighted %) | Exposure | Non-exposure | χ²   | P value |
|---------------------|------------------|----------|--------------|------|---------|
| Outcome of delivery |                  |          |              |      |         |
| Stillbirth          | 873 (1.35)       | 75 (2.00)| 798 (1.33)   | 5.63 | 0.018   |
| Livebirth           | 72,580 (98.65)   | 3340 (98.00)| 69,240 (98.67)|      |         |
| Weeks of gestation  |                  |          |              |      |         |
| Preterm             | 6290 (7.55)      | 503 (14.14) | 5787 (7.32) | 75.82| < 0.001 |
| Term                | 65,337 (92.01)   | 2808 (85.46) | 62,529 (92.24)|      |         |
| Post-term           | 250 (0.44)       | 17 (0.41) | 233 (0.44)   |      |         |
| Birth weight        |                  |          |              |      |         |
| Low                 | 4859 (5.93)      | 392 (10.82) | 4467 (5.76) | 47.91| < 0.001 |
| Normal              | 63,658 (87.44)   | 2833 (83.77) | 60,825 (87.57)|      |         |
| High                | 4728 (6.63)      | 174 (5.42) | 4554 (6.67)  |      |         |
| Birth weight for gestational age |          |          |              |      |         |
| Small               | 6877 (9.58)      | 394 (11.32) | 6483 (9.52) | 4.83 | 0.008   |
| Appropriate         | 59,415 (80.86)   | 2716 (79.93) | 56,699 (80.89)|      |         |
| Large               | 6977 (9.56)      | 292 (8.75) | 6685 (9.59)  |      |         |

### Table 3

| Birth outcome       | Adjusted Risk Ratio a (95% Confidence Interval, P value) |
|---------------------|----------------------------------------------------------|
|                     | Model 1 (All births) | Model 2 (Singleton births) |         |
| Total               |                        |                           |         |
| Stillbirth          | 1.21 (0.71–2.04, 0.484)| 1.29 (0.75–2.23, 0.359) |         |
| Preterm birth       | 1.25 (1.06–1.48, 0.010)| 1.22 (1.02–1.47, 0.033) |         |
| Low birth weight    | 1.08 (0.89–1.31, 0.430)| 1.03 (0.84–1.28, 0.755) |         |
| Small for gestational age | 0.93 (0.71–1.22, 0.583) | 0.91 (0.68–1.21, 0.524) |         |
| Girls               |                        |                           |         |
| Stillbirth          | 1.14 (0.58–2.26, 0.705)| 1.28 (0.63–2.63, 0.494) |         |
| Preterm             | 1.09 (0.84–1.41, 0.535)| 1.07 (0.81–1.42, 0.638) |         |
| Low birth weight    | 1.00 (0.76–1.32, 0.991)| 0.98 (0.72–1.33, 0.897) |         |
| Small for gestational age | 0.85 (0.55–1.33, 0.489) | 0.84 (0.53–1.35, 0.474) |         |
| Boys                |                        |                           |         |
| Stillbirth          | 1.25 (0.57–2.75, 0.579)| 1.28 (0.57–2.86, 0.547) |         |
| Preterm             | 1.40 (1.12–1.75, 0.002)| 1.36 (1.07–1.73, 0.012) |         |
| Low birth weight    | 1.17 (0.89–1.53, 0.253)| 1.09 (0.82–1.47, 0.547) |         |
| Small for gestational age | 1.03 (0.84–1.26, 0.788) | 1.01 (0.82–1.24, 0.943) |         |

*aAdjusted maternal diseases, history of PTB, method of conception, parity, multiple pregnancy or singleton pregnancy, mode of delivery and sex of newborns*
and reproduction [32, 33]. Maternal psychosocial stress, including post-traumatic stress, could serve as an independent trigger for PTB [18, 29], given that a previous study of 1986 Chernobyl accident identified radiation-related anxiety as the prime culprit of PTB [34] and Wenchuan earthquake-related posttraumatic stress symptoms remained especially common among female survivors 5 years after the earthquake [35]. Disaster-related stress, in our opinion, could be main potential factor behind the long-term adverse effects of Wenchuan earthquake exposure on birth outcomes. The heightened maternal stress hormones, including glucocorticoids, may service an important biological mechanism behind the long-lasting earthquake-related adverse birth outcomes [36].

Our study has important implications for policy and clinical practice. Policy makers should acknowledge the long-lasting adverse effects of the earthquake on reproductive health in post-disaster recovery and invest long-term psychological support for the female survivors. Obstetricians also could reduce the risks of adverse birth outcomes by identifying the pregnant women experienced earthquake and providing targeted psychological interventions.

Limitations
There are some important limitations should be noted. First, the exposure assessment was defined by the geographical location of the delivery hospitals, hence we restricted to the pregnant women with Sichuan hukou in stratified analysis to make sure exposure group actually experienced the disaster. Second, although we have adjusted for the confounders available in the dataset, the between-group comparability may still compromise the validity of our findings, given that some inherent non-comparability between groups was less likely to removed. Difference in difference analysis (before-after earthquake comparsion with a parallel design) may be useful, but the datasets before the earthquake are not available. Third, we were unable to distinguish between the earthquake-related stress and other social stressors due to limitations in data availability. Fourth, the dose-response relationship between distance from the epicenter of the earthquake and incidence of PTB could support our findings. However, the sample size in Sichuan province is not large enough to allow us to conduct the dose-response analysis. Last, we only included secondary and tertiary hospitals in this study, which may introduce select bias and underestimate the associations. However, few pregnant women delivery in home or primary hospitals in China, and mostly due to exceptional circumstances [15, 37]. Considering all of the limitations mentioned above, the long-term risk of maternal exposure to Wenchuan earthquake on offspring birth outcomes should be interpreted with caution.

Conclusions
Maternal exposure to the 2008 Wenchuan earthquake elevates the risk of PTB in offspring several years later after the earthquake, and only the male birth is vulnerable to the long-term effect. Maternal acute exposure to disasters could serve as a major monitor for long-term reproductive outcomes, and more research is needed to understand the underlining reasons for long-term disaster-related adverse birth outcomes.

Supplementary information
Supplementary information accompanies this paper at https://doi.org/10.1186/s12884-020-03206-1.

Table 4 Associations of maternal exposure to Wenchuan earthquake with adverse birth outcomes among livebirths

| Birth outcome | Adjusted Risk Ratio* (95% Confidence Interval, P value) |
|--------------|------------------------------------------------------|
|              | Model 1 (All livebirths) | Model 2 (Singleton livebirths) |
| Total        |                                                      |
| Preterm birth| 1.27 (1.06–1.51, 0.009) | 1.24 (1.02–1.50, 0.031) |
| Low birth weight| 1.09 (0.89–1.33, 0.427) | 1.02 (0.81–1.27, 0.857) |
| Small for gestational age| 0.91 (0.69–1.20, 0.505) | 0.89 (0.66–1.19, 0.417) |
| Girls        |                                                      |
| Preterm      | 1.06 (0.81–1.40, 0.675) | 1.04 (0.77–1.41, 0.798) |
| Low birth weight| 0.98 (0.73–1.31, 0.871) | 0.93 (0.67–1.29, 0.676) |
| Small for gestational age| 0.85 (0.54–1.35, 0.499) | 0.83 (0.52–1.34, 0.451) |
| Boys         |                                                      |
| Preterm      | 1.47 (1.18–1.84, 0.001) | 1.43 (1.12–1.82, 0.004) |
| Low birth weight| 1.22 (0.93–1.61, 0.154) | 1.13 (0.83–1.54, 0.420) |
| Small for gestational age| 0.99 (0.80–1.22, 0.913) | 0.96 (0.78–1.20, 0.738) |

*Adjusted maternal diseases, history of PTB, method of conception, parity, multiple pregnancy or singleton pregnancy, mode of delivery and sex of newborns.
Wenchuan earthquake with adverse birth outcomes among all livebirths of local pregnant women.

Abbreviations
CEM: Coarsened exact matching; PTB: Preterm birth; LBW: Low birthweight; SGA: Small for gestational age; CLDS: China Labor and Delivery Survey; DCC: Data coordination center; WHO: World Health Organization; BMI: Body mass index; aRR: Adjusted risk ratio; CI: Confidence interval

Acknowledgments
We are grateful to the members of CLDS data coordination center for their support of this project.

Authors’ contributions
LZ conceived the study and had primary responsibility for editing of the manuscript. JN and SL contributed to study design and were involved in curation of data. QL developed the analysis plan, analysed the data and drafted the manuscript. JZ and JL assisted with interpretation of the results. All authors participated in manuscript revision and approval of the final version.

Funding
The study was supported by Shanghai Municipal Health Commission (201640032) and Shanghai Hospital Development Center (SHDC12016204), and approved by World Health Organization (A65899).

Availability of data and materials
The datasets used and/or analysed during the current study are available from the corresponding author on reasonable request.

Ethics approval and consent to participate
The CLDS was approved by the WHO Research Ethics Review Committee and the corresponding author on reasonable request. The datasets used and/or analysed during the current study available from the corresponding author on reasonable request.

Consent for publication
Not applicable.

Competing interests
None declared.

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Received: 2 July 2020 Accepted: 24 August 2020

Published online: 22 September 2020

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