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Validity of Body Mass Index as a Measure of Adiposity in Infancy

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Abstract

Objectives—To assess the validity of BMI and age- and sex-standardized BMI z-score (BMIZ) as surrogates for adiposity [body fat percentage (BF%), fat mass, and fat mass index (FMI, kg/ $(m²)$] at three time points in infancy (1, 4, and 7 months), and to assess the extent to which change in BMIZ represents change in adiposity.

Study design—We performed a secondary analysis of 447 full-term infants in a previous trial of maternal vitamin D supplementation during lactation. Study staff measured infant anthropometrics and assessed body composition with dual-energy x-ray absorptiometry at 1, 4, and 7 months. We calculated Spearman correlations (r_s) among BMI, BMIZ, and adiposity at each time point, and between change in BMIZ and change in adiposity between time points.

Results—Infants (N=447) were 52% male; 37% white, 31% black, 30% Hispanic. BMIZ was moderately correlated with BF% $(r_s=0.43, 0.55, 0.48$ at 1, 4, and 7 months respectively). BMIZ correlated more strongly with fat mass and FMI, particularly at 4 and 7 months (fat mass $r_s = 0.72-$

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0.76; FMI r_s =0.75–0.79). Changes in BMIZ were moderately correlated with adiposity changes from 1–4 months (r_s =0.44 with BF% change; r_s =0.53 with fat mass change), but only weakly correlated from 4–7 months $(r_s=0.21 \text{ with BF\% change}; r_s=0.27 \text{ with fat mass change}).$

Conclusions—BMIZ is moderately correlated with adiposity in infancy. Changes in BMIZ are a poor indicator of adiposity changes in later infancy. BMI and BMIZ are limited as surrogates for adiposity and especially adiposity changes in infancy.

Keywords

body composition; infant

Anthropometric measures of body proportionality are widely used in research and clinical practice as surrogates for body composition. In particular, body mass index (BMI, kg/m²) is often used as a proxy for adiposity, based on high correlations (correlation coefficients of 0.79–0.92) between BMI and directly measured adiposity in adults (1), adolescents (2,3), and school-aged children (2–5). In infants, BMI and other indices of body proportionality such as weight-for-length (WFL, kg) have similarly been interpreted as surrogates for adiposity, with higher BMI or WFL measurements interpreted as reflecting higher adiposity $(6-11)$.

BMI in infancy is predictive of subsequent obesity, with higher peak BMI, rapid increases in BMI, and BMI z-score (BMIZ) values above the 85th percentile all associated with obesity in early childhood or adulthood (10,12–15). However, the extent to which BMI, BMIZ, and their changes over time accurately reflect adiposity gains in infancy is poorly understood.

Although a few prior studies have examined associations between BMI or BMIZ and adiposity in infancy, most were limited by small sample sizes (<70 infants) (18–20), limitation to a single ethnic group (21), or methods of body composition assessment (such as isotope dilution or total body electrical conductivity) (18–20,22,23) that are potentially less accurate than newer methods such as dual-energy x-ray absorptiometry (DXA) (24,25). One recent study, using air displacement plethysmography, overcomes most of those limitations, but only assessed infants at 2 time points and only followed infants to 5 months (27). Therefore, we analyzed the associations of BMI and BMIZ, and their changes over time, with adiposity in a large, racially diverse cohort of healthy infants during the first 7 months of life.

METHODS

This study was conducted as a secondary analysis of data collected during a previously completed study of maternal vitamin D supplementation (28). Mother-infant dyads were recruited from the newborn nursery at the Medical University of South Carolina in Charleston, South Carolina or Rochester General Hospital in Rochester, New York, as well as from local community hospitals in Charleston and Rochester, between January 2007 and December 2011. Eligibility criteria included healthy singleton infant born at ≥35 weeks gestational age, age of infant at enrollment <6 weeks, and mother planning to exclusively breastfeed or exclusively formula feed for 6 months. Mothers were excluded if they had pre-

existing type 1 or 2 diabetes mellitus, hypertension, parathyroid disease, uncontrolled thyroid disease, hypocalcemia, hypercalcemia, or were taking diuretics or cardiac medications. Infants were excluded if they had an inborn error of metabolism, congenital anomalies, or admission to the neonatal intensive care unit lasting >72 hrs. Eligibility criteria for this secondary analysis included any infant with at least one visit with both anthropometric data and body composition measurement. Of the 460 infants enrolled in the original study, 447 (97%) were included in the analysis; 13 were excluded because they had no DXA measurements performed.

Birthweight was obtained from the infants' medical records. Subsequent weights and lengths were measured monthly by trained study staff. At each study visit, infants were weighed on a standard infant scale (Scale-Tronix, Inc, Wheaton, IL) to the nearest gram and length was measured to the nearest 0.1 cm on either a vinyl or plexiglass infant length board (Perspective Enterprises, Portage, MI) by the two-person technique. BMI was calculated using the standard formula: weight/length² (kg/m²). Birthweight z-scores were determined from the 2010 Olsen growth charts accounting for gestational age and sex (29). Z-scores were computed for weight, length, BMI, and WFL at 1, 4, and 7 months using the World Health Organization growth standards via the macro 'WHO Child Growth Standards SPSS Syntax File (igrowup.sps)' (World Health Organization, Geneva, Switzerland, 2005).

At 1, 4, and 7 months, infants underwent whole body dual-energy radiograph absorptiometry using a Hologic Discovery A (Hologic, Inc, Waltham, MA) with the Hologic infant wholebody software (version 12.7.3:3). Both study sites used a Hologic Discovery A and a spine phantom standard was sent to each site twice during the study period for cross-calibration of the scanners; 20 scans were performed at each site on the phantom, with correlation 0.998 and no statistically significant differences in body composition of the phantom as measured by the two different scanners. A standard procedure was followed to limit infant movement and only scans without motion artifact were used. DXA is safe (radiation exposure equivalent to about 1 day's exposure to background radiation), as accurate as more sophisticated methods for measuring infant body composition such as magnetic resonance imaging (30), and is superior to anthropometric measures such as skinfold thicknesses (25).

We obtained 2 measures of adiposity—body fat percentage (BF%) and fat mass (in kg) from each scan. We then calculated a third measure of adiposity, fat mass index (FMI), from the standard formula: fat mass/length² (kg/m²), using the length measured by study staff. BF % is the most commonly used measure of adiposity in research (31), but has been criticized as a poor measure of adiposity because it is affected by changes in lean mass even if fat mass does not change (31,32). Measuring fat mass alone does not account for proportionality of the fat mass for body size, whereas the FMI adjusts for the patient's height or length (31,32). It is unknown which measure of adiposity is most predictive of subsequent health outcomes; therefore, we examined all three measures. Lean mass (in kg) also was obtained directly from each scan and used to calculate the lean mass index (LMI) using the formula: lean mass/length² (kg/m²).

The trial from which these data were obtained was approved by the Institutional Review Boards of the Medical University of South Carolina (#16536) and the University of

Rochester (#14460) and was registered via clinicaltrials.gov (NCT00412074). Parents of infants who participated in the study provided informed consent. This secondary analysis of de-identified data was classified as exempt by the Institutional Review Board of Brigham and Women's Hospital.

Statistical Analyses

Using the DXA-derived adiposity measures as the gold standard for adiposity, we calculated correlation coefficients to examine the extent to which anthropometric measurements were correlated with adiposity at each time point. We chose Spearman (r_s) rather than Pearson correlation coefficients because they do not assume normality of data or a linear relationship between the measurements. To evaluate whether changes in BMI and BMIZ were associated with changes in adiposity, we calculated Spearman correlation coefficients between pairs of changes in anthropometric measurements and concurrent changes in body composition. Pearson correlations were almost identical to Spearman correlations (data not shown).

We used stratification to assess for differences in correlations among racial/ethnic groups, between sexes, and by feeding type (breastmilk or formula feeding). We used the method of Zou (33) to assess for significant differences between correlated correlation coefficients with a variable in common (such as whether BMI is more strongly correlated with fat mass than lean mass). To compare the correlation between 2 given variables across subsamples (such as whether correlations differ between males and females), we applied the Fisher ztransformation (tanh⁻¹r) and performed an asymptotic z-test (for 2 groups) or chi-squared test (for 3 groups), using the sample-specific standard errors $((n-3)^{-1/2})$. Significance was set at $P = .05$.

Our primary focus was to examine associations of BMI, rather than other possible indices of body proportionality such as WFL or ponderal index $(kg/m³)$, with DXA-assessed adiposity because BMI has been shown to correlate with adiposity more strongly than ponderal index does in infants (21). Further, BMIZ changes are more strongly correlated than WFLZ with body fat changes in infancy (27), and BMIZ is more strongly predictive of early childhood obesity than WFLZ (13). Although the CDC recommends the use of WFLZ until 24 months of age (34) in part due to the inaccuracy of clinical length measurements in infants, BMI has recently been recommended over WFL for monitoring body proportionality in infancy (13) and is commonly used in clinical practice (35). For completeness, we explored WFL and WFLZ in addition to BMI and present those results as well. A secondary focus was on the relationship between anthropometrics and measures of lean mass.

Statistical analysis was performed using IBM SPSS Statistics software version 22.0 (IBM Corp, Armonk, New York) and SAS software version 9.4 (SAS Institute Inc, Cary, North Carolina).

RESULTS

The 447 infants were 38% Caucasian, 31% African-American, 29% Hispanic, and 2% Asian; 52% were male (Table 1). Of the included infants, DXA and anthropometric data were available for 437 infants (98%) at 1 month of age (mean \pm standard deviation of exact

age at visit: 36 ± 7 days), 331 infants (74%) at 4 months of age (125 \pm 7 days), and 277 infants (62%) at 7 months of age (214 \pm 10 days) (Figure 1; available at www.jpeds.com). Infants who dropped out of the study before the last visit had lower birth weights (3254g vs. 3405g, p=0.001), were more likely to be African-American or Hispanic than Caucasian $(78\% \text{ vs. } 50\%, \text{ p} < 0.001)$, and had less educated mothers $(18\% \text{ vs. } 43\% \text{ with college degree})$. p<0.001), but no significant difference in maternal BMI (median 27.8 vs. 26.1, p=0.07), compared with infants who completed the study.

Table 2 shows Spearman correlations among anthropometric variables and body composition at each time point. BMIZ was moderately correlated with body fat percentage at each time point (r_s =0.43, 0.55, and 0.48 respectively). BMIZ was more strongly correlated (p<0.001 for each comparison) with fat mass and FMI than with BF% at all time points (r_s = 0.43, 0.55, 0.48 for BF%; for fat mass and FMI, r_s = 0.56–0.61, 0.76–0.79, and 0.72–0.75 at 1, 4, and 7 months respectively). Correlations of anthropometrics with adiposity were lowest at 1 month for all adiposity measures, such that the correlation of BMIZ and adiposity was significantly stronger at 4 and 7 months than at 1 month ($p=0.03$ for BF%; $p<0.001$ for fat mass and FMI). At the individual level, a wide range of adiposity was associated with a given BMIZ (Figure 2). For example, a 1 month old infant with BMIZ of zero could have a BF% ranging from 9% to 39%.

In addition to being correlated with adiposity, BMIZ was also positively correlated with lean mass (r_s =0.32, 0.28, and 0.30) and LMI (r_s =0.34, 0.28, and 0.29) at 1, 4, and 7 months respectively, with similar correlations for BMI (Table 2). Correlations of BMIZ with lean mass and LMI were significantly lower than with fat mass or FMI at each time point (p<0.001 for each comparison).

All 3 measures of adiposity were more strongly correlated with BMIZ than with WFLZ at 1 month and 4 months (p 0.001 for each comparison). At 7 months, adiposity correlated similarly with BMIZ and WFLZ (p>0.2 for each comparison; Table 2).

Stratification by sex, race/ethnicity, infant feeding type (predominantly breastfed or exclusively formula fed), or maternal feeding intention did not alter correlations of anthropometric with adiposity measures (data available upon request). However, stratification reduced the power of these analyses.

Changes in BMIZ were moderately correlated with changes in adiposity from 1–4 months $(r_s=0.44$ with BF%; $r_s=0.53$ with fat mass; $r_s=0.62$ with FMI). In contrast, from 4–7 months, changes in BMIZ were only weakly correlated with changes in body fat percentage $(r_s=0.21)$ and fat mass $(r_s=0.27)$, and moderately correlated with fat mass index $(r_s=0.49)$ (Table 3).

Changes in BMI and BMIZ between 1–4 months were more strongly correlated with changes in BF%, fat mass, and FMI than they were with lean mass or lean mass index (p<0.001 for all comparisons). However, from 4–7 months, changes in BMI or BMIZ correlated equally well with changes in adiposity as with changes in lean mass or LMI (p=0.10–0.19 for each comparison), except that changes in FMI were significantly more strongly correlated than for LMI (r_s =0.53 vs 0.23; p=0.004).

Changes in adiposity (BF%, fat mass, and FMI) were more strongly correlated with changes in BMIZ than with changes in WFLZ during each period (Table 3). But, the difference was only statistically significant for changes between 1–4 months in BF% (p=0.002; r_s =0.44 with BMIZ, r_s =0.36 with WFLZ) and FMI (p<0.001; r_s =0.62 with BMIZ, r_s =0.53 with WFLZ).

DISCUSSION

The moderate correlations we observed between infant BMI or BMIZ and adiposity $(r_s=0.43-0.55)$ contrast with high correlations of BMI and DXA-derived adiposity measures in older children. For BF%, correlation coefficients in school age children and adolescents range from r=0.79–0.92 (2,3,5) although a correlation as low as r=0.63 was reported in one study (4). Similarly, among school age children, correlations are high between BMI and DXA-derived fat mass (r=0.83–0.95) or FMI (r=0.80) (3–5). Although we found relatively strong correlations of BMI with both fat mass and FMI $(r_s=0.72-0.79)$ at 4 and 7 months, and moderate correlations $(r_s=0.56-0.66)$ at 1 month, these were low relative to previously published studies of older children (2–5). Technical limitations of performing DXA on infants, such as motion artifact, can limit its accuracy (25) and could theoretically contribute to the lower correlations we observed; however, the moderate correlations we observed between BMI or BMIZ and DXA-derived adiposity are consistent with the few prior studies conducted in infants, which used different techniques to measure body composition. De Cunto et al showed BMIZ was moderately correlated $(r=0.65)$ with BF% at birth as measured by air displacement plethysmography (21), and de Bruin et al found a similarly moderate correlation (R^2 =0.44, corresponding to r=0.66) between BMI and BF% using total body electrical conductivity in a grouped sample of infants aged 1–12 months (23). Our findings, taken together with prior studies, suggest that during infancy, BMI and BMIZ are limited as surrogate markers of adiposity.

We showed that prospective changes in BMI were moderately associated with changes in DXA-measured adiposity between 1 and 4 months of life and were more strongly correlated with fat mass gains than lean mass gains. Conversely, between 4 and 7 months, correlations between BMI changes and adiposity changes were weak, and although correlations were generally higher for fat gains than for lean mass gains, the difference was not statistically significant. This discrepancy is likely partly attributable to the typical growth pattern in infancy, in which infants experience a rapid accrual of fat during the first several months of life followed by a period of relatively slower fat mass accrual and relatively larger changes in lean mass (18,36).

BMI and BMIZ trajectories are often interpreted as surrogates for changes in adiposity (10,13,14). However, the associations between changes in anthropometrics and concurrent changes in adiposity have rarely been studied in infancy. One study performed over 20 years ago found that weight z-score changes from birth to 12 weeks were positively correlated with both FMI z-score and lean mass index z-score at 12 weeks, but did not compare associations between FMI and LMI, and did not assess changes beyond 12 weeks (22). A recent large study of >1000 infants showed a positive association between changes in BMIZ and changes in BF% from birth to 5 months of age (27), similar to the positive association we found from 1–4 months. We extend their findings by showing that BMIZ changes

Accurate identification of excess adiposity in infancy is important in the context of the rising prevalence of obesity in childhood and the well-established contribution of early infant weight gain to later-life risk of obesity and its associated complications (12–14,37). Public health efforts aimed at obesity prevention increasingly target early childhood populations, extending even into infancy (13,38,39). For example, changing infant feeding practices by reducing the protein content in formula (40), increasing breastfeeding rates (41), and delaying solid food introduction (38), along with behavioral interventions for parents to address infant sleeping and crying (38), have all been tested as possible interventions to reduce childhood obesity.

Most studies of obesity risk have examined infant weight or BMI gain without assessing body composition (6,10–13), although one small study suggests that rapid gain in fat mass in early infancy is a better predictor of subsequent excess adiposity in childhood than weight gain alone (16). Clinicians and researchers engaged in efforts to reduce infant weight gain as a means of preventing subsequent obesity need accurate methods of identifying adiposity in order to best identify and intervene for infants at high risk of developing adiposity-related complications later in childhood and adulthood.

Our finding that infant BMI or BMI gain do not accurately represent adiposity or adiposity gain is important in the context of several studies linking early rapid BMI gain with later obesity (10,12–15). These studies interpreted BMI gain as a surrogate for excess accrual of adipose tissue in early life, predisposing the infant to subsequent excess adiposity (16,17). Our results suggest that this interpretation may not be correct. Instead, the link between rapid BMI gain in infancy and later obesity may involve changes in lean mass or a combination of lean and adipose tissue (43,44). Studies incorporating directly measured infant body composition are needed to establish the extent to which rapid fat and/or lean mass gain during infancy explains this association.

The use of DXA for body composition measurement in infants has several limitations, including dependence of the results on type of scanner and software, sensitivity to motion artifacts, and overestimates of fat mass compared with other body composition techniques (including ADP and the 4-compartment model) (25). To minimize these limitations, we used the same DXA model and software throughout the study and only used scans free from motion artifact. The racial/ethnic diversity of our sample and recent time course of the study increase its generalizability over most prior studies that were conducted over two decades ago in racially and ethnically homogenous populations. However, the proportionately greater dropout of infants from racial/ethnic minorities and lower socioeconomic status may limit generalizability of the findings in later infancy. Also, our finding may not be generalizable to infants in resource-poor settings, where correlations between infant growth and adiposity differ from those in industrialized countries (46) .

BMI and BMIZ are moderately correlated with adiposity in infancy, as compared with high correlations in older children and adults. An increasing BMI or BMIZ is a poor indicator of increasing adiposity in later infancy. Therefore, measurement of BMI and BMIZ may not be sufficient for accurate assessment of adiposity or adiposity changes in infancy.

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ABBREVIATIONS

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Figure 2.

Scatterplots of body fat percentage and BMI z-score in full term infants at 1 month (panel A; N=437), 4 months (panel B; N=331), and 7 months (panel C; N=277).

Table 1

Participant characteristics.

* Data are displayed as mean ± standard deviation.

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Table 2

Correlations among anthropometric measurements and body composition measurements in 447 full term infants aged 1–7 months.

J Pediatr. Author manuscript; available in PMC 2019 May 01.

*

Values shown are Spearman correlation coefficients. Values shown are Spearman correlation coefficients. BF%, body fat percentage; BMIZ, BMI z-score; FMI, fat mass index; LMI, lean mass index; WFLZ, weight-for-length z-score. N=437 at 1 month, 331 at 4 months, and 277 at 7 months. BF%, body fat percentage; BMIZ, BMI z-score; FMI, fat mass index; LMI, lean mass index; WFLZ, weight-for-length z-score. N=437 at 1 month, 331 at 4 months, and 277 at 7 months.

 NS Denotes the correlation is not statistically significant (p>0.05). All other correlations without this superscript are statistically significant with p<0.01. NS Denotes the correlation is not statistically significant (p>0.05). All other correlations without this superscript are statistically significant with p<0.01.

Table 3

Correlations between concurrent changes in anthropometrics and changes in body composition for 323 full term infants aged 1–7 months.

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 $*$ Values shown are Spearman correlation coefficients. N=323 from 1-4 months; N=275 from 4-7 months. Values shown are Spearman correlation coefficients. N=323 from 1–4 months; N=275 from 4–7 months.

BF%, body fat percentage; BMZ BMI z-score; FMI, fat mass index; LMI, lean mass index; WFLZ, weight-for-length z-score; , change. BF%, body fat percentage; BMIZ, BMI z-score; FMI, fat mass index; LMI, lean mass index; WFLZ, weight-for-length z-score; Δ, change.

NS Denotes correlation is not statistically significant (p>0.05). All other correlations are significant with p<0.05. NS Denotes correlation is not statistically significant (p>0.05). All other correlations are significant with p<0.05.

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