Development of population-specific prediction equations for bioelectrical impedance analyses in Vietnamese children

Phuong Hong Nguyen^{1,2}, Melissa F. Young³, Long Quynh Khuong⁴, Usha Ramakrishnan³, Reynaldo Martorell³ and Daniel J. Hoffman⁵*

¹Poverty, Health and Nutrition Division, International Food Policy Research Institute (IFPRI), Washington, DC 20006, USA ²Thai Nguyen University of Pharmacy and Medicine, Thai Nguyen, 24000, Vietnam

³Hubert Department of Global Health, Rollins School of Public Health, Emory University, Atlanta, GA, USA

⁴Center for Population Health Science, Hanoi University of Public Health, Hanoi, 10000, Vietnam

⁵Department of Nutritional Sciences, Program in International Nutrition, New Jersey Institute for Food, Nutrition, and Health, Center for Childhood Nutrition Research, Rutgers, the State University of New Jersey, New Brunswick, NJ, USA

(Submitted 25 November 2019 - Final revision received 2 June 2020 - Accepted 10 June 2020 - First published online 3 July 2020)

Abstract

There is a need for accurate, inexpensive and field-friendly methods to assess body composition in children. Bioelectrical impedance analysis (BIA) is a promising approach; however, there have been limited validation and use among young children in resource-poor settings. We aim to develop and validate population-specific prediction equations for estimating total fat mass (FM), fat free-mass (FFM) and percentage body fat (PBF) in Vietnamese children (4–7 years) using reactance and resistance from BIA, anthropometric variables and demographic information. We conducted a cross-sectional survey of 120 children. Body composition was measured using dual-energy X-ray absorptiometry (DXA), BIA and anthropometry. To develop prediction equations, we split all data into development (70%) and validation datasets (30%). The model performance was evaluated using predicted residual error sum of squares, root mean squared error (RMSE), mean absolute error (MAE) and R^2 . We identified a top performing model with the least number of parameters (age, sex, weight and resistance index or resistance and height), low RMSE (FM 0·70, FFM 0·74, PBF 3·10), low MAE (FM 0·55, FFM 0·62, PBF 2·49), high R^2 (FM 0·95, FFM 0·92, PBF 0·82) and the least difference between predicted values and actual values from DXA (FM 0·03 kg or 0·01 sp, FFM 0·06 kg or 0·02 sp, PBF 0·27% or 0·04 sp). In conclusion, we developed the first valid and highly predictive equations to estimate FM, FFM and PBF in Vietnamese children using BIA. These findings have important implications for future research on the double burden of disease and risks associated with overweight and obesity in young children.

Key words: Body composition: Bioelectric impedance analysis: Children: Prediction: Fat mass



Child obesity is a growing public health epidemic across the globe in which currently 50 million girls and 74 million boys are obese⁽¹⁾. The increased prevalence of overweight and obesity is especially alarming in low- and middle-incomes countries (LMIC), such as Vietnam, that are undergoing the nutrition transition and shifting from traditional to obesogenic diets⁽²⁾. These countries also face the dual burden of a high prevalence of undernutrition and obesity and often accompanied by micronutrient deficiencies^(3–5). Poor nutrition and growth early in life can also increase the risk for non-communicable chronic diseases later in life^(6–10). However, existing tools that are used to measure child growth and body composition, especially in resource poor settings, are often either non-field friendly and/or inaccurate and unreliable⁽¹¹⁾. Therefore, the objective of the

present study was to develop and validate a relatively inexpensive and 'field-friendly' tool to assess body composition in children in LMIC.

Historically, large cohort studies and national surveys have relied on relatively inexpensive and generally reliable anthropometric measures, such as weight, height, waist circumference and skinfold measures, to estimate body composition in children and adults^(12–15). However, these methods are generally not generalisable and may create biased estimates especially when body composition compartments, such as percentage body fat, have been calculated using prediction equations developed from populations that differ by age, race and/or ethnicity^(16–21). For example, Hoffman *et al.*⁽¹⁵⁾ reported that skinfold prediction equations consistently underestimated body fat mass (FM)

Abbreviations: BIA, bioelectrical impedance analysis; DXA, dual-energy X-ray absorptiometry; FM, fat mass; FFM, fat-free mass; LMIC, low- and middle-incomes countries; MAE, mean absolute error; PBF, percentage body fat; RMSE, root mean squared error.

* Corresponding author: Daniel J. Hoffman, email dhoffman@aesop.rutgers.edu



compared with dual-energy X-ray absorptiometry (DXA) and 2H dilution. In addition, Rodríguez et al. (14) found that skinfold prediction equations had poor agreement with body composition measurements estimated by DXA. More recently, many studies are now relying on simpler and more reliable methods that have minimal operator error, such as bioelectric impedance analysis (BIA). While BIA may be considered to less prone to operator error compared with anthropometrics, it is not without limitations, and a recent review described varying degrees of bias attributed to the use of different prediction equations and the inconsistent use of protocols between studies⁽²²⁾.

Estimating body composition with BIA is based on the principle that electricity is impeded as it is conducted through a cylinder and passes easily through hydrated tissue, that is, fat-free mass (FFM), yet resisted by tissue with little to no water, fat mass (FM)⁽²³⁾. Yet, the human body is not one cylinder and can be thought to be composed of five cylinders, the trunk, arms and legs, and differences in proportions between these areas can yield different results for the same BIA. Moreover, the use of BIA is not without challenges as most manufacturers create proprietary equations that cannot be altered and have not been widely validated in children⁽²⁴⁾. Still, a number of groups have published accurate prediction equations for paediatric uses of BIA⁽²⁵⁻²⁷⁾. Developing population-specific prediction equations for BIA is typically done⁽²⁸⁻³⁰⁾ by comparing estimates against a 'gold standard', such as ²H dilution or DXA to assess body composition. For example, DXA was used to develop a nationally representative BIA prediction equation for children in the USA⁽²⁸⁾. These approaches have also been used for special populations, such as neonates⁽³¹⁾ and severely obese adolescents⁽³²⁾. However, gaps remain in the use of BIA to assess body composition in young children from LMIC that may differ by race and ethnicity which have been related to body composition⁽³³⁾.

Given the rise of obesity in LMIC, there is an urgent need for simple yet valid measures of body composition that can be used easily to estimate body fat in young children in resourcepoor environments and better understand how specific early life factors influence the development of adiposity across the lifespan. The objective of this paper was to develop and validate population-specific prediction equations for estimating FM and FFM in young Vietnamese children using reactance, resistance from bioelectrical impedance as well as anthropometric variables.

Methods

Study setting and sample

The present study is a collaboration between investigators based at Emory University, Rutgers University and Thai Nguyen University of Medicine and Pharmacy. We conducted a crosssectional survey of 120 healthy children aged 4-7 years who were recruited from local schools and daycare centres in the city of Thai Nguyen, Vietnam. The study site was selected based on an existing partnership between Emory University and Thai Nguyen University of Medicine and Pharmacy and availability of resources, especially trained personnel and ease of access to a DXA machine at Thai Nguyen National Hospital, teaching hospital for Thai Nguyen University of Medicine and Pharmacy. We employed a convenience sampling framework to recruit children with an equal distribution of BMI z-score (underweight, normal and overweight), sex and age. Written informed consent was obtained from all mothers and/or primary caregiver, and all study protocols were approved by the Human Investigations Committees in Vietnam and Emory University.

Measures

Seven anthropometric measurements, height, weight, triceps and subscapular skinfolds and waist, hip, and mid upper arm circumferences, were obtained in duplicate by trained staff using standardised procedures (34-36). Weight and height were measured using a calibrated digital scale (Seca Corporation) and a fixed stadiometer (Seca Corporation), respectively. Skinfold thicknesses and circumferences were measured using Lange skinfold calipers and flexible, non-stretchable measuring tape. BMI z-scores were calculated by comparing each child's weight and height measurements with the WHO standards (37), and children were categorised as low BMI (<-1 sD), normal (-1 to 1 sd) or overweight (>1 sd).

Body composition was measured using two different methods. First, we assessed body composition using a Seca mBCA 525 multifrequency BIA (Seca Corporation). All subjects had been fasting for at least 4 h with no vigorous exercise for 24 h before the measurement. All children wore minimal clothing and rested in the prone position for 5 min prior to the measurement. The position of each child's arms and legs was done according to the manufacturer's protocol. Specifically, the feet and thighs are not in contact with each other, and the hands and arms are placed beside the body without touching the body. Finally, the children were told to lie still and remain relaxed during the measurement. Using an 8-polar tactile-electrode impedance meter, four electrodes were placed on the palm and thumb of both hands, and four electrodes were placed on the anterior and posterior aspects of the soles of both feet. The raw values of total body resistance and reactance at 50 kHz were recorded for the data presented. Second, wholebody DXA scanning was used as the reference body composition measurement (Hologic Discovery DXA System, Hologic Inc). Scans were conducted using paediatric software according to the manufacturer-recommended protocol to obtain estimates of total FM, total FFM and percentage body fat (PBF).

Statistical methods

Descriptive analyses were conducted for general characteristics of study sample. The candidate predictors for the models were selected based on their clinical importance as reported in previous studies (25,38). Predictor variables included age, sex, five anthropometric variables (height, weight, waist circumference, hip circumference, subscapular skinfold), resistance and reactance as well as resistance index (RI, height in cm²/resistance).

We developed prediction equations by following the four steps of model development that have been used in previous validation studies^(25,28). In step 1, the random-split method was used to create the development and validation datasets with 70 and 30% of the total observations in the original dataset,



respectively. All potential models were built in the development set and tested in the validation set. In step 2, models were generated in the development dataset using the Least Absolute Shrinkage and Selection Operator technique (39). Estimation of the penalty parameter for the Least Absolute Shrinkage and Selection Operator was based on grid search with 10-fold cross-validation as the optimisation criterion. In step 3, all final model equations were tested in the validation dataset. The predicted FM, FFM and PBF (FM_{PRED}, FFM _{PRED} and PBF_{PRED}) were calculated in the validation data using the coefficients of the models selected in step 2 and then compared with the actual values from DXA (denoted as DXA_{FM}, DXA_{FFM} and DXA_{PBF}). The model performance was evaluated using the following four metrics: predicted residual error sum of squares, root mean squared error (RMSE), mean absolute error (MAE) and R^2 . The PRESS, RMSE and MAE measure the discrepancy between the predicted and the actual value, thus the smaller value, the closer the fit between the model and the data. The R^2 provides the proportion of the variance of an outcome variable that is explained by predictor variables in a multiple linear regression model such that a higher \mathbb{R}^2 indicates a better fit. In step 4, the final equations for FM_{PRED}, FFM_{PRED} and PBF_{PRED} were produced based on the selected model using the full dataset.

Sensitivity analyses were conducted to compare the performance of models that were developed from a list of candidate variables in which seven different subsets of candidate predictors were tested (models 1–7). All candidate predictors were first included in model 1 and then were removed one by one for each subsequent model until model 6 that contained only age, sex and weight and model 7 that included only RI and reactance. We also developed two sets of models, the first set included RI without height (as height in already included in the RI) and the second set included resistance and height to determine if using RI provided a better fit than the including the variables independently.

All statistical analyses were performed using Stata (version 15.2) and R (version 3.5.0), and statistical significance was set at P < 0.05.

Results

Descriptive statistics for anthropometric measurements and other sample characteristics are presented in Table 1, while estimates of body composition from BIA and DXA are shown in Table 2. There were no statistical differences in mean age, sex, height, weight, skinfold thicknesses or anthropometric

Table 1. Anthropometric characteristics of children in Vietnam by total sample and development and validation groups (Mean values and standard deviations; numbers and percentages)

	Total (n 119)			Development (n 83)			Validation (n 36)			
	Mean		SD	Mean		SD	Mean		SD	Р
Age (months)	74.3		11.8	74.7		11.5	73.3		12.5	0.573
Min-max		53-2-95-3			54-6-95-3			53.2-89.5		
Sex (%)										
Female		49.6			47.0			55.6		0.510
Male		50.4			53.0			44.4		
Height (cm)	114.4		7.9	114.9		7.9	113.2		8.0	0.311
Weight (kg)	21.1		5.6	21.3		5.8	20.6		5.3	0.503
Weight for age (z-score)	-0.2		1.6	-0.2		1.5	-0.3		1.6	0.634
Height for age (z-score)	-0.4		1.0	-0.4		1.0	-0.5		1.1	0.431
BMI (z-score)	0.0		1.6	0.1		1.6	-0.0		1.6	0.846
Triceps skinfold (mm)	10.3		4.9	10-4		5.0	9.9		4.9	0.607
Subscapular skinfold (mm)	8.4		4.9	8.5		5.0	8.0		4.8	0.644
Mid-upper arm circumference (cm)	18-6		3.0	18-8		3.1	18.2		2.9	0.346
Waist circumference (cm)	53.5		7.1	53.8		7.3	52.9		6.6	0.499
Hip circumference (cm)	57.8		6.9	58-0		7.0	57.3		6.7	0.621

Table 2. Body composition data of children in Vietnam by total sample and development and validation groups using bioelectrical impedance analysis (BIA) or dual-energy X-ray absorptiometry (DXA) (Mean values and standard deviations)

	Total (n 119)		Developm	ent (n 83)	Validation (n 36)		
	Mean	SD	Mean	SD	Mean	SD	Р
Resistance (50 kHz)	745.5	82-6	748-0	82-8	739-8	82.9	0.620
Reactance (50 kHz)	61.3	6.9	61.9	7.4	60.0	5.4	0.107
RI	17.9	3.6	18.0	3.7	17.7	3.4	0.634
DXA fat mass (kg)	6.6	3.2	6.7	3.2	6.3	3.1	0.559
DXA percentage fat (%)	30.0	7.2	30.2	7.1	29.6	7.4	0.700
DXA fat-free mass (kg)	14.5	2.9	14-6	3.0	14.1	2.6	0.355
DXA weight (kg)	21.1	12.5	21.3	5.8	20.5	5.3	0.435
Bone mineral density (mg/cm²)	0.7	0.1	0.7	0.1	0.7	0.1	0.873

RI, resistance index (height (cm)²/resistance)



indicators including WAZ, WHZ or HAZ or skinfold measures between the development and validation sub-samples. There were also no differences in mean values of resistance and reactance, RI, FM, FFM or PBF between the development and validation sub-samples.

The development of prediction equations for total FM resulted in seven models that were applied to the validation sub-sample as shown in Table 3. Models 6 and 7 had the poorest performance with highest RMSE and MAE and lowest R^2 . Although there were modest differences between the RMSE, MAE and R^2 for models 1–5, model 5 performed the best with the least number of parameters (age, sex, weight and resistance index) compared with 5-8 parameters in models 1-4, while maintaining a low RMSE (0.70), low MAE (0.55), high R^2 (0.95) and least difference between FM_{PRED} and DXA_{FM} (0.03 kg). We obtained similar results for FFM (Table 4) with model 5 (using resistance and height instead of resistance index). Results for the development of a prediction equation for PBF are provided in Table 5 in which model 5 also performed the best, but with a lower R^2 (0.82) compared with the model for FM or FFM (>0.90).

Results by sex are provided in the supplemental tables. There were no significant differences between sex for raw anthropometric measures except waist circumference which was higher in boys than girls (online Supplementary Table S1). Boys also had higher resistance, lower reactance and a higher RI compared

with girls. As well, boys had greater LBM and BMD compared with girls.

For sensitivity analyses, we tested different variables for prediction models using height and resistance as two independent variables or as RI. For prediction of FM (online Supplementary Table S2), we obtained models with only slightly higher RMSE and MAE and minimally lower R^2 , but larger mean differences between FM_{PRED} and DXA_{FM}. For prediction of FFM (online Supplementary Table S3), the models with RI generated greater mean differences between FFM_{PRED} and DXA_{FFM} compared with models using height and resistance as two independent predictors. In contrast, for prediction of PBF (online Supplementary Table S4), the models with RI generated smaller mean differences between PBF_{PRED} and DXA_{PBF} compared with models using height and resistance as two independent predictors.

Discussion

Given the increased prevalence of the double burden of malnutrition (DBM) in many LMIC, it is imperative that validated techniques be developed to assess body composition in children to better understand the aetiology of the DBM and assess programmes to prevent the DBM. To that end, we successfully developed a prediction equation that allows us to use BIA to obtain accurate estimates of FFM, FM and PBF in a sample of young children in Vietnam. The final equation that included

Table 3. Prediction models developed for total fat mass (FM) and statistics from the validation sample using novel models in the validation sample of children in Vietnam

(Mean values and standard deviations; coefficient values and 95 % confidence intervals)

	Model 1	Model 2		Model 4 Coefficient	Model 5 Coefficient	Model 6	Model 7	Applied model*
	Coefficient	Coefficient				Coefficient	Coefficient	
Regression coefficients to	predict FM from	selected varial	oles in the deve	lopment sub-sa	mple			
Intercept	− 2·130	-2.379	-2.039	1.645	-0.255	−1 ·577	-2.282	-0.579
Age	-0.025	-0.031	-0.030	-0.030	-0.040	-0.061	_	-0.039
Female	0.271	0.413	0.456	0.317	0.503	0.798	_	0.501
Weight	0.519	0.655	0.661	0.800	0.788	0.587	_	0.796
Resistance index	-0.276	-0.375	-0.369	-0.438	-0.391	_	0.598	-0.385
Reactance	-0.019	-0.021	-0.021	-0.032	_	_	-0.029	_
Waist circumference	0.060	0.070	0.087	_	_	_	_	_
Hip circumference	0.024	0.027	_	_	_	_	_	_
Subscapular skinfold	0.122	_	_	_	_	_	_	_
Statistics from the validation	on sub-sample ι	using models cre	eated in the dev	elopment sub-s	ample			
PRESS	14.58	18.89	18.94	18-65	17.6	30.73	173.45	
RMSE	0.63	0.72	0.73	0.72	0.70	0.92	2.20	
MAE	0.51	0.60	0.60	0.58	0.55	0.75	1.65	
R ²	0.958	0.945	0.945	0.946	0.949	0.911	0.498	
DXA _{FM}								
Mean	6.34	6.34	6.34	6.34	6.34	6.34	6.34	
SD	3.14	3.14	3.14	3.14	3.14	3.14	3.14	
FM _{PRED}								
Mean	6.39	6.39	6.39	6.40	6.36	6.43	6.57	
SD	2.88	2.86	2.85	2.84	2.83	2.77	2.10	
MSD (DXAFM - FMPRED)	-0.05	-0.06	-0.06	-0.06	-0.03	-0.09	-0.23	
MSD 95 % CI (LB)	-0.35	-0.33	-0.33	-0.66	− 0·71	−1 ·52	− 3·78	
MSD 95 % CI (UB)	0.07	0.15	0.15	-0.18	-0.25	− 0·91	-2.33	
MSD _{boy}	- 0·16	- 0·16	− 0·15	-0.12	0.00	-0.07	-0.69	
MSD _{girl}	0.03	0.03	0.02	-0.02	-0.05	-0.12	0.14	

PRESS, predicted residual error sum of squares; RMSE, root mean square error; MAE, mean absolute error; DXA_{FM}, FM from dual-energy X-ray absorptiometry; FM_{PRED}, predicted FM; MSD, mean signed difference; LB, lower bound; UB, upper bound.



^{*} Final equation after model 5 was applied to all data. FM = -0.039 x age (months) + 0.501 x female (1) + 0.796 x weight (kg) - 0.385 x resistance index - 0.579

Table 4. Prediction models developed for total fat-free mass (FFM) and statistics from the validation sample using novel models in the validation sample of

(Mean values and standard deviations; coefficient values and 95 % confidence intervals)

	Model 1	Model 2	Model 3 Coefficient	Model 4 Coefficient	Model 5 Coefficient	Model 6	Model 7	
Predictors	Coefficient	Coefficient				Coefficient	Coefficient	Applied model*
Results from development s	sample							
Intercept	-5.134	-6.248	-6.248	-6.227	-6.380	-10.830	26.669	-5 ⋅539
Age	0.013	0.012	0.012	0.012	0.013	0.008	_	0.008
Female	-0.374	-0.422	-0.422	-0.424	-0.546	-0.741	_	-0.638
Height	0.170	0.189	0.189	0.188	0.193	0.171	_	0.189
Weight	0.231	0.169	0.169	0.169	0.169	0.262	_	0.167
Resistance	-0.008	-0.008	-0.008	-0.008	-0.007	_	-0.026	-0.007
Reactance	0.022	0.019	0.019	0.020	_	_	0.122	
Waist circumference	-0.007	X	X	_	_	_	_	
Hip circumference	X†	X	_	_	_	_	_	
Subscapular skinfold	-0.040	_	_	_	_	_	_	
Statistics from the validation	sample using	models created	in the developr	ment sample				
PRESS	19.22	19.79	19.79	19.87	19.73	25.55	151.34	
RMSE	0.73	0.74	0.74	0.74	0.74	0.84	2.05	
MAE	0.60	0.61	0.61	0.61	0.62	0.72	1.63	
R^2	0.920	0.918	0.918	0.918	0.918	0.894	0.373	
DXA _{FFM}								
Mean	14.12	14.12	14.12	14.12	14.12	14.12	14.12	
SD	2.63	2.63	2.63	2.63	2.63	2.63	2.63	
FFM _{PBED}								
Mean	14.18	14.18	14.18	14.17	14.18	14.08	14.60	
SD	2.79	2.80	2.80	2.80	2.83	2.75	1.95	
MSD (DXAFFM-FFMPRED)	-0.06	-0.06	-0.06	-0.05	-0.06	0.04	-0.48	
MSD 95 % CI (LB)	-0.33	-0.37	-0.37	-0.37	-0.40	-0.22	0.57	
MSD 95 % CI (UB)	0.15	0.12	0.12	0.12	0.09	0.34	1.89	
MSD _{bov}	0.14	0.13	0.13	0.13	0.06	0.12	0.18	
MSD _{girl}	-0.23	-0.21	-0.21	-0.21	-0.17	-0.04	-1.02	

PRESS, predicted residual error sum of squares; RMSE, root mean square error; MAE, mean absolute error; DXA_{FFM}, FFM from dual energy X-ray absorptiometry; FFM_{PRED}, predicted FFM: MSD, mean signed difference; LB, lower bound; UB, upper bound.

raw data from BIA along with sex, age and body weight predicted FM within 25 g of DXA_{FM} with an R^2 of 0.94 and FFM within 8 g of DXA_{FFM} with an R^2 of 0.93. As well, the prediction equation developed for PBF had an R^2 of 0.82 with that estimated from DXA. It is also important to note that the standard deviations of the absolute values of the predicted FM and FFM compared with DXA_{FM} and DXA_{FFM} were similar, suggesting good agreement between the predicted and actual measures. The correlation between our prediction equations developed in Vietnamese children is similar with studies that used other population-specific BIA prediction equations and reported R^2 between 0.80 and $0.93^{(16,29,30)}$

Techniques to estimate body composition in children that are logistically simple for field and survey research have generally relied on skinfold measures and other anthropometric techniques (34,36). Skinfold measurements are found to be predictive of body composition in a diverse population of children with correlation coefficients as high as 0.85-1.00 for DXA_{FM} and DXA_{FFM}, respectively⁽⁴⁰⁾. However, skinfold measures are prone to operator error and can differ by population groups as reported by Hoffman et al. (15) in which three skinfold predictions were found to underestimate PBF by 2-4 percentage points compared with that measured by DXA. Thus, the use of BIA may reduce operator error, but without consistent protocols, bias between BIA and 'gold standards' may persist (41-43). As well, BIA requires a specific prediction equation to generate the estimate of body composition from the requisite resistance and reactance data of the BIA.

The benefits of BIA are many, but mainly centered around ease of use and the relatively inexpensive equipment compared with DXA or stable isotopes. However, a key challenge for the use of BIA in children has been the lack of universal prediction equations for estimating body composition. To address this issue, investigators throughout the world have developed and validated specific prediction equations for children living in different populations. For example, in a study of Asian neonates, the use of weight, length, sex and RI (height (cm)^{2/}resistance) had an R^2 of 0.90 compared with the FFM estimate from airdisplacement plethysmography⁽³¹⁾. A study of 3-year-old children in Denmark reported the estimation of FFM using weight, height, sum of skinfolds and RI had an R² of 0.85 compared with the FFM estimate from DXA⁽⁴⁴⁾. As well, in a study of 2-year-old multi-ethnic children, RI, weight, height and sex were used to predict FFM with an R^2 of 0.89 compared with the FFM estimate from DXA⁽⁴⁵⁾. Finally, in a study of Mexican children, using FFM estimated from ²H dilution compared with a BIA prediction equation that included only RI and weight yielded an R^2 of 0.96⁽⁴⁶⁾. Thus, our results, using age, sex, weight and RI, resulted



Final equation after model 5 was applied to all data. FFM = 0.008 x age (months) - 0.638 x female (1) + 0.189 x height (cm) + 0.167 x weight (kg) - 0.007 x resistance - 5.539. † X: Predictor was excluded by LASSO.



Table 5. Prediction models developed for percentage body fat (PBF) and statistics from the validation sample using novel models in the validation sample of children in Vietnam

(Mean values and standard deviations; coefficient values and 95 % confidence intervals)

	Model 1	Model 2 Coefficient	Model 3 Coefficient	Model 4 Coefficient	Model 5	Model 6	Model 7	
Predictors	Coefficient				Coefficient	Coefficient	Coefficient	Applied model*
Regression coefficients to	predict PBF fro	m selected varia	ables in the dev	elopment sub-s	ample			
Intercept	23.70	22.91	24.26	38-39	29.96	24.63	30.34	28.00
Age	-0.13	-0.15	-0.15	-0.15	-0.19	-0.28	0.65	-0.19
Female	1.90	2.32	2.55	2.03	2.85	4.00	-0.19	2.84
Weight	1.06	1.45	1.46	2.00	1.95	1.14	_	2.01
Resistance index†	−1 ·26	−1 ·54	−1 ·49	−1 .77	−1 ·56	_	_	-1.54
Reactance	-0.09	− 0·10	-0.10	-0.14	_	_	_	_
Waist circumference	0.20	0.23	0.33	_	_	_	_	_
Hip circumference	0.13	0.14	_	_	_	_	_	_
Subscapular skinfold	0.36	_	_	_	_	_	_	_
Statistics from the validation	on sub-sample ι	using models cre	eated in the dev	elopment sub-s	ample			
PRESS	314-36	353.06	355.15	347.70	345.01	540.05	1547-2	
RMSE	2.95	3.13	3.14	3.11	3.10	3.87	6.56	
MAE	2.34	2.52	2.58	2.50	2.49	3.39	5.23	
R^2	0.836	0.816	0.815	0.819	0.820	0.719	0.194	
DXA _{PBF}								
Mean	29.59	29.59	29.59	29.59	29.59	29.59	29.59	
SD	7.41	7.41	7.41	7.41	7.41	7.41	7.41	
PBF _{PRED}								
Mean	29.90	29.91	29.90	29.94	29.77	30.04	30-31	
SD	5.88	5.84	5.76	5.73	5.75	5.37	2.70	
MSD _{DXAPBF - PBFPRED}	-0.31	-0.32	-0.31	-0.35	-0.27	-0.45	-0.72	
MSD 95 % CI (LB)	-2.88	-2.78	-2.84	-4.03	-4.30	−7 ·48	–11 ⋅57	
MSD 95 % CI (UB)	-0.93	-0.72	-0.77	-1.98	-2.25	-4.93	−7 ·25	
MSD _{boy}	-0.69	-0.67	-0.61	-0.53	0.04	-0.26	-2.49	
MSD _{qirl}	-0.02	-0.04	-0.07	-0.22	-0.36	-0.62	0.68	

PRESS, predicted residual error sum of squares; RMSE, root mean square error; MAE, mean absolute error; DXA_{PBF}, PBF from dual-energy X-ray absorptiometry; PBF_{PRED}, predicted PBF; MSD, mean signed difference; LB, lower bound; UB, upper bound.

* Final equation after model 5 was applied to all data. PBF = $-0.19 \times \text{age}$ (months) $+2.84 \times \text{female}$ (1) $+2.01 \times \text{weight}$ (kg) $-1.54 \times \text{resistance}$ index +28.00.

† Resistance index (height in cm²/resistance).

in a R^2 of 0.93 compared with the FFM estimate from DXA are comparable to other studies using similar techniques in different ethnic and geographic groups.

Our findings are consistent with other studies, but some limitations remain that merit discussion. First, the choice of outcome variable is important as it has implications for subsequent uses. For example, we relied on a two-compartment model that estimated FM and FFM and did not consider the contribution of other compartments, such as total body water, that would have required additional assessments, such as isotope dilution, that are more invasive and time consuming. However, DXA is considered a 'gold standard' for human body composition that is comparable to three- or four-compartment models of body composition. Second, including key variables that are meaningful and inherently related to the outcome of interest, such as height, weight, resistance, and accompanying skinfold measures, increases the validity, but caution needs to be exercised when such measures may introduce bias and operator error. While a number of other papers on this topic have developed prediction equations that include both height and RI(31,45), we followed the protocol that did not include height in any models that also contained RI⁽²⁸⁾ as doing so overcontrols for the influence of height on the relationship between other variables and the DXA outcome given that it is already accounted for in the RI. Moreover, some anthropometric variables that were dropped in our analyses, such as waist circumference, may still be relevant given they are risk factors for hypertension, elevated TAG and insulin resistance. Yet, it should be noted there are other valid reasons for dropping variables when considering models of similar statistical properties, such as subscapular skinfold measures that required additional equipment the partial removal of clothes, a protocol that may be poorly accepted in many cultures, especially when measuring women and adolescent girls. Third, it has been suggested to correct for the disparity in body composition compartments owing to DXA measures that may underestimate one compartment over the other and estimate total body weights that are not equal to digital scales. We did not use a correction factor as the body weights estimated by DXA were statistically equal to that of the digital scale. Finally, it is important to note that the prediction equations developed in this cohort of healthy Vietnamese children may not be generalisable for other ethnicities but the fact that such equations are not generalisable underscores the need and importance of developing population-specific prediction equations. We tested our equations with another equation developed in a multi-national sample of children⁽²⁶⁾ that had a much lower R^2 (0.73) compared with ours (0.91) as summarised in online Supplementary Table S5.

Our findings will be useful to better characterise the DBM that is a major public health problem in many LMIC, especially in Asia as well as immigrant groups in many developed countries, such as the USA^(5,47). In fact, one study from Vietnam reported that up to 22 % of children were found to be obese⁽⁴⁸⁾. As well, improving our understanding of specific biological or environmental





factors that promote the DMB remains a priority for research and development programmes. Therefore, developing techniques and research tools to more accurately address the issue of paediatric body composition in LMIC will greatly enhance the ability to address the DMB.

The results of the present study have a number of broad implications for research and policies in LMIC. For example, groups in a number of countries in South East Asia will benefit from this work by having access to accurate estimates of body composition in children. Both national and international organisations need simple and inexpensive methods to better evaluate the impact of programmes that are designed to promote healthy growth and lower the prevalence of childhood obesity. Finally, improving the ability to measure body composition in LMIC sets the stage for improving protocols and programmes to reverse the DBM.

In summary, we developed the first valid and highly predictive equations to estimate FM, FFM and PBF in Vietnamese children using BIA. This work is a major contribution that will allow ongoing research studies and national surveys to better estimate the burden and risks associated with overweight and obesity in young children. There is an urgent need to have these methods to support global efforts to prevent DBM as well as understand changes in body composition that occur over the life course especially in settings experiencing during rapid economic development to design appropriate interventions.

Acknowledgements

The authors would like to acknowledge field team at Thai Nguyen University of Medicine and Pharmacy for their work in field supervision and data collection.

Rollins School of Public Health (RSPH) Dean's Pilot and Innovation Grant and the New Jersey Institute for Food, Nutrition, and Health.

P. H. N., M. F. Y., L. Q. K., R. M. and U. R. conceived and designed the study and collected all data. P. H. N. and D. J. H. analysed the data and wrote the manuscript, and M. F. Y., L. Q. K., R. M. and U. R. provided editorial input. All authors reviewed and approved the final version of the manuscript.

None of the authors has any conflicts of interest to declare.

Supplementary material

For supplementary material/s referred to in this article, please visit https://doi.org/10.1017/S000711452000241X

References

- 1. NCD Risk Factor Collaboration (NCD-RisC) (2017) Worldwide trends in body-mass index, underweight, overweight, and obesity from 1975 to 2016: a pooled analysis of 2416 population-based measurement studies in 128-9 million children, adolescents, and adults. Lancet 390, 2627-2642.
- Winichagoon P (2015) Transition of maternal and child nutrition in Asia: implications for public health. Curr Opin Clin Nutr Metab Care 18, 312-317.

- 3. Biswas T, Magalhaes RJS, Townsend N, et al. (2020) Double burden of underweight and overweight among women in South and Southeast Asia: a systematic review and metaanalysis. Adv Nutr 11, 128-143.
- Giao H, Le An P, Truong Vien N, et al. (2019) Stunting and overweight among 12-24-month-old children receiving vaccination in Ho Chi Minh City, Vietnam. Biomed Res Int 2019, 1547626.
- 5. Rachmi CN, Li M & Baur LA (2018) The double burden of malnutrition in Association of South East Asian Nations (ASEAN) countries: a comprehensive review of the literature. Asia Pac I Clin Nutr **27**, 736–755.
- 6. Emmett PM & Jones LR (2015) Diet, growth, and obesity development throughout childhood in the Avon Longitudinal Study of Parents and Children. Nutr Rev, 73, Suppl. 3, 175-206.
- 7. Popkin BM, Richards MK & Montiero CA (1996) Stunting is associated with overweight in children of four nations that are undergoing the nutrition transition. J Nutr 126, 3009–3016.
- Krishnaveni GV & Yajnik CS (2017) Developmental origins of diabetes-an Indian perspective. Eur J Clin Nutr 71, 865-869.
- 9. Eriksson JG (2016) Developmental Origins of Health and Disease - from a small body size at birth to epigenetics. Ann Med 48, 456-467.
- 10. Hoffman DJ, Reynolds RM & Hardy DB (2017) Developmental origins of health and disease: current knowledge and potential mechanisms. Nutr Rev 75, 951-970.
- 11. Kyle UG, Earthman CP, Pichard C, et al. (2015) Body composition during growth in children: limitations and perspectives of bioelectrical impedance analysis. Eur J Clin Nutr 69,
- 12. Lohman TG & Going SB (2006) Body composition assessment for development of an international growth standard for preadolescent and adolescent children. Food Nutr Bull 27, S314-S325.
- 13. Slaughter MH, Lohman TG, Boileau RA, et al. (1988) Skinfold equations for estimation of body fatness in children and youth. Hum Biol 60, 709-723.
- 14. Rodríguez G, Moreno LA, Blay MG, et al. (2005) Body fat measurement in adolescents: comparison of skinfold thickness equations with dual-energy X-ray absorptiometry. Eur J Clin Nutr 59, 1158-1166.
- 15. Hoffman DJ, Sawaya AL, Martins PA, et al. (2006) Comparison of techniques to evaluate adiposity in stunted and nonstunted children. Pediatrics 117, e725-732.
- 16. Nasreddine L, Naja F, Hills AP, et al. (2012) Validity of predictive equations developed to estimate body fat from anthropometry and bioelectrical impedance analysis in 8-10 year-old children. Clin Nutr **31**, 364–371.
- 17. Vicente-Rodríguez G, Rey-López JP, Mesana MI, et al. (2012) Reliability and intermethod agreement for body fat assessment among two field and two laboratory methods in adolescents. Obesity 20, 221-228.
- 18. Gonçalves EM, Silva AM, Santos DA, et al. (2012) Accuracy of anthropometric measurements in estimating fat mass in individuals with 21-hydroxylase deficiency. Nutrition 28, 984-990.
- 19. Deurenberg P & Deurenberg-Yap M (2002) Validation of skinfold thickness and hand-held impedance measurements for estimation of body fat percentage among Singaporean Chinese, Malay and Indian subjects. Asia Pac J Clin Nutr 11,
- 20. Doña E, Olveira C, Palenque FJ, et al. (2018) Body composition measurement in bronchiectasis: comparison between bioelectrical impedance analysis, skinfold thickness measurement, and dual-energy X-ray absorptiometry before and after pulmonary rehabilitation. J Acad Nutr Diet 118, 1464-1473.





- 21. Cameron N, Griffiths PL, Wright MM, et al. (2004) Regression equations to estimate percentage body fat in African prepubertal children aged 9 y. Am J Clin Nutr 80, 70-75.
- Ward LC (2019) Bioelectrical impedance analysis for body composition assessment: reflections on accuracy, clinical utility, and standardisation. Eur J Clin Nutr 73, 194-199.
- Lukaski HC, Bolonchuk WW, Hall CB, et al. (1986) Validation of tetrapolar bioelectrical impedance method to assess human body composition. J Appl Physiol 60,
- 24. Haroun D, Croker H, Viner RM, et al. (2009) Validation of BIA in obese children and adolescents and re-evaluation in a longitudinal study. Obesity 17, 2245-2250.
- Stevens J, Ou F-S, Cai J, et al. (2016) Prediction of percent body fat measurements in Americans 8 years and older. Int J Obes 40, 587-594.
- Liu A, Byrne NM, Ma G, et al. (2011) Validation of bioelectrical impedance analysis for total body water assessment against the deuterium dilution technique in Asian children. Eur J Clin Nutr 65, 1321-1327.
- Kehoe SH, Krishnaveni GV, Lubree HG, et al. (2011) Prediction of body-fat percentage from skinfold and bio-impedance measurements in Indian school children. Eur J Clin Nutr 65, 1263-1270.
- Stevens J, Truesdale KP, Cai J, et al. (2017) Nationally representative equations that include resistance and reactance for the prediction of percent body fat in Americans. Int J Obes 41,
- Hoffman DJ, Toro-Ramos T, Sawaya AL, et al. (2012) Estimating total body fat using a skinfold prediction equation in Brazilian children. Ann Hum Biol 39, 156-160.
- Diouf A, Diongue O, Nde M, et al. (2018) Validity of bioelectrical impedance analysis in predicting total body water and adiposity among Senegalese school-aged children. PLOS ONE 13, e0204486.
- Tint M-T, Ward LC, Soh SE, et al. (2016) Estimation of fat-free mass in Asian neonates using bioelectrical impedance analysis. Br J Nutr 115, 1033-1042.
- Steinberg A, Manlhiot C, Li P, et al. (2019) Development and validation of bioelectrical impedance analysis equations in adolescents with severe obesity. J Nutr 149, 1288-1293
- Liu A, Byrne NM, Kagawa M, et al. (2011) Ethnic differences in the relationship between body mass index and percentage body fat among Asian children from different backgrounds. Br J Nutr 106, 1390-1397.
- 34. Gibson R (2005) Principles of Nutritional Assessment. New York, NY: Oxford University Press.

- 35. Cogill B (2003) Anthropometric Indicators Measurement Guide. Washington, DC: Food and Nutrition Technical Assistance Project, Academy for Educational Development.
- 36. Lohman TG (1988) Anthropometric Standardization Reference Manual. Champaign, IL: Human Kinetics Books.
- World Health Organization (2010) The WHO Child Growth Standards. Geneva: WHO.
- Wan CS, Ward LC, Halim J, et al. (2014) Bioelectrical impedance analysis to estimate body composition, and change in adiposity, in overweight and obese adolescents: comparison with dual-energy X-ray absorptiometry. BMC Pediatr 14, 249.
- Tibshirani R (1996) Regression shrinkage and selection via the Lasso. J R Stat Soc Series B 58, 267-288.
- Wendel D, Weber D, Leonard MB, et al. (2017) Body composition estimation using skinfolds in children with and without health conditions affecting growth and body composition. Ann Hum Biol 44, 108–120.
- 41. McLester CN, Nickerson BS, Kliszczewicz BM, et al. (2018) Reliability, agreement of various inbody body composition analyzers as compared to dual-energy X-ray absorptiometry in healthy men, women. J Clin Densitom (epublication ahead of print version 3 November 2018).
- 42. Ramírez-Vélez R, Tordecilla-Sanders A, Correa-Bautista JE, et al. (2018) Validation of multi-frequency bioelectrical impedance analysis versus dual-energy X-ray absorptiometry to measure body fat percentage in overweight/obese Colombian adults. Am J Hum Biol 30, e23071.
- 43. Lee L-W, Liao Y-S, Lu H-K, et al. (2017) Validation of two portable bioelectrical impedance analyses for the assessment of body composition in school age children. PLOS ONE 12, e0171568.
- 44. Ejlerskov KT, Jensen SM, Christensen LB, et al. (2014) Prediction of fat-free body mass from bioelectrical impedance and anthropometry among 3-year-old children using DXA. Sci Rep 4, 3889.
- 45. Rush EC, Bristow S, Plank LD, et al. (2013) Bioimpedance prediction of fat-free mass from dual-energy X-ray absorptiometry in a multi-ethnic group of 2-year-old children. Eur J Clin Nutr 67, 214-217.
- 46. Ramírez E, Valencia ME, Bourges H, et al. (2012) Body composition prediction equations based on deuterium oxide dilution method in Mexican children: a national study. Eur J Clin Nutr **66** 1099–1103
- 47. Khan NC & Khoi HH (2008) Double burden of malnutrition: the Vietnamese perspective. Asia Pac J Clin Nutr 17, Suppl. 1,
- 48. Hoang NTD, Orellana L, Le TD, et al. (2018) Anthropometric status among 6-9-year-old school children in rural areas in Hai Phong City, Vietnam. Nutrients 10, 1431.

