Interaction between delivery mode and maternal age in predicting overweight and obesity in 1,123 Chinese preschool children

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Background: Pediatric overweight/obesity has escalated to epidemic proportions worldwide. In this study, we aimed to assess the association of delivery mode and maternal age, both individually and interactively, with the risk of being overweight or obese among Chinese preschool children.

Methods: We cross-sectionally recruited 1,123 preschool children from five kindergartens in Beijing. Data were collected by a pre-validated self-developed questionnaire. Overweight and obesity are defined according to the World Health Organization (WHO), International Obesity Task Force (IOTF), and China criteria, respectively.

Results: Cesarean delivery was significantly associated with pediatric overweight/obesity under the WHO [adjusted odds ratio (aOR), 95% confidence interval (CI): 1.60, 1.12–2.29], IOTF (1.77, 1.23–2.53), and China (1.43, 1.06–1.94) criteria, respectively. Maternal age <28 years reached statistical significance under both WHO (1.69, 1.09–2.61) and IOTF (1.69, 1.09–2.61) criteria in predicting pediatric overweight/obesity. The interaction between cesarean delivery and maternal age <28 years was remarkably significant under the WHO (2.26, 1.10–4.67), IOTF (2.92, 1.43–5.96), and China (2.36, 1.24–4.50) criteria.

Conclusions: Our findings indicate that the interaction between cesarean delivery and maternal age <28 years can remarkably increase the risk of overweight/obesity among Chinese preschool children.

Keywords: Delivery mode; maternal age; overweight; obesity; preschool children

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Introduction

The past decades have witnessed a dramatic rise in the prevalence of pediatric overweight and obesity around the world (1). In China, an estimated 10% and 12% young children aged 2–6 years are separately classified as overweight and obese (2). The problem of pediatric overweight/obesity has received wide scientific attention,

and we must expand this attention by developing effective strategies to curb this global problem. Hence, efforts to identify potential risk factors attributable to pediatric overweight/obesity must be proceeded.

Growing evidence indicates that cesarean delivery can increase the risk of overweight and obesity in childhood (3-6), whereas other studies failed to reveal any supportive

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evidence (7-9). Li and colleagues meta-analyzed the results of 9 studies, and found that relative to vaginal birth, cesarean delivery was significantly associated with a 33% increased risk of being overweight or obese in offspring (10). Another meta-analysis by Sutharsan and colleagues showed a relative risk of 1.15 for overweight and obesity due to cesarean delivery in early childhood (0 to 5 years), and a similar trend was noted for mid-childhood and adolescence (5 to 18 years) (11). Above evidence collectively indicates that cesarean delivery is a significant risk factor for pediatric overweight and obesity.

Additionally, pregnancy at advanced maternal age is common in both developed and developing countries (12). Some studies have shown that maternal age was closely related to cesarean delivery, and cesarean delivery rate increased consistently with advancing maternal age (12-14). Maternal age seemed to be the strongest predictor of cesarean delivery for the first birth of "low risk" women (15). In a cohort study, Triunfo and colleagues reported that maternal age had a significant and independent contribution to cesarean delivery rate (16). Maternal age not only increased the risk of cesarean delivery, but also associated with pediatric overweight and obesity. Although little research is available on the relationship between maternal age and pediatric obesity, the results of previous studies were inconsistent and inconclusive (17,18). A recent study indicated that the risk of overweight and obesity decreased with increasing maternal age (19). The role of delivery mode and maternal age, acting independently or in an interactive manner, in pediatric overweight and obesity is still unclear.

We here developed a hypothesis that delivery mode and maternal age might act interactively in predisposition to pediatric overweight and obesity. To verify the hypothesis, we examined the association of delivery mode and maternal age, both individually and interactively, with the risk of overweight/obesity among Chinese preschool children using data from a cross-sectional survey.

Methods

Study population

The cross-sectional study recruited preschool children aged 3–6 years from five kindergartens in Chaoyang District, Beijing during the period between March and May in 2017.

This study was approved by the Ethics Committee of China-Japan Friendship Hospital, and the parents or supervisors of all involved children provided written informed consent prior to participation.

Data collection and quality control

Using a stratified cluster random sampling strategy, 1,333 preschool-aged children were selected in this study, and data were collected by a pre-validated self-developed questionnaire. Initially, we collected 1,327 questionnaires, and after excluding questionnaires with invalid gender, body height, and weight, 1,123 validated questionnaires were left in the final analysis. Of all eligible preschool-aged children, 49.8% were boys and 50.2% were girls.

The parents or supervisors of all involved children were requested to complete the self-developed questionnaire to collect data on age of both children and parents, gender, weight, height, and birth weight of children, gestational age, cesarean delivery, feeding patterns, gravidity, parity, and breastfeeding duration, if available. Body weight and height of children and birth history were reported by their parents or supervisors. Gestational age was divided into <37 weeks, 37-42 weeks, and >42 weeks. Delivery mode included cesarean delivery and vaginal delivery. Feeding patterns were classified into breast feeding, mixed feeding, and artificial feeding. Gravidity was defined as a number of pregnancies, while parity was defined as the number of births. The dates of maternal and paternal birth were collected, and their parents' age was calculated as the difference between the child's date of birth and the date of the parents' birth. Maternal age was divided as <28 years, 28-32 years, and >32 years by interquartile range, and paternal age was classified as <30 years, 30-35 years, and >35 years. In cases of children whose mother was dead, relevant data were not recorded and analyzed in this study.

There was strict quality control in process of questionnaires distribution and recovery, as described in our recent publication (20). We provided the same training on investigation methods to health physicians from five kindergartens before distributing questionnaires. The questionnaires were distributed to children and then to their parents or supervisors by kindergarten teachers who were responsible for distributing and collecting the questionnaires. After collecting the questionnaires, health physicians conducted the first check, and we double checked for questionnaires by telephone interview. In addition, questionnaires with missing or unreasonable data were returned to kindergarten teachers, who contacted parents to request or validate the relevant information.

Overweight and obesity definition

In this study, we calculated children's body mass index (BMI) as weight (kg) divided by the square of height (m^2) . Overweight and obesity were defined by BMI using different criteria from the World Health Organization (WHO), the International Obesity Task Force (IOTF), and China, respectively (Table S1). In detail, for the WHO Child Growth Standards, we defined overweight and obesity by age- and sex-specific BMI z-scores and age was considered on account of a difference between over and under 5 years. Under 5 years old, overweight was defined as more than two standard deviations (SD) above the reference median, and obesity was defined as more than three SD (21). Over 5 years old, overweight was greater than one SD above reference median, and obesity was greater than two SD above reference median (22). In the IOTF criteria, we adopted international cut-off points for BMI for overweight and obesity by sex between 2 and 18 years (overweight as over BMI of 25 kg/m²; obesity as over BMI of 30 kg/m² at age 18) (23). With the China criteria, we defined overweight and obesity as over BMI of 24 and 28 kg/m² at age 18 respectively (24).

Statistical analyses

We classified all study children as non-overweight, overweight and obesity groups separately by BMI using the WHO, IOTF, and China criteria. We described categorical variables using count (percentage) and compared between two groups by the use of Chi-squared test. We described continuous variables using the median (interquartile range) comparing among groups by t-test or rank-sum test according to the normality of data distribution (conforming to a normal distribution by t-test, and otherwise by ranksum test). For delivery mode and maternal age, we used the univariate and multivariable logistic regression model to examine the association with overweight and obesity, individually and interactively, before and after adjusting for age, gender, paternal age, gestational age, feeding patterns, gravidity, parity, breastfeeding duration, and birth weight. Additionally, we explored the interaction by conducting subgroup analysis between gender and age before and after other potential confounders.

A restricted cubic spline regression model was fitted to delineate the association between maternal age and overweight/obesity risk among preschool children. To affiliate clinical explanation, we created a three-dimension interactive surface between maternal age and birthweight accounting for the predict risk of overweight and obesity by delivery mode and maternal age.

Three-dimension interactive surface was plotted using the "rgl.surface" program in the "rgl" package, which is a library of functions that offers three-dimensional, realtime visualization functionality to the R programming environment. The core of the "rgl" package is a shared library that acts as an interface between R and OpenGL.

Unless otherwise indicated, the STATA software Release 14.0 (Stata Corp, TX) were applied in statistical analyses. P value of less than 0.05 was considered as statistically significant.

Results

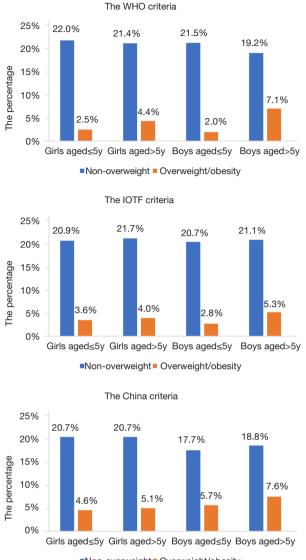
Prevalence of overweight/obesity in preschool children

A total of 1,123 children based on gender and age were divided as four subgroups—girls aged ≤ 5 years, girls aged >5 years, boys aged ≤ 5 years, and boys aged >5 years groups. The prevalence of overweight and obesity in each subgroup under three growth criteria is described in *Figure 1*. High rates of overweight and obesity in a group of boys aged over 5 years were seen (WHO: 7.12%; IOTF: 5.25%; China: 7.57%).

Baseline characteristics

Age, gender, maternal age, paternal age, gestational age, delivery mode, feeding patterns, gravidity, parity, breastfeeding duration, and birth weight were obtained in baseline characteristics (Table 1). Age and gender showed inconsistent difference between the two groups under various growth references. The overweight and obese children born by cesarean delivery was 56.74%, 57.56%, and 54.90% in total overweight and obese children by WHO, IOTF, and China criteria. Compared with vaginal delivery, cesarean delivery rate in overweight and obese group was higher than that in non-overweight group, and reach the statistical difference under three criteria (all P<0.05). Birth weight in overweight and obese group was also high compared with that in non-overweight group, and achieve statistical difference among groups under three criteria (all P<0.05). The overweight and obese children with maternal age <28 years was more than that in nonoverweight children, and showed no statistical significance under both WHO and China criteria, as well as reached

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Non-overweight Overweight/obesity

Figure 1 The prevalence of overweight and obesity in preschool children based on age and gender by WHO, IOTF, and China criteria, respectively. WHO, World Health Organization; IOTF, International Obesity Task Force.

marginal significance under the IOTF criteria (P=0.042). Parental BMI in overweight and obese children were higher than that in non-overweight children, and reached statistical significance under three criteria (all P<0.05).

Risk factors for delivery mode and maternal age

The odds of delivery mode and maternal age, as risk factors for overweight and obesity in childhood, were shown

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in Table 2, and the risk trend of maternal age predicting pediatric overweight and obesity under three criteria (Figure 2). Before and after adjusting for age, gender, paternal age, gestational age, feeding patterns, gravidity, parity, breastfeeding duration and birth weight, cesarean delivery was significant associated with pediatric overweight and obesity under WHO (OR: 1.51 and 1.60, 95% CI: 1.09-2.09 and 1.12-2.29, P=0.012 and 0.010, respectively), IOTF (OR: 1.57 and 1.77, 95% CI: 1.13-2.18 and 1.23-2.53, P: 0.007 and 0.002, respectively), and China (OR: 1.42 and 1.43, 95% CI: 1.07-1.88 and 1.06-1.94, P=0.014 and 0.021, respectively) criteria. Maternal age <28 years showed no statistical difference in increasing the risk of pediatric overweight and obesity by China criteria (OR: 1.40, 95% CI: 0.95-2.05, P=0.086), and reached statistical significance by WHO and IOTF criteria (OR: 1.69, 95% CI: 1.09-2.61, P=0.019). Additionally, the restricted cubic spline regression model indicated that the predict risk for overweight and obesity in childhood was decreased with increase of maternal age, and the trend was moderate until maternal age with 32 years under WHO (Figure 2A), IOTF (Figure 2B), and China (Figure 2C) criteria.

Interaction between delivery mode and maternal age

The effect-size estimates of the interaction between delivery mode and maternal age using three criteria were presented in Table 3. According to categories of delivery mode and maternal age, we recombined six new variables and considered vaginal delivery and maternal age >32 years as a reference to explore their potential interaction. The OR of the single presence of either maternal age <28 years and 28-32 years or cesarean delivery had an increasing trend until the largest effect-size in cesarean delivery and maternal age <28 years by WHO (OR: 2.33, 95% CI: 1.19-4.58), IOTF (OR: 2.61, 95% CI: 1.34-5.08), China (OR: 2.35, 95% CI: 1.29-4.29) criteria. After multivariable adjustment, the preschool children with cesarean delivery and maternal age <28 years was 2.26, 2.92, and 2.36 times for the risk of being overweight and obese under WHO, IOTF, and China criteria, respectively. In view of a higher prevalence of overweight and obesity in boys aged over 5 years, we examined the interaction of delivery mode and maternal age in the risk of overweight and obesity for boys aged over 5 years, and found a rise in odds of overweight and obesity for boys aged over 5 years cesarean delivery with and maternal age <28 years before and after adjustment under WHO (OR: 3.86 and 3.5, 95% CI: 1.20-12.39 and

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Table 1 Baseline characteristics of preschool children

	W	HO criteria	IO [.]	TF criteria		China criteria			
Variables	Non-overweight	Overweight/ obesity	Ρ	Non-overweight	Overweight/ obesity	Р	Non-overweight	Overweight/ obesity	Ρ
Age (years)			<0.001			0.034			0.28
≤5	488 (51.69)	50 (27.93)		467 (49.26)	71 (40.57)		422 (48.79)	116 (44.96)	
>5	456 (48.31)	129 (72.07)		481 (50.74)	104 (59.43)		443 (51.21)	142 (55.04)	
Gender			0.035			0.634			0.004
Male	457 (48.41)	102 (56.98)		469 (49.47)	90 (51.43)		410 (47.40)	149 (57.75)	
Female	487 (51.59)	77 (43.02)		479 (50.53)	85 (48.57)		455 (52.60)	109 (42.25)	
Maternal age (years)		0.054			0.042			0.207
<28	187 (19.81)	49 (27.37)		187 (19.73)	49 (28)		172 (19.88)	64 (24.81)	
28–32	448 (47.46)	82 (45.81)		452 (47.68)	78 (44.57)		411 (47.51)	119 (46.12)	
>32	309 (32.73)	48 (26.82)		309 (32.59)	48 (27.43)		282 (32.6)	75 (29.07)	
Paternal age (y	/ears)		0.954			0.89			0.952
<30	270 (28.6)	53 (29.61)		271 (28.59)	52 (29.71)		247 (28.55)	76 (29.46)	
30–35	411 (43.54)	76 (42.46)		414 (43.67)	73 (41.71)		377 (43.58)	110 (42.64)	
>35	263 (27.86)	50 (27.93)		263 (27.74)	50 (28.57)		241 (27.86)	72 (27.91)	
Gestational ag	e (weeks)		0.132			0.097			0.088
<37	67 (7.17)	146 (82.95)		68 (7.23)	12 (7.06)		60 (6.98)	20 (7.94)	
37–42	815 (87.17)	13 (7.39)		820 (87.14)	141 (82.94)		752 (87.54)	209 (82.94)	
>42	53 (5.67)	17 (9.66)		53 (5.63)	17 (10)		47 (5.47)	23 (9.13)	
Cesarean delivery	436 (46.48)	101 (56.74)	0.012	438 (46.4)	99 (57.56)	0.007	397 (46.11)	140 (54.90)	0.014
Feeding patter	ns		0.905			0.776			0.964
Breast feeding	483 (51.66)	89 (50.28)		485 (51.54)	87 (50.88)		444 (51.63)	128 (50.79)	
Mixed feeding	369 (39.47)	73 (41.24)		371 (39.43)	71 (41.52)		340 (39.53)	102 (40.48)	
Artificial feeding	83 (8.88)	15 (8.47)		85 (9.03)	13 (7.6)		76 (8.84)	22 (8.73)	
Gravidity	1 [1–2]	1 [1–2]	0.703	1 [1–2]	1 [1–2]	0.398	1 [1–2]	1 [1–2]	0.36
Parity	1 [1–1]	1 [1–1]	0.405	1 [1–1]	1 [1–1]	0.722	1 [1–1]	1 [1–1]	0.915
Breastfeeding duration	8 [4–12]	8 [4–12]	0.929	8 [4–12]	8 [4–13]	0.424	8 [4–12]	8 [4–13]	0.402
Birth weight (×500 g)	6.8 (6.06–7.38)	7 (6.4–7.6)	0.009	6.8 (6.06–7.36)	7 (6.4–7.6)	<0.001	6.8 (6.02–7.34)	7 (6.4–7.6)	0.005
Maternal BMI (kg/m²)	20.76 (19.43–22.48)	21.81 (19.94–23.87)	<0.001	20.76 (19.47–22.48)	21.99 (19.92–23.95)	<0.001	20.70 (19.30–22.41)	21.71 (19.96–23.59)	<0.001
Paternal BMI (kg/m²)	24.48 (22.60–26.42)	25.25 (23.40–26.87)	0.003	24.49 (22.60–26.37)	25.62 (23.55–27.47)	<0.001	24.39 (22.60–26.28)	25.47 (23.51–27.40)	<0.001

Data are expressed as median (interquartile range) or count (percent). P values were calculated by the *t*-test or the rank-sum test and the Chisquared test, where appropriate. WHO, World Health Organization; IOTF, International Obesity Task Force.

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		0	1	0					
Variables		WHO criteria			IOTF criteria			China criteria	
variables	OR	95% CI	Р	OR	95% CI	Р	OR	95% CI	Р
Unadjusted									
Delivery mode									
Vaginal delivery		Reference			Reference			Reference	
Cesarean delivery	1.51	1.09-2.09	0.012	1.57	1.13–2.18	0.007	1.42	1.07–1.88	0.014
Maternal age (years)									
>32		Reference			Reference			Reference	
28–32	1.18	0.80–1.73	0.403	1.11	0.75–1.64	0.595	1.09	0.79–1.51	0.610
<28	1.69	1.09-2.61	0.019	1.69	1.09–2.61	0.019	1.40	0.95–2.05	0.086
Multivariable adjusted									
Delivery mode									
Vaginal delivery		Reference			Reference			Reference	
Cesarean delivery	1.60	1.12-2.29	0.010	1.77	1.23–2.53	0.002	1.43	1.06–1.94	0.021
Maternal age (years)									
>32		Reference			Reference			Reference	
28–32	1.27	0.83–1.95	0.275	1.16	0.75–1.79	0.507	1.12	0.78–1.61	0.535
<28	1.72	1.05–2.80	0.030	1.77	1.09–2.87	0.022	1.40	0.92-2.13	0.120

Table 2 The risk of delivery mode and maternal age for pediatric overweight and obesity

P values were calculated before and after adjusting for age, gender, paternal age, gestational age, feeding patterns, gravidity, parity, breastfeeding duration, and birth weight. WHO, World Health Organization; IOTF, International Obesity Task Force; OR, odds ratio; 95% Cl, 95% confidence interval.

1.05–11.60, P: 0.023 and 0.041, respectively), IOTF (OR: 6.78 and 6.08, 95% CI: 1.73–26.59 and 1.49–24.90, P: 0.006 and 0.012, respectively), and China (OR: 3.86 and 3.43, 95% CI: 1.20–12.39 and 1.04–11.31, P: 0.023 and 0.043, respectively) criteria, respectively.

Three-dimension interactive surface

To affiliate clinical explanation, we created a threedimension interactive surface between maternal age and birthweight in *Figure 3*, showing that the younger the mother is, the more weight the child is born with, and the greater the risk of being overweight or obese, and on the contrary the residual age of the mother and the small birthweight of the child were protective factors of overweight and obesity under WHO (*Figure 3A*) and IOTF criteria (*Figure 3B*). The association between maternal age and birthweight was more obvious in China criteria (*Figure 3C*) than the other two criteria.

Discussion

The study aimed to examine the individual and interactive association of delivery mode and maternal age with the risk of overweight and obesity in childhood. The results found that cesarean delivery and maternal age <28 years might be predictors for overweight and obesity, the interaction of which increased at least double risk of overweight and obesity compared with others, especially in boys aged over 5 years. As far as we know, this is the first report on the synergistic effect of delivery mode and maternal age in predicting preschool overweight and obesity. Its important clinical significance lies in the reduction of cesarean delivery rate and providing information of the benefits of ageappropriate birth to help prevent preschool children from being overweight and obesity.

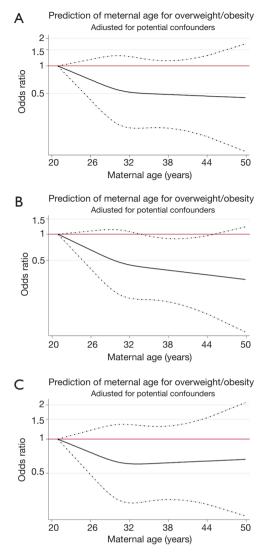


Figure 2 Association between maternal age and overweight/obesity risk among preschool children under the WHO (A), IOTF (B), and China criteria (C), respectively. In panels, navy line was plotted for maternal age, and red line for reference. Dotted lines represent 95% confidence intervals. WHO, World Health Organization; IOTF, International Obesity Task Force.

In this study, the rate of cesarean delivery was nearly 58% in the overweight and obesity group, and higher than that in the non-overweight group. A number of studies have reported that birth by cesarean delivery was closely associated with pediatric overweight and obesity (25-27). The influence of cesarean delivery for offspring overweight and obesity was found in multiple stages of growth and development, such as infancy period, preschool stages, adolescence, and early adulthood (3,6,27,28). Compared

to vaginal delivery, cesarean delivery influenced the newborn acquisition of maternal vaginal microbiota, result in the microbiota dysbiosis (29). An early study showed the primary gut microbiota in infants born by cesarean delivery might be disturbed for up to six months after the birth (30). There was a significant difference in the gut microbiota composition of important bacterial species between obese and lean children (31). Additionally, Gut microbiome transplantation was considered as a protective strategy against obesity and obesity-related diseases while the method was still experimental (32,33). It is possible that intestinal microbiota plays an essential role in the association between delivery mode and pediatric obesity. Besides, there were other underlying mechanisms independent of the intestinal microbiota in the pathophysiology of obesity, such as inflammation and immune factors.

In view of maternal age, we found the rate of overweight and obesity in preschool children decreased with advancing maternal age, which was consistent with a previous study (19). It was a possible mechanism that older mothers were known to give birth to smaller infants, which might be last in later in life. Higher maternal age (OR: 1.11, 95% CI: 1.01–1.22) was one of the strongest predictors of both onset and resolution of overweight/obesity between the primary school children and adolescents (34), which is consistent with our study results.

It is known that cesarean delivery increases the risk of overweight and obesity in childhood. Maternal age was affected by some other factors so that it had a small contribution to predict overweight and obesity later in life. We hypothesized that interaction of delivery mode and maternal age made a larger impact on predicting the risk for pediatric overweight and obesity. Our study results found that the interaction of cesarean delivery and maternal age <28 years increased 1.26–1.92 times in the risk of being overweight and obese under three criteria compared with vaginal delivery and maternal age >32 years, and the risk in boys over 5 years was higher to reach 3.43–6.08 times.

A strength of this study was its use of three different growth criteria estimating overweight and obesity. Our results were adjusted for several variables ignored by some previous studies, such as gestational age, feeding patterns, and breastfeeding duration. Similar to other cross-sectional studies, this study had following limitations. Firstly, there was recall bias for some variables in questionnaires, such as weight, height, and birth history, which would have impact on risk estimate. Secondly, our results were not adjusted for maternal pre-pregnancy BMI, maternal education,

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Table 3 The interaction of delivery mode and maternal age

Variables	_	WHO criteria			IOTF criteria			China criteria		
Variables	OR	95% CI	Р	OR	95% CI	Р	OR	95% CI	Р	
Unadjusted										
Vaginal delivery/maternal age										
>32 years		Reference			Reference			Reference		
28–32 years	0.89	0.49–1.63	0.703	0.83	0.45–1.52	0.524	1.11	0.67–1.85	0.683	
<28 years	1.56	0.82–2.95	0.177	1.41	0.74–2.71	0.299	1.39	0.79–2.45	0.258	
Cesarean delivery/maternal age										
>32 years	1.17	0.63–2.18	0.614	1.17	0.63–2.18	0.614	1.44	0.85-2.44	0.173	
28–32 years	1.76	0.99–3.13	0.052	1.59	0.89–2.84	0.116	1.55	0.93–2.58	0.089	
<28 years	2.33	1.19–4.58	0.014	2.61	1.34–5.08	0.005	2.35	1.29–4.29	0.005	
Multivariable adjusted										
Vaginal delivery/maternal age										
>32 years		Reference			Reference			Reference		
28–32 years	0.89	0.46–1.73	0.741	0.84	0.43–1.64	0.603	1.26	0.73–2.18	0.410	
<28 years	1.57	0.78–3.17	0.210	1.39	0.68–2.84	0.369	1.50	0.81–2.78	0.201	
Cesarean delivery/maternal age										
>32 years	1.16	0.59–2.29	0.673	1.21	0.61–2.40	0.587	1.60	0.90–2.85	0.110	
28–32 years	1.97	1.06–3.66	0.031	1.84	0.99–3.45	0.056	1.70	0.98–2.93	0.057	
<28 years	2.26	1.10–4.67	0.027	2.92	1.43–5.96	0.003	2.36	1.24-4.50	0.009	
Unadjusted in boys aged over 5 years										
Vaginal delivery/maternal age										
>32 years		Reference			Reference			Reference		
28–32 years	2.47	0.84–7.23	0.100	2.60	0.69–9.81	0.158	2.84	0.98-8.27	0.055	
<28 years	3.39	1.05–10.99	0.042	3.85	0.94–15.85	0.062	3.39	1.05–10.99	0.042	
Cesarean delivery/maternal age										
>32 years	2.63	0.82-8.37	0.103	2.17	0.50–9.31	0.299	3.31	1.06–10.37	0.039	
28–32 years	3.54	1.22–10.24	0.020	4.33	1.19–15.84	0.027	3.78	1.31–10.92	0.014	
<28 years	3.86	1.20–12.39	0.023	6.78	1.73–26.59	0.006	3.86	1.20–12.39	0.023	
Multivariable adjusted in boys aged over 5	years									
Vaginal delivery/maternal age										
>32 years		Reference			Reference			Reference		
28–32 years	2.26	0.74–6.92	0.150	1.93	0.49–7.63	0.349	2.68	0.89-8.08	0.080	
<28 years	3.59	1.08–11.96	0.037	3.56	0.85–15.07	0.083	3.61	1.08–12.0	0.036	
Cesarean delivery/maternal age										
>32 years	2.81	0.85–9.26	0.090	2.39	0.54–10.71	0.253	3.60	1.11–11.61	0.032	
28–32 years	3.37	1.13–10.07	0.030	4.06	1.08–15.27	0.038	3.63	1.22–10.81	0.02	
<28 years	3.50	1.05–11.60	0.041	6.08	1.49–24.90	0.012	3.43	1.04–11.31	0.043	

P values were calculated before and after adjusting for age, gender, paternal age, gestational age, feeding patterns, gravidity, parity, breastfeeding duration, and birth weight. WHO, World Health Organization; IOTF, International Obesity Task Force; OR, odds ratio; 95% CI, 95% confidence interval.

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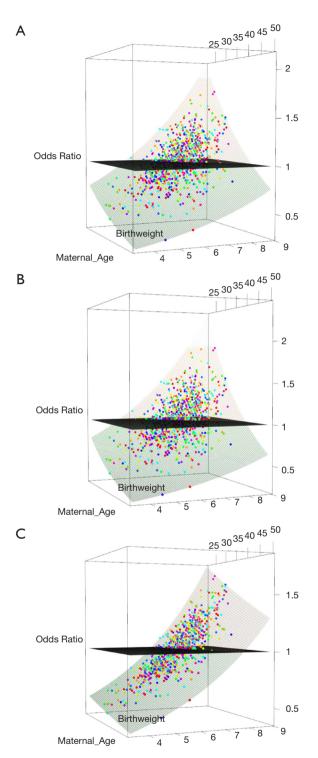


Figure 3 Three-dimension interactive surface of maternal and birthweight in predicting pediatric overweight and obesity under the WHO (A), IOTF (B), China criteria (C), respectively. The grey horizontal grid denotes the reference surface. WHO, World Health Organization; IOTF, International Obesity Task Force.

and antibiotics use during pregnancy due to limited data. Thirdly, cesarean delivery was not divided as planned and unplanned type, although cesarean type was a controversy for the risk of obesity in childhood. A prospective cohort study showed no difference between two types (35), while a study of 8,900 Chinese children found significant difference (3).

Conclusions

Taken together, our findings suggest that the interaction of cesarean delivery and maternal age <28 years, especially in boys over 5 years old, was closely associated with an increased risk of overweight and obesity in Chinese preschool children. We agree that further studies are helpful to call on government and parents to attach importance to control weight in boys over 5 years with maternal age <28 years and cesarean delivery.

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Footnote

Conflicts of Interest: All authors have completed the ICMJE uniform disclosure form (available at http://dx.doi. org/10.21037/atm.2020.03.128). The authors have no conflicts of interest to declare.

Ethical Statement: The authors are accountable for all aspects of the work in ensuring that questions related to the accuracy or integrity of any part of the work are appropriately investigated and resolved. Ethical approval was procured from the ethics committee of China-Japan Friendship Hospital. All child guardians provided written informed consent for the use of data in questionnaires, as well as the subsequent analyses. The children, guardians,

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Supplementary

Table S1 Overweight and obesity definition

WHO criteria (2006)

<5 years

Overweight: BMI z-scores >2 SD

Obesity: BMI z-scores >3 SD

≥5 years

Overweight: BMI z-scores >1 SD

Obesity: BMI z-scores >2 SD

IOTF criteria (2000)

Overweight and obesity are respectively defined to pass through BMI for 25 kg/m² and 30 kg/m² at age 18

China criteria (2009)

Overweight and obesity are respectively defined to pass through BMI for 24 kg/m² and 28 kg/m² at age 18

BMI, body mass index; SD, standard deviation.