



Association between leg length-to-height ratio and metabolic syndrome in Chinese children aged 3 to 6 years

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ABSTRACT

Objective. The aim of this study is to investigate the association between leg-length-to-height ratio (LLHR) and metabolic syndrome (MetS) among Chinese children.

Methods. 1236 children (619 obese and 617 nonobese children) aged 3–6 years participated in a cross-sectional survey in 2005 in Tianjin, China. Information on body adiposity, metabolic traits, and related covariates was obtained using a standardized protocol. LLHR was calculated as the ratio of leg length to stature.

Results. In the multivariable logistic regression analyses, compared with those in the lowest quartile, odds ratios (OR) and 95% confidence intervals (CI) of MetS among children in the second through the highest quartiles of LLHR Z-score were 0.89 (95% CI, 0.64–1.25), 0.45 (95% CI, 0.32–0.63), and 0.37 (95% CI, 0.26–0.53), respectively, (P for trend < 0.0001 across LLHR Z-score quartiles). Compared with children with both higher levels of LLHR and lower levels of adipose indices, the corresponding ORs of MetS for those with both lower levels of LLHR and higher levels of anthropometric indices were 4.51 (95% CI, 3.08–6.62) for BMI Z-score, 3.86 (95% CI, 2.60–5.73) for waist circumference, and 2.75 (95% CI, 1.85–4.10) for waist-to-hip ratio, respectively.

Conclusions. Greater LLHR is inversely associated with MetS in Chinese children.

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Introduction

A growing body of evidence suggests that pediatric obesity and related metabolic abnormalities have profound implications on risk of cardiometabolic disease in adulthood (Gunnell et al., 1998; Guo et al., 2002; Magnussen et al., 2010). Sinha et al. (2002) reported that there is a significant correlation between obesity and metabolic abnormalities in American children and adolescents. A longitudinal study further revealed that pediatric metabolic syndrome (MetS)

was related to severe adult atherosclerosis and an increased risk of type 2 diabetes (T2D) (Magnussen et al., 2010).

Prenatal and postnatal development has been shown to be associated with risk of cardiometabolic disease in adulthood (Barker, 1995). Low birth weight, an indicator of attenuated intrauterine development, has been found to be related to an increased risk of T2D (Kajser et al., 2009) and CVD (Barker et al., 2005). A recent systematic review (Paajanen et al., 2010) reported that shorter stature in adults is associated with increased risks of CVD morbidity and mortality compared with those with relatively greater stature. There is some evidence suggesting an inverse relation of body height to both insulin resistance (Brown et al., 1991; Davey Smith et al., 2001) and risk of T2D (Njolstad et al., 1998; Sayeed et al., 1997). Several studies (Gunnell et al., 1999; Li et al., 2007; Wadsworth et al., 2002) have shown that leg length, but not trunk length, is the height component that is more sensitive to postpartum environmental exposures during infancy. In contrast, birth weight, a proxy of intrauterine development, has an identical impact on both leg and trunk length growth (Gunnell et al., 1999; Wadsworth et al., 2002). Some studies, including ours, have shown that relatively longer leg length

Abbreviations: BMI, body mass index; CI, confidence interval; CVD, cardiovascular disease; HDL, high-density lipoprotein; LDL, low-density lipoprotein; LLHR, leg-length-to-height ratio; MetS, metabolic syndrome; OR, odds ratios; PI, Ponderal index; T2D, type 2 diabetes; WHR, waist-to-hip ratio.

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(Liu et al., 2009) and greater leg length-to-height ratio (LLHR) (Asao et al., 2006) were associated with a reduced risk of T2D in adults. Further, we observed that greater LLHR was associated with a decreased risk of developing childhood overweight or obesity among Canadian children (Liu et al., 2012). We, therefore, hypothesized that height components might be associated with MetS among children. In this analysis, we aimed to investigate the associations between LLHR and MetS in Chinese children aged 3–6 years. In addition, we evaluated the interactions between LLHR and anthropometric indices of adiposity with respect to metabolic abnormalities among our study participants.

Materials and methods

Study participants

The study design and sampling methods of the Tianjin Children's Health Survey 2005 have been described in detail previously (Tian et al., 2010; Zhang et al., 2009). In brief, the cross-sectional survey was carried out in 71 kindergartens randomly sampled from a total of 269 kindergartens in Tianjin, China from March to September 2005. A multistage cluster sampling was used to obtain a random sample of children aged 3 to 6 years in the city. About 20% of kindergartens (29 kindergartens of 151) were randomly selected from 9 urban districts, and about 35% (42 kindergartens of 118) were randomly selected from 9 rural areas because of their small sizes. All children aged 3 to 6 years in the selected kindergartens were invited to participate in the survey. A total of 15,928 children completed the survey, with a response rate of 95.6% (Zhang et al., 2009). Of them, an age-matched (one-for-one in 1 year increment) case-control study was further conducted to investigate neonatal and postnatal factors associated with childhood obesity and obesity-related metabolic abnormalities (Tian et al., 2010). We used the World Health Organization child growth reference (World Health Organization, 2004) to define obesity and nonobesity. The BMI Z-score cutoff point we used was 1.65, which is the 95th percentile of age- and sex-specific distribution. The cases and controls ($n = 1258$) were those with a BMI Z-score of 1.65 or higher and their age-matched counterparts with a BMI Z-score less than 1.65, respectively. The matching was undertaken in each sampled kindergarten. In total, 1236 children (619 obese and 617 nonobese children) were successfully recruited for the study. The study was approved by the institutional review board of the Tianjin Women and Children's Health Center.

Data collection

Trained health workers undertook the data collection procedures. Before the physical examination, a set of self-administered questionnaires was given to the children's parents to be completed at home (Tian et al., 2010). Parents were also instructed not to give any food or beverages to their child after 8 PM the night prior to the morning physical examination. The physical examination was done between 7:30 and 8:30 AM at the kindergarten clinic unit. When children were brought to clinic by their parents, the self-administered questionnaire was examined by a trained health worker, who was from the Women and Children's Health Center at city or local district levels, to identify any uncertain and incomplete questions from the questionnaire and seek clarification from the parents. After anthropometry and other measurements were taken approximately 0.5-mL sample of peripheral blood was taken from the child's middle finger.

Anthropometric measurements

During the physical examination, weight was measured with a beam-balance scale with subjects wearing light indoor clothing without shoes. Height and sitting height were measured by a stadiometer. Sitting height was measured as the child sitting on the seat straight against the wall of the stadiometer and recorded as the distance between the

head piece that was touching the child's head firmly and the seat of the stadiometer. Waist circumference was measured at the level of the umbilicus and hip circumference was measured at the widest point around the left and right greater trochanters. The values of height, sitting height, waist circumference and hip circumference were recorded to 0.1 cm and weight was recorded to 0.1 kg.

Body mass index (BMI) was calculated as weight in kilograms divided by height in meters squared. Ponderal index (PI) was calculated as birth weight in kilograms divided by birth length in meters cubed. Waist-to-hip ratio (WHR) was calculated by dividing waist circumference by hip circumference. Leg-length was determined as the difference between body height and sitting height (Bogin and Varela-Silva, 2010). LLHR was computed as the ratio of leg length to body height: $(\text{leg-length} / \text{height}) \times 100$.

Measurements of metabolic traits

Blood pressure was measured using a standardized mercury sphygmomanometer with an appropriate cuff bladder for children. The fourth Korotkoff sound was adopted for diastolic blood pressure recording. The measurement was taken on the right arm of the participant in a comfortable sitting position after at least 5 minutes' rest. Mean blood pressure was calculated from 2 readings unless the difference between these readings was greater than 10 mm Hg, in which case a third measurement was taken and the mean of the last 2 measurements was used.

Serum fasting glucose, total cholesterol, high-density lipoprotein (HDL)-cholesterol, low-density lipoprotein (LDL)-cholesterol, and triglycerides were measured on an automatic analyzer (RX Daytona; Randox Laboratories Ltd, Antrim, Ireland) with reagents purchased from the manufacturer.

Definition of metabolic syndrome

In this analysis, the pediatric MetS was defined as having three or more of the following components: 1) waist circumference ≥ 75 percentile of age- and sex-specific waist circumference distribution (Fernandez et al., 2004); 2) triglycerides ≥ 0.85 mmol/L; 3) HDL cholesterol < 1.55 mmol/L (Jolliffe and Janssen, 2006); 4) either systolic or diastolic blood pressure ≥ 75 percentile age-, sex-, and height-specific blood pressure distribution (National High Blood Pressure Education Program Working Group on High Blood Pressure in Children and Adolescents, 2004); and 5) fasting glucose ≥ 5.6 mmol/L (Magnussen et al., 2010).

Assessment of covariates

Parents' educational attainment was categorized into three groups: 9 years or less, 10 to 12 years, and 13 years or more. Both mother's and father's BMI were derived from information of weight and height collected in the self-administered questionnaire. Based on the responses to relevant questions from the questionnaire, the following categorical variables were created and dichotomized as yes or no: breast-feeding at age of 12 months, complementary food introduction before age of 6 months, sleep duration less than 9 h, sweetened beverage consumption more than 500 ml/week, high-fat meat intake defined as if fat content was more than half of total meat, everyday intake of vegetables and fruits, duration of television viewing 60 minutes/day or more, and duration of any type of physical activity 30 min/day or less. Disease status was classified as yes for 13 participants who reported having pneumonia, cold, or fever during the past 30 days (Tian et al., 2010).

Statistical analysis

In order to account for age and sex differences in LLHR, we computed age- and sex-specific standard deviation scores (Z-score) of LLHR for

each child. Differences in the variables across LLHR Z-score quartiles were assessed by either a general linear regression model for continuous variables or the *Chi*-square test for categorical variables. Multivariable logistic regression models were utilized to assess the associations of LLHR with the prevalence of MetS. The analyses were performed with adjustment for age and sex (Model one) and then further controlling for other potential confounders (Model two) that we have previously reported to be associated with obesity and metabolic disorders in Chinese children (Bowers et al., 2011; Tian et al., 2010; Yu et al., 2008; Zhang et al., 2009). To examine whether the association between LLRH and MetS was affected by different measurements used to describe adiposity, i.e., BMI Z-score, waist circumference, and WHR, we redefined modified MetS as having two or more MetS components excluding enlarged waist circumference. Levels of LLHR were classified as low and high by the median of LLHR Z-score and the anthropometric indices were grouped into tertiles. In addition, we evaluated the associations of LLHR Z-score with MetS components by using multivariable linear regression models. When appropriate, natural log-transformation was performed to improve the normality of data on MetS components. The statistical significance was defined as $P < 0.05$ (2-sided). All statistical analyses were performed with SAS version 9.2 (SAS Institute, Cary, North Carolina).

Results

General characteristics of study participants

The general characteristics of participants by quartiles of LLHR Z-score are listed in Table 1. In general, compared with children with lower levels of LLHR, those with higher levels of LLHR were more likely to be girls and nonobesity and consume high fat meat. A higher percentage reported high levels of mother's educational attainment. Meanwhile, they had lower levels of body weight, BMI, waist and hip circumference, WHR, blood pressure, triglycerides, and mother's BMI and higher levels of body height, leg length, and HDL-cholesterol.

Associations of leg length-to-height ratio with metabolic syndrome

When adjusting for age and sex, results from logistic regression analyses indicated that greater LLHR was associated with decreased odds of having MetS in children aged 3 to 6 years ($P_{\text{for trend}} < 0.0001$ across LLHR Z-score quartiles, Table 2). In the multivariable adjusted model, the odds ratios (OR, 95% confidence interval [CI]) of having MetS for children in the second through top quartiles were 0.89 (95% CI, 0.64–1.25), 0.45 (95% CI, 0.32–0.63), and 0.37 (95% CI,

Table 1
General characteristics of study participants according to quartiles of leg length-to-height ratio Z-score^a.

Characteristics	Quartiles of age- and sex-specific leg length-to-height ratio Z-score				P
	Q1 (lowest) (n = 309)	Q2 (n = 309)	Q3 (n = 309)	Q4 (highest) (n = 309)	
Age (yr)	5.2 (0.9)	5.3 (0.9)	5.3 (0.8)	5.4 (0.9)	0.22
Girls (n, %)	114 (36.9)	152 (49.2)	125 (40.5)	135 (43.7)	0.016
Obesity (n, %)	207 (67.0)	178 (57.6)	143 (46.3)	91 (29.5)	<0.0001
Birth weight (g)	3530 (464)	3552 (567)	3550 (625)	3503 (446)	0.63
Birth length (cm)	54.4 (12.3)	53.5 (10.9)	52.9 (10.1)	54.2 (11.5)	0.32
Ponderal Index (kg/m ³)	25.2 (7.7)	25.5 (6.3)	26.3 (7.6)	24.8 (6.9)	0.071
Gestational age (weeks)	39.2 (1.2)	39.5 (3.6)	39.3 (3.7)	39.5 (3.6)	0.63
Body weight (kg)	25.7 (6.4)	25.2 (6.0)	24.2 (5.8)	22.9 (4.9)	<0.0001
Body height (cm)	114.7 (7.3)	115.9 (7.1)	116.3 (6.9)	116.6 (6.8)	0.004
Leg length (cm)	48.5 (3.8)	50.4 (3.6)	51.4 (3.5)	52.7 (3.6)	<0.0001
Body mass index (kg/m ²)	19.3 (3.3)	18.5 (3.0)	17.7 (2.9)	16.7 (2.4)	<0.0001
Waist circumference (cm)	61.4 (8.5)	60.9 (8.2)	59.2 (8.0)	57.6 (7.0)	<0.0001
Hip circumference (cm)	66.2 (7.7)	65.6 (7.2)	64.2 (7.0)	62.6 (6.3)	<0.0001
Waist-to-hip ratio	0.93 (0.04)	0.93 (0.05)	0.92 (0.04)	0.92 (0.04)	0.025
Systolic blood pressure (mm Hg)	99.2 (11.1)	98.0 (10.6)	96.5 (9.9)	94.6 (9.3)	<0.0001
Diastolic blood pressure (mm Hg)	63.8 (8.2)	63.7 (8.3)	62.6 (7.6)	61.4 (6.7)	0.0003
Glucose (mmol/L)	5.02 (0.47)	4.97 (0.58)	5.00 (0.51)	4.95 (0.57)	0.38
Total cholesterol (mmol/L)	4.01 (0.64)	4.07 (0.64)	4.05 (0.66)	4.14 (0.69)	0.10
HDL cholesterol (mmol/L)	1.47 (0.30)	1.47 (0.27)	1.52 (0.34)	1.52 (0.31)	0.022
Triglycerides (mmol/L)	0.91 (0.47)	0.88 (0.38)	0.84 (0.43)	0.82 (0.36)	0.046
LDL cholesterol (mmol/L)	2.33 (0.58)	2.38 (0.56)	2.32 (0.52)	2.38 (0.61)	0.38
Mothers' body mass index (kg/m ²)	22.8 (3.3)	22.7 (3.7)	22.5 (2.9)	22.1 (3.0)	0.033
Fathers' body mass index (kg/m ²)	24.9 (4.0)	25.1 (4.3)	25.0 (3.6)	24.5 (3.9)	0.25
Mothers' education (yr, n, %)					0.033
≤9	86 (27.8)	94 (25.2)	78 (25.2)	69 (22.3)	
10–12	108 (35.0)	80 (28.7)	86 (27.8)	93 (30.1)	
≥13	115 (37.2)	135 (43.9)	145 (46.9)	147 (47.6)	
Fathers' education (yr, n, %)					0.84
≤9	80 (25.9)	73 (23.6)	70 (22.7)	66 (21.4)	
10–12	92 (29.8)	98 (31.7)	92 (29.8)	92 (29.8)	
≥13	137 (44.3)	138 (44.7)	147 (47.6)	157 (48.9)	
Breast feeding cessation <12 months (yes, n, %)	107 (34.6)	106 (34.3)	117 (37.9)	115 (37.2)	0.73
Age at complementary food introduction <6 months (yes, n, %)	164 (53.6)	163 (53.6)	172 (56.0)	164 (53.6)	0.91
Sleep duration <9 h (yes, n, %)	125 (40.5)	129 (41.8)	137 (44.3)	127 (41.0)	0.78
Disease during past month ^b (yes, n, %)	5 (1.6)	4 (1.3)	3 (1.0)	1 (0.3)	0.51
Sweetened beverage drinking >500 ml/week (yes, n, %)	101 (32.7)	104 (33.7)	120 (38.8)	104 (33.7)	0.36
Vegetable intake everyday (yes, n, %)	138 (44.7)	127 (41.1)	136 (44.0)	133 (43.0)	0.82
Fruit intake everyday (yes, n, %)	149 (48.2)	123 (39.8)	137 (44.3)	131 (42.4)	0.19
High fat meat intake (yes, n, %)	174 (56.3)	203 (65.7)	207 (67.0)	210 (68.0)	0.009
Duration of television viewing ≥60 min/day (yes, n, %)	147 (47.6)	140 (45.3)	132 (42.7)	120 (38.8)	0.15
Duration of physical activity <30 min/day (yes, n, %)	70 (22.7)	67 (21.7)	83 (26.9)	86 (27.8)	0.20

^a Data are mean (standard deviation) and number of participants (percentage); percentages do not total 100 due to rounding.

^b Parent-reported pneumonia, cold or fever during past month.

Table 2

Associations between leg length-to-height ratio Z-score and metabolic syndrome in children aged 3–6 years in Tianjin, China in 2005.

	Metabolic syndrome (ORs, 95% CIs)				P for trend	P for interaction	Continuous ^c
	Q1	Q2	Q3	Q4			
<i>Entire samples</i>							
Cases/participants	153/309	140/309	96/309	78/309			
Model 1 ^a	1.0 (Reference)	0.87 (0.63–1.20)	0.47 (0.34–0.65)	0.35 (0.25–0.50)	<0.0001		0.66 (0.58–0.75)
Model 2 ^b	1.0 (Reference)	0.89 (0.64–1.25)	0.45 (0.32–0.63)	0.37 (0.26–0.53)	<0.0001		0.67 (0.59–0.76)
<i>By gender</i>							
<i>Boys</i>							
Cases/participants	95/195	75/157	59/184	43/174			
Model 1 ^a	1.0 (Reference)	0.90 (0.59–1.38)	0.46 (0.30–0.70)	0.33 (0.21–0.51)	<0.0001	0.89	0.64 (0.54–0.76)
Model 2 ^b	1.0 (Reference)	0.88 (0.56–1.37)	0.44 (0.29–0.69)	0.34 (0.21–0.54)	<0.0001	0.91	0.65 (0.55–0.77)
<i>Girls</i>							
Cases/participants	54/114	65/152	37/125	35/135			
Model 1 ^a	1.0 (Reference)	0.85 (0.52–1.38)	0.48 (0.28–0.82)	0.40 (0.23–0.68)	<0.0001		0.69 (0.57–0.83)
Model 2 ^b	1.0 (Reference)	0.92 (0.55–1.57)	0.43 (0.24–0.76)	0.41 (0.23–0.72)	0.0001		0.68 (0.55–0.83)
<i>By age</i>							
<i>3–4 years</i>							
Cases/participants	61/110	52/100	32/94	30/104			
Model 1 ^a	1.0 (Reference)	0.90 (0.52–1.56)	0.42 (0.24–0.74)	0.34 (0.19–0.61)	<0.0001	0.94	0.60 (0.48–0.75)
Model 2 ^b	1.0 (Reference)	0.93 (0.52–1.68)	0.40 (0.22–0.74)	0.36 (0.20–0.66)	<0.0001	0.98	0.62 (0.49–0.78)
<i>5–6 years</i>							
Cases/participants	92/199	88/209	64/215	48/205			
Model 1 ^a	1.0 (Reference)	0.86 (0.58–1.28)	0.50 (0.33–0.74)	0.36 (0.24–0.55)	<0.0001		0.69 (0.59–0.81)
Model 2 ^b	1.0 (Reference)	0.86 (0.57–1.30)	0.46 (0.30–0.71)	0.37 (0.24–0.59)	<0.0001		0.69 (0.59–0.81)

CI = confidence interval; OR = odds ratio.

^a Model 1, adjusted for age and sex (when appropriate).^b Model 2, adjusted for age, sex (when appropriate), Ponderal index, gestational age, parents' educational status and body mass index, sleep duration, timing of breast feeding cessation and complementary food introduction, self-reported disease during past month, sweetened beverage drinking, intakes of vegetable, fruit and high fat meat, duration of television viewing, and physical activity.^c The ORs were calculated including leg length-to-height ratio Z-score as a continuous variable in the logistic regression models.

0.26–0.53), respectively, in comparing with those in the first quartile ($P_{\text{for trend}} < 0.0001$). Each a 1-standard deviation (Z-score) increment in LLHR was associated with about 30% of decreased likelihoods of MetS. These associations were consistent among different genders and age groups.

We observed an additive effect of lower levels of LLHR and higher levels of adiposity on the risk of the modified MetS (Fig. 1, A to C). Compared to children with both higher levels of LLHR and lower levels of adipose indicators, the corresponding odds (95% CI) of having the modified MetS for those with both lower levels of LLHR and higher levels of anthropometric indices were 4.51 (95% CI, 3.08–6.62) for BMI Z-score, 3.86 (95% CI, 2.60–5.73) for waist circumference, and 2.75 (95% CI, 1.85–4.10) for WHR, respectively.

For MetS components, LLHR Z-score was inversely and significantly associated with waist circumference, blood pressure, and triglycerides (Supplementary Table 1). These associations were more consistent among boys than girls.

Discussion

In this large-scale cross-sectional analysis, we found that the ratio of leg length to body stature was inversely associated with MetS in Chinese children as young as three to six years of age. This association appeared to be independent of a number of both parental and postnatal factors known to be related to obesity and metabolic disorders in Chinese children (Bowers et al., 2011; Tian et al., 2010; Yu et al., 2008; Zhang et al., 2009). Importantly, we observed an additive effect of shorter leg length and all dimensions of adiposity on MetS risk, suggesting that LLHR might be a marker for pathogenesis of cardiometabolic disorders in children (Grundy et al., 2005; Savva et al., 2000).

Many studies have repeatedly observed stature components in adults, particularly leg length, to be associated with cardiometabolic disorders, including obesity (Gunnell et al., 2003; Lawlor et al., 2004), insulin resistance (Asao et al., 2006; Davey Smith et al., 2001; Lawlor et al., 2004), dyslipidemia (Gunnell et al., 2003; Han et al., 1997;

Lawlor et al., 2002), elevated blood pressure (Langenberg et al., 2005), T2D (Asao et al., 2006; Lawlor et al., 2002; Liu et al., 2009), and CVD (Davey Smith et al., 2001; Ferrie et al., 2006; Lawlor et al., 2004). Although the underlying mechanism of the stature–disease risk association is not fully understood, it is generally believed that it reflects an impaired nutrition status in early life (Leitch, 1951), which may modify the metabolic pathway and influence risk of disease development. Recently, some studies reported that LLHR was negatively associated with the risk of childhood overweight and obesity (Frisancho, 2007; Liu et al., 2012; Pliakas and McCarthy, 2009). To our knowledge, this is the first analysis investigating the association of relative leg length with MetS in relation to obesity status—the known cardiometabolic risk factors in children.

It is commonly believed that the relatively shorter stature or leg length may indicate an adverse environmental exposure before prepubescence, but the actual mechanism is unclear. Several hypotheses have been proposed for the possible explanations of this observed stature–disease relationship (Bogin et al., 2007). It is pointed out that human growth is highly sensitive during the prepubertal growth and development in responding to the overall quality of living conditions and the legs are growing faster relative to other body parts from birth to age 7 years (Bogin and Varela-Silva, 2010). A relatively shorter leg length may indicate the occurrence of adversity during infancy and childhood leading to competition between body parts. The vital organs of head, thorax, and abdomen of the body will be protected from any adversity at the expense of the less vital tissues including the limbs. Therefore, the cost of the competition may not just lead to shorter limbs, but may also change the metabolic patterns that may influence the risk of developing cardiometabolic disease (Varela-Silva et al., 2007). This observation has been supported by data from several experimental studies (Ferron et al., 2010; Fulzele et al., 2010) suggesting that some proteins, (e.g., osteocalcin) produced during bone turn over might have endocrine function that were involved in the regulation of glucose metabolism and might play an important role in the development of T2D.

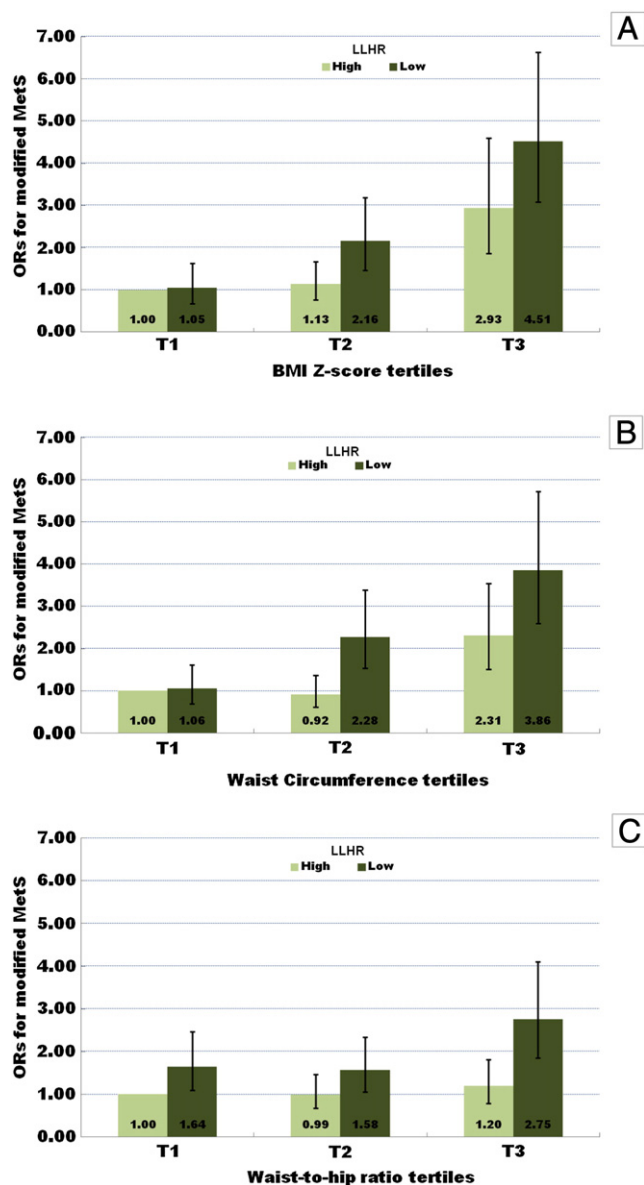


Fig. 1. Joint associations of leg length-to-height ratio and body compositions with modified metabolic syndrome in children 3–6 years of age in Tianjin, China in 2005. A, B, and C: Multivariable adjusted ORs (95% CIs) of having the modified metabolic syndrome according to median of leg length to height ratio Z-score and tertiles of BMI Z-score (A), waist circumference (B), and waist-to-hip ratio (C). The modified metabolic syndrome was defined as having more than two of the four metabolic syndrome components, i.e., elevated blood pressure, hyperglycemia, low HDL cholesterol, and hypertriglyceridemia. The Odds ratios were computed with adjustment for age, sex, Ponderal index, gestational age, parents' educational status and body mass index, sleep duration, timing of breast feeding cessation and complementary food introduction, self-reported disease during past month, sweetened beverage drinking, intakes of vegetable, fruit and high fat meat, duration of television viewing, and physical activity.

Further, our findings are in accordance with a recent study reporting that the associations of smaller LLHR with worse insulin sensitivity and β cell function were more evident among adults with relatively higher waist circumference compared with those with lower waist circumference (Johnston et al., 2013). We also observed that children with higher levels of LLHR were more likely to have mothers with high levels of education compared with those with lower levels. Substantial evidence suggests that childhood environmental conditions are strong determinants of leg and trunk length in adults (Li et al., 2007; Wadsworth et al., 2002). Parental socioeconomic status (SES) may be a proxy of childhood environmental exposure. Further studies are warranted to

investigate factors related to parental SES that may influence children's body growth.

Our study has some limitations. Firstly, the cross-sectional design of the study did not allow us to make inferences based on temporal sequence. Secondly, there is no consensus on the definition of pediatric MetS (Paul et al., 2007). The cut-offs adopted in our study could only indicate elevated levels of these measurements. Further studies are warranted to evaluate the long-term health effects of childhood clustering of multiple metabolic abnormalities. Thirdly, the assessment of covariables was based on the parents' responses to questionnaires. This might be subject to recall bias, which might lead to underestimation of the observed associations. Finally, our study participants were from a case-control study that may limit the generalizability of our findings. Notwithstanding these limitations, the population-based larger sample size and actual anthropometric and laboratory measurements were the strengths of the study.

Conclusions

We find that LLHR is inversely associated with MetS in Chinese children as young as 3 to 6 years of age. LLHR and adiposity may have an additive effect in relation to pediatric MetS. These findings may suggest that further investigations are warranted to delineate the biological pathways underlying the observed association and to identify appropriate means toward enhancing prepubertal LLHR for reducing individual risk of developing cardiometabolic disease.

Supplementary data to this article can be found online at <http://dx.doi.org/10.1016/j.pmedr.2014.11.002>.

Disclosure

The authors have full access to the data in this study and take complete responsibility for the integrity of the data and the accuracy of the data analysis.

The sponsors were not involved in the study design, data collection, analysis, or interpretation.

Conflict of interest statement

The authors have no conflicts of interest to declare.

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