

SCIENTIFIC REPORTS



OPEN

Associations of maternal pre-pregnancy body mass index and gestational weight gain with birth outcomes in Shanghai, China

Lingli Xiao^{1,*}, Guodong Ding^{2,*}, Angela Vinturache³, Jian Xu⁴, Yifang Ding¹, Jialin Guo¹, Liping Huang¹, Xuelei Yin¹, Jing Qiao², Inesh Thureraja⁵ & Xiaoming Ben¹

Received: 17 October 2016
Accepted: 12 December 2016
Published: 25 January 2017

Recent data suggests that abnormal maternal pre-pregnancy body mass index (BMI) or gestational weight gain (GWG) is associated with unfavorable delivery outcomes. However, limited clinical evidence is available to support this correlation in China. Participating 510 mother-infant pairs were recruited from the Shanghai First Maternity and Infant Hospital, China, between January 1st and 30th 2016. Maternal pre-pregnancy BMI was categorized according to the China's classification and GWG according to the 2009 Institute of Medicine recommendations (IOM). Linear regression tested the associations between pre-pregnancy BMI or GWG and length of gestation, birthweight, length, and head circumference. Logistic regression assessed the associations between pre-pregnancy BMI or GWG and macrosomic, small- (SGA) and large- (LGA) for-gestational-age infants. Overweight/obese women showed increased length of gestation and birthweight, but did not have a higher risk of macrosomic and LGA infants compared with normal weight women. Women with excessive GWG showed increased length of gestation, birthweight, length, and head circumference, and were more likely to deliver macrosomic and LGA infants compared with women with adequate GWG. Although a relatively low proportion of women from Shanghai area are overweight/obese or exhibit excessive GWG, both high pre-pregnancy BMI and excessive GWG influence perinatal outcomes.

Maternal nutritional status, reflected by pre-pregnancy body mass index (BMI) and gestational weight gain (GWG), is perceived as an important predictor of perinatal and even long-term outcomes for mothers and children. Accumulating evidence suggests that higher BMI prior to pregnancy is associated with an increased risk of preeclampsia, gestational diabetes, caesarean section, postpartum hemorrhage, and fetal macrosomia^{1–4}, whereas maternal underweight is associated with an increased risk of preterm delivery and small-for-gestational-age (SGA)^{5,6}. Maternal GWG has also been widely studied as an independent indicator of adverse pregnancy outcomes. Excessive GWG is associated with large-for-gestational-age (LGA), gestational diabetes, caesarean section, and postpartum weight retention^{7–9}, while insufficient GWG is linked with low birthweight, preterm delivery, and SGA birth^{10,11}. However, of the majority of the studies on the association between non-optimal maternal BMI or GWG and adverse pregnancy outcomes were conducted in developed countries of Europe and North America^{12–14}. Furthermore, the extent that maternal BMI or GWG affects pregnancy outcomes varies among studies, among factors contributing to these differences are variations in the cut-offs used for BMI and GWG values, and characteristics of population studied such as demographics and ethnicity. For instance, women belonging to ethnic groups characterized by a small body size have been reported to gain less weight on average during pregnancy than larger women (Caucasians)¹⁵. In developing Asian countries, women generally have a lower BMI

¹Department of Neonatology, Shanghai First Maternity and Infant Hospital, Tongji University School of Medicine, Shanghai 201204, China. ²Department of Pediatrics, Shanghai East Hospital, Tongji University School of Medicine, Shanghai 200120, China. ³Department of Obstetrics & Gynaecology, John Radcliffe Hospital, Oxford University Hospital NHS Foundation Trust, Headley Way, Oxford, OX3 9DU, UK. ⁴Department of Pediatrics, Punan Hospital of Pudong New District, Shanghai 200125, China. ⁵Bachelor of medicine and bachelor of surgery (MBBS) 2012, Tongji University School of Medicine, Shanghai 200092, China. *These authors contributed equally to this work. Correspondence and requests for materials should be addressed to X.B. (email: benxm@163.com)

and a smaller GWG than those reported in developed countries from Europe and North America¹⁶. To date the information on the impact of maternal pre-pregnancy BMI and GWG is scarce, especially in developing Asian countries like China.

China, the world's largest developing country, once regarded to have one of the leanest populations, is gradually catching up with western countries in terms of prevalence of overweight and obesity. The increases in the prevalence of overweight and obesity may be attributable to the China's economic development, which has brought with it a high degree of westernization of the traditional Chinese diet and lifestyle¹⁷. The National Nutrition and Health Survey published in 2005 showed that, in China, the prevalence of overweight (BMI ≥ 24 kg/m²) in women 18–44 years of age increased from 16.8 to 21.8%, and the prevalence of obesity (BMI ≥ 28 kg/m²) increased from 3.1 to 6.1% from 1992 to 2002¹⁸. According to the 2009 Institute of Medicine (IOM) recommendations, more than half of childbearing-aged women living in north China exhibited excessive GWG^{19,20}. The increasing prevalence of unhealthy preconception BMI and inappropriate GWG among women is of great concern for the Chinese public health community. Until now, however, there have been only a few reports evaluating the influence of maternal pre-pregnancy BMI and GWG on delivery outcomes in Chinese women^{4,19–21}. In the present study, we aimed to investigate the current weight status of childbearing-aged women in Shanghai and to examine the possible relationships of non-optimal GWG and pre-pregnancy BMI with birth outcomes.

Methods

Study population. This retrospective cohort study was conducted in the Shanghai First Maternity and Infant Hospital, Tongji University School of Medicine, Shanghai, China. The hospital is one of the largest maternity hospitals with an annual delivery volume of approximately 25,000 babies, accounting for 12% of total number of deliveries in Shanghai. Eligibility criteria included: singleton pregnancy; age over 18 years old; residence in the metropolitan Shanghai area for at least 3 years prior to pregnancy; no report of assisted reproduction (spontaneous conception), pre-existing diabetes mellitus or gestational diabetes, chronic or pregnancy-associated hypertension, thyroid disorders, HIV infection or AIDS, and illicit drug use. Women were recruited to the study after delivery. From January 1st to 30th 2016, a total of 605 women met the eligibility criteria, among whom 519 women agreed to take part in this study (response rate 85.8%). Of these women, 9 women with missing values for major confounders were excluded. Therefore, 510 mothers were included in this study. Each woman participating in the study signed an informed consent form, and the research protocol for this study was approved by the Medical Ethics Committee of Shanghai First Maternity and Infant Hospital, Tongji University School of Medicine. All methods were performed in accordance with the relevant guidelines and regulations.

Maternal interviews. Standardized face-to-face interviews were conducted with the women shortly after delivery by specially trained registered nurses in the hospital. The questionnaire collected information about: demographics (maternal age, education, marital status, and household income) and maternal characteristics (pre-pregnancy weight, and smoking and alcohol use during pregnancy). Other relevant information such as previous pregnancies and any medical conditions, current pregnancy complications, self-reported last menstrual period (LMP), and weight prior to delivery was obtained by interview and confirmed by maternal medical records. Clinical estimates of gestational age (ultrasound) were also obtained from maternal medical records. Gestational age was estimated based on the date of the self-reported LMP; if the LMP was unreliable or if there was a significant discordance between the clinical estimate and LMP (>2 weeks), the clinical estimation of gestational age was used²². Maternal height was measured during the interview. If the questionnaire was incomplete, after discharge, the mothers were contacted by telephone to obtain the missing information.

Exposure information. The main exposures of interest were maternal pre-pregnancy BMI and GWG. At enrollment, from the maternal self-reported pre-pregnancy weight and measured height, we calculated pre-pregnancy BMI [weight (kg)/height (m)²]. In China, the two most commonly used BMI classifications for adults are the World Health Organization (WHO) standard and the recently developed Chinese BMI standard²³. However, the Chinese standard that has lower cutoff points for BMI categories is more recommended in China because of recent evidence suggesting that Chinese people, as well as several other Asian Pacific populations, have an elevated risk for obesity-related diseases or conditions at a lower BMI than Caucasians²⁴. Thus, the Chinese BMI classification standard was used in this study due to its better sensitivity and specificity for identifying risk factors such as hypertension, diabetes, and dyslipidemia in the Chinese population²³. Therefore, women were divided into four groups based on pre-pregnancy BMI according to categories defined by the Working Group on Obesity in China as follows: underweight (BMI < 18.5 kg/m²), normal weight (18.5 kg/m² \leq BMI < 24.0 kg/m²), overweight (24.0 kg/m² \leq BMI < 28.0 kg/m²), and obese (BMI \geq 28.0 kg/m²)²³. Given the small number of women meeting the criteria for “obese” (n = 6), for meaningful comparisons this category was combined with the “overweight” category and collectively referred as overweight/obese. Statistical comparisons were carried out between three BMI groups: (1) “underweight”, (2) “normal weight”, and (3) “overweight/obese”. Women with normal BMI (18.5–24.0 kg/m²) were used as reference group for the analysis.

Maternal GWG was calculated as the difference between self-reported pre-pregnancy weight and the last clinically measured weight recorded prior to delivery. Since the Chinese Society of Obstetrics and Gynecology did not have an official guideline for weight gain during pregnancy, maternal GWG was categorized as inadequate, adequate, and excessive according to the 2009 Institute of Medicine (IOM) recommendations²⁵. Inadequate GWG was defined as a weight gain during pregnancy of <12.5 kg in underweight women, <11.5 kg in normal weight women, <7 kg in overweight women, and <5 kg in obese women. Adequate GWG was defined as a weight gain during pregnancy of 12.5–18 kg in underweight women, 11.5–16 kg in normal weight women, 7–11.5 kg in overweight women, and 5–9 kg in obese women. Excessive GWG was defined as a weight gain during pregnancy of

>18 kg in underweight women, >16 kg in normal weight women, >11.5 kg in overweight women, and >9 kg in obese women. Women with adequate GWG were used as reference group for the analysis.

Outcome assessment. Neonatal outcomes included birthweight, length, head circumference, length of gestation, macrosomia, and SGA and LGA births. We were not able to analyze low birthweight ($n = 5$) and preterm delivery ($n = 8$) because of small number of infants who met these criteria. Infant sex, birth date, method of delivery, birthweight, crown-heel length, and head circumference were obtained from hospital delivery logs and medical records.

Low birthweight was defined as birthweight <2,500 g, whereas fetal macrosomia was defined as birthweight >4,000 g regardless of gestational age. A SGA infant was defined as an infant having a birthweight <10th percentile, whereas a LGA infant was defined as an infant having a birthweight >90th percentile for gestational age and sex according to a Chinese reference curve²⁶. Preterm delivery was defined as birth at <37 completed weeks of gestation.

Statistical analysis. Descriptive statistics were used to present the characteristics of the study population. Linear regression models were conducted to calculate beta ($\hat{\alpha}$) coefficients and 95% confidence intervals (CIs) evaluating the associations between exposures and length of gestation, birthweight, length, and head circumference (continuous variables). Logistic regression models were conducted to calculate odds ratios (ORs) and 95% CIs evaluating the associations between exposures and macrosomia, SGA and LGA births (dichotomous variables, yes/no).

We selected potential confounders for the multivariate linear or logistic models based on the associations reported in the literature and on the statistical considerations. We included in the final models as potential confounders those variables that were associated at p -values ≤ 0.2 with two or more birth outcomes in bivariate analysis. Confounders included were: maternal age, education, parity, smoking, alcohol use, infant sex, and household monthly salary. Multivariate models of birthweight, length, head circumference, macrosomia, SGA, and LGA were also adjusted for gestational age. Maternal pre-pregnancy BMI and GWG were mutually adjusted.

All results of regression models are expressed as comparisons with women with normal pre-pregnancy BMI or adequate GWG. Statistical analyses were carried out using SPSS software (SPSS Inc., Chicago, IL) based on two-tailed tests, and $p < 0.05$ indicated statistical significance.

Results

Table 1 presents the socio-demographic characteristics of the study population. The mean (SD) pre-pregnancy BMI was 20.5 (2.6) kg/m², and GWG was 15.1 (4.5) kg. With respect to pre-pregnancy BMI, 23.5% of the women were underweight, 65.9% were normal weight, and 10.6% were overweight/obese prior to pregnancy. Based on the IOM recommendations (2009), 20.6% of the women had inadequate GWG, 43.5% had adequate (normal) GWG, and 35.9% had excessive GWG. Maternal age ranged from 18 to 44 years old, and almost 80% of the women were between 25 and 34 years old. The vast majority (89.2%) of women had at least 13 years of education, three-quarters (75.3%) were primiparous, and two-thirds (65.5%) lived in households with a monthly income between RMB 10,000 and 30,000 yuan (per capita monthly income was 12,900 yuan in Shanghai reported in 2015). Nearly one-third (30.4%) of the women lived with a smoker during pregnancy or smoked themselves, although a few (6.3%) consumed alcohol regularly (at least once every two weeks).

Overall, 53.5% of the newborn infants were male. The mean (SD) birthweight, length, and head circumference were 3376.9 (447.7) g, 50.1 (0.7) cm, and 34.2 (1.1) cm, respectively. A total of 6.9% ($n = 35$) of infants were macrosomic, 10.6% ($n = 54$) were LGA, and 6.3% ($n = 32$) were SGA. However, very few infants were born of low birthweight (birthweight <2,500 g) (1.0%, $n = 5$) or preterm (1.6%, $n = 8$) (gestational age <37 weeks).

The unadjusted and adjusted linear regression results for the association of maternal pre-pregnancy BMI and GWG with birth outcomes are shown in Table 2. After adjusting for confounders, increased length of gestation ($\beta = 0.40$, 95% CI = 0.11 to 0.49) was found in overweight/obese women when compared with normal-weight women, but no associations were seen for underweight women. Similarly, women with excessive GWG also showed increased length of gestation ($\beta = 0.23$, 95% CI = 0.03 to 0.43) compared with women who had adequate GWG, but no associations were found in women who had inadequate GWG.

There were all significant positive associations of maternal pre-pregnancy BMI and GWG with birthweight (Table 2). Infants of overweight/obese women had increased birthweight ($\beta = 144.8$, 95% CI = 144.5 to 145.1), whereas infants of underweight women showed decreased birthweight ($\beta = -104.2$, 95% CI = -104.4 to -104.0), when compared with those of normal-weight women. Similarly, infants of women with excessive GWG showed increased birthweight ($\beta = 216.3$, 95% CI = 216.1 to 216.5), and infants with women with inadequate GWG showed decreased birthweight ($\beta = -120.5$, 95% CI = -120.8 to -120.3), when compared with those of women who had adequate GWG.

No significant associations were found between maternal pre-pregnancy BMI and birth length or head circumference. However, increased birth length ($\beta = 0.29$, 95% CI = 0.09 to 0.49) and head circumference ($\beta = 0.33$, 95% CI = 0.13 to 0.52) were observed in infants of women with excessive GWG when compared with women who had adequate GWG, but no associations were found in women with inadequate GWG (Table 2).

Because of the strong positive associations between pre-pregnancy BMI and GWG and birthweight found in the linear regression analyses, we further examined the potential relationships of pre-pregnancy BMI and GWG with macrosomia, SGA or LGA infants in logistic regression analysis. Table 3 summarizes the odds of macrosomia, SGA or LGA infants in women with non-optimal BMI or GWG. Overweight/obese women did not show an increase in the risk of macrosomia and LGA infants. Conversely, underweight women did not have an increased risk of SGA infants when compared with normal-weight women. However, women with excessive GWG exhibited

| | Total sample (n = 510) | Pre-pregnancy BMI category | | | Gestational weight gain category | | |
|--------------------------------|---------------------------|----------------------------|----------------------------|----------------------------------|----------------------------------|-----------------------|---------------------|
| | | Underweight (n = 120) | Normal weight (n = 336) | Overweight and obese (n = 54) | Inadequate (n = 105) | Adequate (n = 222) | Excessive (n = 183) |
| Maternal characteristic | | | | | | | |
| Age (years) | | | | | | | |
| <25 | 23 (4.5%) | 4 (3.3%) | 17 (5.1%) | 2 (3.7%) | 5 (4.8%) | 9 (4.1%) | 9 (4.9%) |
| 25–29 | 199 (39.0%) | 63 (52.5%) | 116 (34.5%) | 20 (37.0%) | 35 (33.3%) | 89 (40.1%) | 75 (41.0%) |
| 30–34 | 206 (40.4%) | 41 (34.2%) | 144 (42.9%) | 21 (38.9%) | 50 (47.6%) | 86 (38.7%) | 70 (38.3%) |
| ≥35 | 82 (16.1%) | 12 (10.0%) | 59 (17.6%) | 11 (20.4%) | 15 (14.3%) | 38 (17.1%) | 29 (15.8%) |
| Education (years) | | | | | | | |
| ≤12 (High school) | 55 (10.8%) | 12 (10.0%) | 33 (9.8%) | 10 (18.5%) | 5 (4.8%) | 19 (8.6%) | 31 (16.9%) |
| 13–16 (Undergraduate school) | 349 (68.4%) | 82 (68.3%) | 230 (68.5%) | 37 (68.5%) | 62 (59.0%) | 163 (73.4%) | 124 (67.8%) |
| ≥17 (Postgraduate school) | 106 (20.8%) | 26 (21.7%) | 73 (21.7%) | 7 (13.0%) | 38 (36.2%) | 40 (18.0%) | 28 (15.3%) |
| Parity | | | | | | | |
| 0 (Primiparous) | 384 (75.3%) | 99 (82.5%) | 241 (71.7%) | 44 (81.5%) | 80 (76.2%) | 163 (73.4%) | 141 (77.0%) |
| ≥1 (Multiparous) | 126 (24.7%) | 21 (17.5%) | 95 (28.3%) | 10 (18.5%) | 25 (23.8%) | 59 (26.6%) | 42 (23.0%) |
| Household monthly salary (¥) | | | | | | | |
| <10000 | 85 (16.7%) | 22 (18.3%) | 51 (15.2%) | 12 (22.2%) | 17 (16.2%) | 33 (14.9%) | 35 (19.1%) |
| 10000–20000 | 196 (38.4%) | 50 (41.7%) | 126 (37.5%) | 20 (37.0%) | 36 (34.3%) | 82 (36.9%) | 78 (42.6%) |
| 20001–30000 | 138 (27.1%) | 30 (25.0%) | 95 (28.3%) | 13 (24.1%) | 31 (29.5%) | 64 (28.8%) | 43 (23.5%) |
| >30000 | 91 (17.8%) | 18 (15.0%) | 64 (19.0%) | 9 (16.7%) | 21 (20.0%) | 43 (19.4%) | 27 (14.8%) |
| Smoking during pregnancy | | | | | | | |
| Yes/Lived with smoker | 155 (30.4%) | 35 (29.2%) | 106 (31.5%) | 14 (25.9%) | 76 (72.4%) | 152 (68.5%) | 127 (69.4%) |
| No | 355 (69.6%) | 85 (70.8%) | 230 (68.5%) | 40 (74.1%) | 29 (27.6%) | 70 (31.5%) | 56 (30.6%) |
| Alcohol use during pregnancy | | | | | | | |
| Yes | 32 (6.3%) | 9 (7.5%) | 18 (5.4%) | 5 (9.3%) | 7 (6.7%) | 12 (5.4%) | 13 (7.1%) |
| No | 478 (93.7%) | 111 (92.5%) | 318 (94.6%) | 49 (90.7%) | 98 (93.3%) | 210 (94.6%) | 170 (92.9%) |
| Infant characteristic | | | | | | | |
| Sex | | | | | | | |
| Male | 273 (53.5%) | 62 (51.7%) | 186 (55.4%) | 25 (46.3%) | 65 (61.9%) | 103 (46.4%) | 105 (57.4%) |
| Female | 237 (46.5%) | 58 (48.3%) | 150 (44.6%) | 29 (53.7%) | 40 (38.1%) | 119 (53.6%) | 78 (42.6%) |
| Gestational age (weeks) | | | | | | | |
| <37 | 8 (1.6%) | 3 (2.5%) | 5 (1.5%) | 0 (0.0%) | 3 (2.9%) | 4 (1.8%) | 1 (0.5%) |
| ≥37 | 502 (98.4%) | 117 (97.5%) | 331 (98.5%) | 54 (100.0%) | 102 (97.1%) | 218 (98.2%) | 182 (99.5%) |
| Birthweight (g) | | | | | | | |
| <2500 | 5 (1.0%) | 2 (1.7%) | 3 (0.9%) | 0 (0.0%) | 2 (1.9%) | 1 (0.5%) | 2 (1.1%) |
| 2500–4000 | 470 (92.2%) | 115 (95.8%) | 307 (91.4%) | 48 (88.9%) | 97 (92.4%) | 217 (97.7%) | 156 (85.2%) |
| ≥4000 | 35 (6.9%) | 3 (2.5%) | 26 (7.7%) | 6 (11.1%) | 6 (5.7%) | 4 (1.8%) | 25 (13.7%) |
| Fetal growth | | | | | | | |
| SGA | 32 (6.3%) | 8 (6.7%) | 23 (6.8%) | 1 (1.9%) | 15 (14.3%) | 14 (6.3%) | 3 (1.6%) |
| AGA | 424 (83.1%) | 107 (89.2%) | 272 (81.0%) | 45 (83.3%) | 83 (79.0%) | 197 (88.7%) | 144 (78.7%) |
| LGA | 54 (10.6%) | 5 (4.2%) | 41 (12.2%) | 8 (14.8%) | 7 (6.7%) | 11 (5.0%) | 36 (19.7%) |
| Birthweight (g) (mean ± SD) | 3376.9 ± 447.7 | 3268.2 ± 3367.9 | 3390.2 ± 465.0 | 3535.9 ± 447.8 | 3180.3 ± 397.8 | 3302.9 ± 365.5 | 3579.5 ± 487.4 |
| Birth length (cm) | 50.1 ± 0.7 | 50.0 ± 0.4 | 50.1 ± 0.7 | 50.2 ± 0.9 | 50.0 ± 0.6 | 50.0 ± 0.5 | 50.3 ± 0.9 |
| Head circumference (cm) | 34.2 ± 1.1 | 34.1 ± 0.9 | 34.2 ± 1.1 | 34.4 ± 1.4 | 34.0 ± 1.1 | 34.1 ± 1.0 | 34.5 ± 1.1 |

Table 1. Maternal and infant's characteristics according to pre-pregnancy body mass index (BMI) and gestational weight gain (GWG) (n = 510).

increased risk of macrosomia (OR = 7.49, 95% CI = 2.42 to 23.19) and LGA infants (OR = 3.79, 95% CI = 1.82 to 7.90), whereas women with inadequate GWG exhibited increased risks of SGA infants (OR = 3.60, 95% CI = 1.57 to 8.30), when compared with women who had adequate GWG.

Discussion

To the best of our knowledge, this is the first study from the metropolitan Shanghai area to assess birth outcomes with respect to pre-pregnancy BMI and GWG. The main findings of this study are that maternal excessive GWG but not overweight/obesity was associated with increased risk of macrosomic and LGA infants, whereas maternal inadequate GWG but not underweight was associated with an increased risk of SGA infants in a sample of women representative for the population of Shanghai. We also found that the proportion of underweight women prior to pregnancy was relatively high, whereas the proportion of women with excessive GWG was low.

| Maternal weight status (No.) | Birth outcomes | | | | | | | |
|--------------------------------------------------|-----------------------------|-------------------------------------|------------------------------|-------------------------------------|------------------------|-------------------------------------|-------------------------|-------------------------------------|
| | Length of gestation (weeks) | | Birthweight (g) | | Length (cm) | | Head circumference (cm) | |
| | Crude β (95% CI) | Adjusted ^b β (95% CI) | Crude β (95% CI) | Adjusted ^c β (95% CI) | Crude β (95% CI) | Adjusted ^c β (95% CI) | Crude β (95% CI) | Adjusted ^c β (95% CI) |
| Pre-pregnancy BMI | | | | | | | | |
| Underweight (120) | 0.02 (-0.19, 0.23) | -0.03 (-0.24, 0.18) | -122.1 (-122.3, -121.9)** | -104.2 (-104.4, -104.0)** | -0.15 (-0.36, 0.06) | -0.14 (-0.35, 0.08) | -0.18 (-0.39, 0.02) | -0.16 (-0.37, 0.06) |
| Normal weight (336) | Referent | Referent | Referent | Referent | Referent | Referent | Referent | Referent |
| Overweight/Obese (54) | 0.37 (0.08, 0.66)** | 0.40 (0.11, 0.69)** | 145.7 (145.4, 145.9)** | 144.8 (144.5, 145.1)** | 0.12 (-0.17, 0.41) | 0.09 (-0.21, 0.38) | 0.21 (-0.08, 0.49) | 0.23 (-0.07, 0.52) |
| GWG by the 2009 IOM recommendations ^a | | | | | | | | |
| Inadequate (105) | -0.13 (-0.36, 0.10) | -0.14 (-0.37, 0.10) | -122.6 (-122.9, -122.4)** | -120.5 (-120.8, -120.3)** | 0.07 (-0.17, 0.30) | 0.08 (-0.15, 0.32) | -0.05 (-0.29, 0.18) | -0.07 (-0.31, 0.17) |
| Adequate (222) | Referent | Referent | Referent | Referent | Referent | Referent | Referent | Referent |
| Excessive (183) | 0.26 (0.07, 0.46)** | 0.23 (0.03, 0.43)* | 276.5 (276.3, 276.7)** | 216.3 (216.1, 216.5)** | 0.35 (0.16, 0.55)** | 0.29 (0.09, 0.49)** | 0.42 (0.23, 0.62)** | 0.33 (0.13, 0.52)** |

Table 2. Associations (95% CIs) of pre-pregnancy body mass index (BMI) and gestational weight gain (GWG) with birth outcomes (n = 510). ^aGWG within the IOM recommendations were 12.5–18, 11.5–16, 7–11.5, and 5–9 kg for underweight, normal weight, overweight, and obese women, respectively. ^bPre-pregnancy BMI and GWG were mutually adjusted and also adjusted for maternal age, education, parity, smoking, alcohol use, infant sex, and household monthly salary. ^cPre-pregnancy BMI and GWG were mutually adjusted and also adjusted for maternal age, education, parity, smoking, alcohol use, infant sex, household monthly salary, and gestational age. * $p < 0.05$; ** $p < 0.01$.

| Maternal weight status | Birth outcomes | | | | | | | | |
|--------------------------------------------------|----------------------------------------------------|-------------------------|-----------------------------------|------------------------------------------------------|-----------------------|-----------------------------------|------------------------------------------------------|------------------------|-----------------------------------|
| | High birthweight (Macrosomia) n = 505 with 35 case | | | Small for gestational age (SGA) n = 456 with 32 case | | | Large for gestational age (LGA) n = 478 with 54 case | | |
| | n | Crude OR (95% CI) | Adjusted OR (95% CI) ^b | n | Crude OR (95% CI) | Adjusted OR (95% CI) ^b | n | Crude OR (95% CI) | Adjusted OR (95% CI) ^b |
| Pre-pregnancy BMI | | | | | | | | | |
| Underweight | 118 (3) | 0.31 (0.09, 1.04) | 0.34 (0.10, 1.22) | 115 (8) | 0.88 (0.38, 2.04) | 0.94 (0.40, 2.23) | 112 (5) | 0.31 (0.12, 0.81)* | 0.32 (0.12, 0.86)* |
| Normal weight | 333 (26) | Referent | Referent | 295 (23) | Referent | Referent | 313 (41) | Referent | Referent |
| Overweight/Obese | 54 (6) | 1.48 (0.58, 3.78) | 1.28 (0.44, 3.75) | 46 (1) | 0.26 (0.04, 2.00) | 0.17 (0.02, 1.34) | 53 (8) | 1.18 (0.52, 2.68) | 1.04 (0.42, 2.56) |
| GWG by the 2009 IOM recommendations ^a | | | | | | | | | |
| Inadequate | 103 (6) | 3.36 (0.93, 12.16) | 4.28 (0.95, 17.03) | 98 (15) | 2.54 (1.18, 5.50)* | 3.60 (1.57, 8.30)** | 90 (7) | 1.51 (0.57, 4.03) | 1.76 (0.64, 4.85) |
| Adequate | 221 (4) | Referent | Referent | 211 (14) | Referent | Referent | 208 (11) | Referent | Referent |
| Excessive | 181 (25) | 8.69 (2.97, 25.48)** | 7.49 (2.42, 23.19)** | 147 (3) | 0.29 (0.08, 1.04) | 0.28 (0.08, 1.02) | 180 (36) | 4.48 (2.20, 9.09)** | 3.79 (1.82, 7.90)** |

Table 3. Odds ratios (ORs) for birthweight according to pre-pregnancy body mass index (BMI) and gestational weight gain (GWG). ^aGWG within the IOM recommendations were 12.5–18, 11.5–16, 7–11.5, and 5–9 kg for underweight, normal weight, overweight, and obese women, respectively. ^bPre-pregnancy BMI and GWG were mutually adjusted and also adjusted for maternal age, education, parity, smoking, alcohol use, infant sex, household monthly salary, and gestational age. * $p < 0.05$; ** $p < 0.01$.

Weight status of childbearing-aged women in Shanghai. In the current study, we used the Chinese criteria instead of the WHO standard for BMI classification to categorize the women into different groups. Our preference is explained by the fact that: i) the WHO guidelines for BMI are based on data from Western countries, ii) increased awareness that, by the WHO guidelines, many Asian countries have a low prevalence of obesity, yet high rates of obesity-related diseases. The expert group of WHO recommended that individual countries should set their own criteria for BMI classification on the basis of their own morbidity and mortality data²⁷. Therefore, more sensitive and specific criteria were developed for the Chinese women of childbearing age to estimate the risk of perinatal outcomes²³. As shown in Table 4, we found that the proportion of underweight women in Shanghai (Southern China) (23.5%) was substantially higher than those reported from Tianjin (Northern China) (11.2%)¹⁹ and Western countries such as USA (5.5%)²⁸ and Canada (3.9%)²⁹, but was comparable to those reported from Jiangsu and Zhejiang provinces (nearby Shanghai) (22.7%)²¹ and Wuhan (Central China) (16.9%)³⁰, and other Asian country such as Japan (14.8–20.0%)^{31–33}. On the other hand, the proportion of women with excessive GWG in Shanghai (35.9%) was substantially lower than those reported from other domestic areas such as Tianjin (57.1%)¹⁹ and Wuhan (57.5%)³⁰, and Western countries such as USA (57.9%)²⁸ and Canada (57.7%)²⁹, but higher than the reported proportions from Japan (7.1%–10.9%)^{31,33}.

| Author/year | Country/city | Sample size | Pre-pregnancy BMI (mean \pm SD) | | | GWG (mean \pm SD)* | | |
|--------------------------------------------------------------|--------------------------------------------------------|-------------|-----------------------------------|-------------------|----------------------|----------------------|--------------|---------------|
| | | | Underweight (%) | Normal weight (%) | Overweight/Obese (%) | Inadequate (%) | Adequate (%) | Excessive (%) |
| Yang <i>et al.</i> ⁵ 2011–2013 ³⁰ | Wuhan, China | 85,765 | 20.4 \pm 2.3 kg/m ² | | | 17.4 \pm 7.2 kg | | |
| | | | 16.9% | 76.4% | 6.7% | 17.5% | 25.0% | 57.5% |
| Li <i>et al.</i> ⁵ 2009–2011 ¹⁹ | Tianjin, China | 33,973 | NA | | | NA | | |
| | | | 11.2% | 64.6% | 24.2% | 9.8% | 33.0% | 57.1% |
| Liu <i>et al.</i> [*] 1993–2005 ²¹ | Jiangsu and Zhejiang, China | 273,365 | NA | | | NA | | |
| | | | 22.7% | 75.5% | 1.9% | NA | NA | NA |
| The present study ⁵ 2016 | Shanghai, China | 510 | 20.5 \pm 2.6 kg/m ² | | | 15.1 \pm 4.5 kg | | |
| | | | 23.5% | 65.9% | 10.6% | 20.6% | 43.5% | 35.9% |
| Tanaka <i>et al.</i> [*] 2010–2013 ³¹ | Osaka, Japan | 1883 | 20.7 \pm 2.9 kg/m ² | | | 10.3 \pm 3.7 kg | | |
| | | | 20.0% | 63.9% | 16.1% | 57.9% | 31.2% | 10.9% |
| Murakami <i>et al.</i> ^{*,32} | Sakata, Japan | 633 | 20.9 \pm 2.8 kg/m ² | | | 10.5 \pm 3.4 kg | | |
| | | | 14.8% | 77.2% | 8.0% | NA | NA | NA |
| Enomoto <i>et al.</i> [*] 2013 ³³ | Yokohama, Japan | 97,157 | NA | | | NA | | |
| | | | 18.2% | 71.1% | 10.6% | 63.8% | 29.1% | 7.1% |
| Deierlein <i>et al.</i> [*] 2001–2005 ²⁸ | the Pregnancy Infection and Nutrition (PIN) study, USA | 363 | 24.2 \pm 5.6 kg/m ² | | | 16.0 \pm 5.4 kg | | |
| | | | 5.5% | 65.4% | 29.1% | 12.7% | 29.4% | 57.9% |
| Ferraro <i>et al.</i> [*] 2002–2009 ²⁹ | the Ottawa and Kingston (Oak) Birth Cohort, Canada | 4,321 | 25 \pm 5.6 kg/m ² | | | 16.1 \pm 6.8 kg | | |
| | | | 3.9% | 56.2% | 39.9% | 13.0% | 29.3% | 57.7% |

Table 4. Between-study comparison of pre-pregnancy body mass index (BMI) and gestational weight gain (GWG) in childbearing-aged women. NA, not applicable. *GWG within the IOM recommendations were 12.5–18, 11.5–16, 7–11.5, and 5–9 kg for underweight, normal weight, overweight, and obese women, respectively. ⁵Women were categorized as underweight (<18.5 kg/m²), normal weight (18.5–23.9 kg/m²), overweight (24.0–27.9 kg/m²), or obese (\geq 28.0 kg/m²) according to the standard of Working Group on Obesity in China. ¹⁹Women were categorized as underweight (<18.5 kg/m²), normal weight (18.5–24.9 kg/m²), overweight (25.0–29.9 kg/m²), or obese (\geq 30.0 kg/m²) according to the WHO criteria. ^{*}Women were categorized as underweight (<18.5 kg/m²), normal weight (18.5–22.9 kg/m²), overweight (23.0–24.9 kg/m²), or obese (\geq 25.0 kg/m²) according to the Japan Society for Study of Obesity.

Several factors may explain the relatively thin body image of women observed in this study. Firstly, regional differences in dietary culture exist throughout China, particularly between southern and northern provinces due to climate and agriculture. Traditional Southern diet (e.g., Shanghai) is characterized by high intakes of rice and low intakes of wheat as staple foods. In contrast, Northern China (e.g., Tianjin) cultivates predominantly wheat, which is the staple food of the populations of this region³⁴. Rice is a low-energy food that contains twice the amount of water and half of the energy compared with the same bulk of steamed bread (wheat)³⁵. In addition, the colder climate in Northern China is associated with less physical activity and more energy-rich diets, which may increase body weight. These factors may explain, at least in part, higher prevalence of overweight and obesity in Northern than in Southern China. Secondly, women belonging to ethnic groups characterized by a small body size have been reported to gain less weight on average during pregnancy than larger women such as Caucasians¹⁵. Asian women generally have a lower BMI and a smaller GWG than those reported in North America¹⁶. However, a recent systematic review and meta-analysis found no significant association between dietary diversity and BMI status³⁶. Furthermore, in the context of globalization, the Western fast food diet had become a major dietary pattern in modern China³⁷. Some evidence also shows that the heterogeneity of food intake patterns is less often associated with BMI and obesity in women³⁸. While differences in genetic makeup, environment, and behaviors across populations could explain discrepancies in maternal body weight in obstetric population from different reports, the knowledge regarding the seasonal food intake, dietary and activity patterns and obesity during pregnancy is still limited and require further interrogation.

Association of pre-pregnancy BMI and GWG with birth outcomes. Both pre-pregnancy BMI and GWG are considered to be key determinants of infant health. Although these associations are extensively studied in the literature published from Western countries, much less is known about the association between BMI and GWG and birth outcomes in Chinese populations.

Only few studies examining the associations between maternal pre-pregnancy BMI and GWG and birth outcomes in Asian populations have been conducted, with inconsistent findings. For example, one study conducted in Nagasaki, Japan revealed that GWG below the IOM recommendation was associated with an increased risk of SGA births, while underweight showed no association with SGA births³⁹. In contrast, another Japanese study from Oosaka, found that underweight was associated with an elevated risk of low birthweight infants, but inadequate GWG showed no association with low birthweight³². The same is true for studies of maternal pre-pregnancy BMI. Some studies reported that overweight/obese women had a higher incidence of fetal macrosomia and LGA infants^{16,40}, or overweight/obese women were at increased risk of SGA infants²⁰, while others

reported that overweight/obesity was not associated with either low birthweight infants or fetal macrosomia⁴¹. However, two large Chinese studies conducted in Tianjin (n = 33,973) and Hebei, Jiangsu, and Zhejiang provinces (n = 292,568), respectively, showed that both overweight/obesity and excessive GWG increased the risks of fetal macrosomia and LGA infants, and both underweight and inadequate GWG were the risk factors for low birthweight and SGA infants^{19,21}. A systematic review and meta-analysis indicates that the effect of pre-pregnancy BMI and GWG on delivery outcomes varies significantly among different groups and populations⁴². These differences may be attributed to population-specific genetic determinants, variable definitions used for of BMI and GWG categories, as well as differences in methodologies.

Our study has several limitations. Firstly, pre-pregnancy body weight was self-reported and not measured objectively. Overweight/obese women tend to underestimate their weight; it is possible that reporting bias occurred which may have influenced the grouping on women in the BMIs categories. Secondly, this study was based on data collected from a single hospital located in Shanghai area, which although reflects the characteristics of the pregnant population from this area, may affect the external validity of the findings. Thirdly, because the sample size was relatively small, we could not perform further subgroup analyses of all BMI categories. Finally, since there are no significant seasonal variations in the number of deliveries in our hospital and our population is representative for the obstetric population of the city of Shanghai, the study enrolled all women who delivered in one month of the year. However, we could not appreciate to what extent, if any, the seasonal variability had an impact on our findings. Although a relationship between seasons and the pattern of nutrition and physical activity may exist, the confounding effect of these factors on our results is unknown.

In conclusion, our study shows that association between adverse perinatal outcomes and non-optimal maternal nutrition exists in an urban Chinese population, supporting the findings from the Western populations. Also, our study draw attention to increasing rates of non-optimal maternal nutrition and rising incidence on obesity in Chinese women of childbearing age. Unlike other potential causes of poor birth outcomes, obesity is a preventable factor. Non-optimal GWG and pre-pregnancy BMI may be prevented with increased awareness and appropriate guidance. There is a misconception in the Chinese culture that increased GWG can be beneficial for the offspring's health and growth. Pregnant women, especially those that are already overweight/obese, should be counseled about the importance of appropriate GWG which represents an underutilized opportunity for a public health intervention to prevent adverse perinatal outcomes. Similarly, underweight and overweight/obese women should also be counseled on the risk of poor delivery outcomes and advised on the importance of attaining optimal pre-pregnancy BMI when planning for pregnancy.

References

- Doherty, D. A., Magann, E. F., Francis, J., Morrison, J. C. & Newnham, J. P. Pre-pregnancy body mass index and pregnancy outcomes. *Int. J. Gynaecol. Obstet.* **95**, 242–247 (2006).
- Vinturache, A., Moledina, N., McDonald, S., Slater, D. & Tough, S. Pre-pregnancy Body Mass Index (BMI) and delivery outcomes in a Canadian population. *BMC Pregnancy Childbirth* **14**, 422 (2014).
- Schummers, L., Hutcheon, J. A., Bodnar, L. M., Lieberman, E. & Himes, K. P. Risk of adverse pregnancy outcomes by prepregnancy body mass index: a population-based study to inform prepregnancy weight loss counseling. *Obstet. Gynecol.* **125**, 133–143 (2015).
- Yang, S. *et al.* Parental Body Mass Index, Gestational Weight Gain, and Risk of Macrosomia: a Population-Based Case-Control Study in China. *Paediatr. Perinat. Epidemiol.* **29**, 462–471 (2015).
- Sebire, N. J., Jolly, M., Harris, J., Regan, L. & Robinson, S. Is maternal underweight really a risk factor for adverse pregnancy outcome? A population-based study in London. *BJOG*. **108**, 61–66 (2001).
- Ronnenberg, A. G. *et al.* Low preconception body mass index is associated with birth outcome in a prospective cohort of Chinese women. *J. Nutr.* **133**, 3449–3455 (2003).
- Stotland, N. E., Cheng, Y. W., Hopkins, L. M. & Caughey, A. B. Gestational weight gain and adverse neonatal outcome among term infants. *Obstet. Gynecol.* **108**, 635–643 (2006).
- Thorsdottir, I., Torfadottir, J. E., Birgisdottir, B. E. & Geirsson, R. T. Weight gain in women of normal weight before pregnancy: complications in pregnancy or delivery and birthoutcome. *Obstet. Gynecol.* **99**, 799–806 (2002).
- Olson, C. M., Strawderman, M. S., Hinton, P. S. & Pearson, T. A. Gestational weight gain and postpartum behaviors associated with weight change from early pregnancy to 1 y postpartum. *Int. J. Obes. Relat. Metab. Disord.* **27**, 117–127 (2003).
- Cogswell, M. E., Serdula, M. K., Hungerford, D. W. & Yip, R. Gestational weight gain among average-weight and overweight women—what is excessive? *Am. J. Obstet. Gynecol.* **172**, 705–712 (1995).
- Nohr, E. A. *et al.* Obesity, gestational weight gain and preterm birth: a study within the Danish National Birth Cohort. *Paediatr. Perinat. Epidemiol.* **21**, 5–14 (2007).
- Sullivan, E. A. *et al.* Maternal super-obesity and perinatal outcomes in Australia: a national population-based cohort study. *BMC Pregnancy Childbirth* **15**, 322 (2015).
- Young, O. M., Twedt, R. & Catov, J. M. Pre-pregnancy maternal obesity and the risk of preterm preeclampsia in the American primigravida. *Obesity (Silver Spring)* **24**, 1226–1229 (2016).
- Walsh, J. M., McGowan, C. A., Mahony, R. M., Foley, M. E. & McAuliffe, F. M. Obstetric and metabolic implications of excessive gestational weight gain in pregnancy. *Obesity (Silver Spring)*. **22**, 1594–1600 (2014).
- Ota, E. *et al.* Maternal body mass index and gestational weight gain and their association with perinatal outcomes in VietNam. *Bull World Health Organ.* **89**, 127–136 (2011).
- Yazdani, S., Yosofniyapasha, Y., Nasab, B. H., Mojaveri, M. H. & Bouzari, Z. Effect of maternal body mass index on pregnancy outcome and newborn weight. *BMC Res. Notes.* **5**, 34 (2012).
- Wu, Y. Overweight and obesity in China. *BMJ*. **333**, 362–363 (2006).
- Ma, G. S. *et al.* The prevalence of body overweight and obesity and its changes among Chinese people during 1992 to 2002. *Zhonghua Yu Fang Yi Xue Za Zhi [in Chinese]* **39**, 311–315 (2005).
- Li, N. *et al.* Maternal prepregnancy body mass index and gestational weight gain on pregnancy outcomes. *PLoS One* **8**, e82310 (2013).
- Liu, L., Hong, Z. & Zhang, L. Associations of prepregnancy body mass index and gestational weight gain with pregnancy outcomes in nulliparous women delivering single live babies. *Sci. Rep.* **5**, 12863 (2015).
- Liu, Y., Dai, W., Dai, X. & Li, Z. Prepregnancy body mass index and gestational weight gain with the outcome of pregnancy: a 13-year study of 292,568 cases in China. *Arch. Gynecol. Obstet.* **286**, 905–911 (2012).
- Ding, G. *et al.* Application of a global reference for fetal-weight and birthweight percentiles in predicting infant mortality. *BJOG*. **120**, 1613–1621 (2013).

23. Zhou, B. F. & Cooperative Meta-Analysis Group of the Working Group on Obesity in China. Predictive values of body mass index and waist circumference for risk factors of certain related diseases in Chinese adults—study on optimal cut-off points of body mass index and waist circumference in Chinese adults. *Biomed. Environ. Sci.* **15**, 83–96 (2002).
24. Wang, Y., Mi, J., Shan, X. Y., Wang, Q. J. & Ge, K. Y. Is China facing an obesity epidemic and the consequences? The trends in obesity and chronic disease in China. *Int. J. Obes. (Lond)* **31**, 177–188 (2007).
25. Rasmussen, K. M. & Yaktine, A. L. *Weight Gain During Pregnancy: Reexamining the Guidelines*. Washington, DC, National Academies Press (US), 2009).
26. Zhu, L. *et al.* Chinese neonatal birthweight curve for different gestational age [in Chinese]. *Zhonghua Er Ke Za Zhi* **53**, 97–103 (2015).
27. Choo, V. WHO reassesses appropriate body-mass index for Asian populations. *Lancet* **360**, 235 (2002).
28. Deierlein, A. L., Siega-Riz, A. M., Adair, L. S. & Herring, A. H. Effects of pre-pregnancy body mass index and gestational weight gain on infant anthropometric outcomes. *J. Pediatr.* **158**, 221–226 (2011).
29. Ferraro, Z. M. *et al.* Excessive gestational weight gain predicts large for gestational age neonates independent of maternal body mass index. *J. Matern. Fetal. Neonatal. Med.* **25**, 538–542 (2012).
30. Yang, S. *et al.* Pre-Pregnancy Body Mass Index, Gestational Weight Gain, and Birthweight: A Cohort Study in China. *PLoS One* **10**, e0130101 (2015).
31. Tanaka, T. *et al.* Associations between the pre-pregnancy body mass index and gestational weight gain with pregnancy outcomes in Japanese women. *J. Obstet. Gynaecol. Res.* **40**, 1296–1303 (2014).
32. Murakami, M. *et al.* Prepregnancy body mass index as an important predictor of perinatal outcomes in Japanese. *Arch. Gynecol. Obstet.* **271**, 311–315 (2005).
33. Enomoto, K. *et al.* Pregnancy Outcomes Based on Pre-Pregnancy Body Mass Index in Japanese Women. *PLoS One* **11**, e0157081 (2016).
34. Yu, C. *et al.* Major dietary patterns in relation to general and central obesity among Chinese adults. *Nutrients* **7**, 5834–5849 (2015).
35. Shi, Z. *et al.* Dietary pattern and weight change in a 5-year follow-up among Chinese adults: results from the Jiangsu Nutrition Study. *Br. J. Nutr.* **105**, 1047–1054 (2011).
36. Salehi-Arbargouei, A., Akbari, F., Bellissimo, N. & Azadbakht, L. Dietary diversity score and obesity: a systematic review and meta-analysis of observational studies. *Eur. J. Clin. Nutr.* **70**, 1–9 (2016).
37. Shu, L. *et al.* Association between Dietary Patterns and the Indicators of Obesity among Chinese: A Cross-Sectional Study. *Nutrients* **7**, 7995–8009 (2015).
38. Togo, P., Osler, M., Sørensen, T. I. & Heitmann, B. L. Food intake patterns and body mass index in observational studies. *Int. J. Obes. Relat. Metab. Disord.* **25**, 1741–1751 (2001).
39. Akahoshi, E. *et al.* Association of maternal pre-pregnancy weight, weight gain during pregnancy, and smoking with small-for-gestational-age infants in Japan. *Early Hum. Dev.* **92**, 33–36 (2016).
40. Vinturache, A. E., McDonald, S., Slater, D. & Tough, S. Perinatal outcomes of maternal overweight and obesity in term infants: a population-based cohort study in Canada. *Sci. Rep.* **5**, 9334 (2015).
41. Fouelifack, F. Y. *et al.* Associations of body mass index and gestational weight gain with term pregnancy outcomes in urban Cameroon: a retrospective cohort study in a tertiary hospital. *BMC Res. Notes.* **8**, 806 (2015).
42. McDonald, S. D., Han, Z., Mulla, S. & Beyene, J. Knowledge Synthesis Group. Overweight and obesity in mothers and risk of preterm birth and low birthweight infants: systematic review and meta-analyses. *BMJ.* **341**, c3428 (2010).

Acknowledgements

We thank the Department of Neonatology's staff, students, hospital partners, participants and families, without whom this study would not have been possible. This study was supported by the National Natural Science Foundation of China (Grant no. 81402645).

Author Contributions

Xiaoming Ben conceived the study and designed it with Lingli Xiao and Guodong Ding. Guodong Ding and Lingli Xiao drafted the manuscript. Angela Vinturache provided critical comments and valuable suggestions to content and statistical analysis, and substantially revised the manuscript. Jian Xu, Yifang Ding, Jialin Guo, Liping Huang, Xuelei Yin, Jing Qiao, and Inesh Thureraja participated in the data collection, and contributed to the writing of the manuscript.

Additional Information

Competing financial interests: The authors declare no competing financial interests.

How to cite this article: Xiao, L. *et al.* Associations of maternal pre-pregnancy body mass index and gestational weight gain with birth outcomes in Shanghai, China. *Sci. Rep.* **7**, 41073; doi: 10.1038/srep41073 (2017).

Publisher's note: Springer Nature remains neutral with regard to jurisdictional claims in published maps and institutional affiliations.



This work is licensed under a Creative Commons Attribution 4.0 International License. The images or other third party material in this article are included in the article's Creative Commons license, unless indicated otherwise in the credit line; if the material is not included under the Creative Commons license, users will need to obtain permission from the license holder to reproduce the material. To view a copy of this license, visit <http://creativecommons.org/licenses/by/4.0/>

© The Author(s) 2017