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Sociodemographic predictors of early postnatal growth: evidence from a Chilean infancy cohort

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ABSTRACT

Objectives Infant anthropometric growth varies across socioeconomic factors, including maternal education and income, and may serve as an indicator of environmental influences in early life with long-term health consequences. Previous research has identified sociodemographic gradients in growth with a focus on the first year and beyond, but estimates are sparse for growth before 6 months. Thus, our objective was to examine the relationship between sociodemographic factors and infant growth patterns between birth and 5 months of age.

Design Prospective cohort study.

Settings Low-income to middle-income neighbourhoods in Santiago, Chile (1991–1996).

Participants 1412 participants from a randomised iron-deficiency anaemia preventive trial in healthy infants.

Main outcome measures Longitudinal anthropometrics including monthly weight (kg), length (cm) and weight-for-length (WFL) values. For each measure, we estimated three individual-level growth parameters (size, timing and velocity) from Superimposition by Translation and Rotation models. Size and timing changes represent vertical and horizontal growth curve shifts, respectively, and velocity change represents growth rate shifts. We estimated the linear association between growth parameters and gestational age, maternal age, education and socioeconomic position (SEP).

Results Lower SEP was associated with a slower linear (length) velocity growth parameter (−0.22, 95% CI −0.31 to −0.13)—outcome units are shifts in days from the average growth curve. Lower SEP was associated with later WFL growth timing as demonstrated through the tempo growth parameter for females (0.25, 95% CI 0.05 to 0.42)—outcome units are shifts in days from the average growth curve. We found no evidence of associations between SEP and the weight size, timing or velocity growth rate parameters.

Conclusion Previous research on growth in older infants and children shows associations between lower SEP with slower length velocity. We found evidence supporting this association in the first 5 months of life, which may inform age-specific prevention efforts aimed at infant length growth.

INTRODUCTION

Interest in early life infant growth has grown as evidence accumulates that it is associated with the development of adult disease, sometimes decades later. Some chronic disease outcomes associated with infant growth characteristics include obesity, endothelial dysfunction and metabolic syndrome. Explanations for these associations include early infancy as a critical window of time for susceptibility to environmental exposures for chronic disease risk factors. Socioeconomic position (SEP) is one such exposure. SEP is associated with child growth patterns, in particular, length and weight. In these studies, lower SEP is generally associated with faster weight gain during childhood, while the inverse holds true for length. These socioeconomic gradients in growth appear to emerge in early life and persist.

Gaps remain in our understanding regarding sociodemographic predictors of growth during infancy and childhood. One such gap relates to the earliest period of infant growth. Most studies to date include three or fewer observations before 6 months, preventing non-linear specifications between weight or height spanning this time. However, curvilinear models...
of growth with more than three observations offer better model fit for early infancy growth. Growth during the first 6 months in the human lifespan is characterised by accelerated growth at the outset and levelling off at around 6 months.17 Given these unique features, early infant growth may yield unique associations with predictors not influential during later periods of growth. Understanding the relationship between early infant growth and sociodemographic factors may yield new information that highlights the potential for earlier interventions to promote optimal health.

Identifying novel associations in this age range can better pinpoint the timing and influence of sociodemographic factors. Given the sparsity of information in the literature focusing on these points, our aim in this study is to examine sociodemographic predictors of infant weight, length and weight-for-length (WFL) growth from 0 to 5 months in an infancy cohort of over 1400 healthy Chilean children. Based on prior research in middle-income to high-income countries applied to a wider range of ages in childhood that is described above, we expected that SEP will be inversely associated with weight gain and positively associated with length growth.

**METHODS**

**Study sample**

The data in this study are drawn from the Santiago Longitudinal Study, a cohort study from low-income to middle-income neighbourhoods in Santiago, Chile. Between 1991 and 1996, infants were recruited for an infancy iron-deficiency anaemia preventive trial18 or neuromaturation study.19 Inclusion criteria for the infancy studies included full-term infants (greater than or equal to 37 weeks gestational age (GA)) with birth weight ≥3.0 kg, vaginal birth, no major health problems for the infant, and, for the preventive trial, no iron-deficiency anaemia present at 5–6 months. Those with iron-deficiency anaemia and the next non-anemic control were invited to participate in the neuromaturation study and are not considered here. Participant eligibility and follow-up information have previously been reported.18

We characterised the growth period prior to treatment randomisation, which occurred at 6 months. Anthropometric measures prior to study enrolment were obtained from the medical chart. The total sample size included 1657 infants who completed the preventive trial.

**Outcome and sociodemographic measures**

Anthropometric measurements included weight (kg), length (cm) and WFL (g/cm). Weight was measured to the nearest 0.01 kg on an electronic scale at local public health clinics. Length was measured on a recumbent board to the nearest 0.1 cm. GA, obtained from the medical chart, was among the set of variables included in the models as a covariate.

Sociodemographic measures were self-reported by the mother, including maternal age (years), total years of education and the modified Graffar Index,20 an index of SEP used in lower income countries.21 The modified Graffar Index represents a sum of 10 measures regarding education, family composition and housing characteristics, which are summed to create a scale with higher values indicating lower social class (online supplementary Appendix Table 1). Mothers self-reported breastfeeding characteristics from birth, including date of first bottle and age at weaning if weaned. From this information, we created variables for breast feeding as the sole source of milk and mixed breast and bottle feeding at 5 months.

**Statistical analyses**

Summary statistics included median and IQRs for continuous variables and per cent with counts for categorical variables. All summary statistics were stratified by child sex. We used two steps to assess the association between infant growth and sociodemographic predictors: (1) Superposition by Translation and Rotation (SITAR) approach22 to estimate infant weight, length and WFL growth characteristics from birth to 5 months followed by (2) linear regression to estimate the relationship between sociodemographic predictors and these growth characteristics. We used a non-linear mixed effects model23 to estimate the growth characteristics with the R nlme package.24 Each model produces up to three different SITAR growth parameters per individual, which have been named ‘size’, ‘tempo’ and ‘velocity’.22 (figure 1). Size indicates a shift of the growth curve up and down for an individual relative to the average growth curve. Tempo indicates a shift of the growth curve to the left or right on the age scale for an individual relative to the average growth curve. Lastly, velocity indicates a transformation of the age scale in the non-linear model, shrinking or enlarging the age scale for an individual relative to the average growth curve. These three parameters are noted as having biologically meaningful interpretations, which are difficult to obtain with other growth models.25 Unless otherwise noted, any references to size, tempo and velocity refer to these parameters from the SITAR construct applied to early infant growth.

Figure 1 Type of change in random effects relative to the sample mean trajectory in weight growth curve trajectories following a shape-invariant model.
The results from the second step analyses are reported. In addition to including males and females and adjusting for sex of the child (in the pooled analyses), sex-stratified analyses were also used for all three anthropometric outcomes, as some estimated associations between SITAR growth parameters and SEP indicators differed by sex of the child.

The adjusted models in the second step started with four covariates: GA, maternal age, total years of maternal education and Graffar Index. We removed covariates from the model based on the least absolute shrinkage and selection operator (lasso) approach. This approach has better performance than conventional model selection methods with a univariate approach, such as stepwise methods. The lasso approach assists in selecting predictors with the strongest coefficients while balancing bias and variation in the model. We used the glmnet package in R to estimate shrunken parameters and the selectiveInference package to provide inference via statistical tests and CIs. Each set of comparisons by outcome, that is, weight, length or WFL was considered separately. Multiple comparisons increase the possibility of statistically significant study findings by chance alone. Therefore, we controlled for multiple comparisons using a Bonferroni correction at an alpha level of 0.05. A coefficient for the predictor of a weight size growth parameter outcome in the second step indicates a change in log(kg) for a one-unit change in the predictor; we multiply this coefficient by 100 to make a symmetric percentage difference on a modified percentage scale. Similarly, a one-unit change in the predictor corresponds to a symmetric percentage change in the velocity growth parameter. Time (days) is not log transformed and the coefficient for this outcome corresponds to a shift in the time scale in days.

For analyses, we used a complete case data set, that is, all participants with non-missing covariates. The proportion of missing data was less than 1% for all variables except the Graffar Index, which had less than 3% missing. The median number of non-missing outcome (anthropometric) values was six out of six monthly measures (birth to 5 months). The per cent of missing outcome values at each time point ranged from 9% at months 1 and 2 to 0.2% at birth. In a post hoc data analysis, we used logistic regression models to estimate associations between SEP (the Graffar Index) as a continuous variable, and binary breastfeeding status outcomes—any or exclusive—at 5 months.

Patient and public involvement
Participants were mothers and infants recruited for research. The mothers were not involved in setting the study design, research questions or outcome measures for this study.

RESULTS
Participants (n=1412) were 53% male and 47% female. Median GA (Q1, Q3) was 40 weeks (39, 40). Median maternal age (Q1, Q3) was 26 years (22, 31), and mothers had a median (IQR) of 10 (8–12) years of education at the time of their infant’s birth (table 1). For the six monthly anthropometric measurements prior to 6 months, each infant had at least two observations, and 72% had measures at all six time points.

We assessed the best model fit for each anthropometric measure via the lowest Bayesian information criterion for growth independent of any covariates. After evaluating all possible combinations of SITAR models from one to three parameters for each of the three anthropometric measures, the best fit (online supplementary Appendix Table 2) models included: (1) all three growth parameters for weight, that is, size, tempo and velocity, (2) sex-specific growth trajectories with tempo and velocity parameters for length and (3) sex-specific growth trajectories with size and tempo parameters for WFL.

The following sections outline the adjusted results of the growth trajectory analyses for the three anthropometric outcomes: weight (kg), length (cm) and WFL (g/cm).

Weight trajectories: size, tempo and velocity
After including all covariates in the model, GA was the only characteristic associated with any weight growth parameters. In the pooled sample, GA was significantly associated with the weight tempo parameter (−2.01, 95% CI −2.98 to −1.70), indicating a leftward shift of about 2 days for each additional week in GA. This indicates earlier timing of weight gain in infants who were born with higher GA (table 2). There was no substantive difference in this association in the sex-stratified analyses.

Length trajectories: tempo and velocity
When evaluating the relationship between deviations from the average length growth characteristics and

| Characteristic                  | Male                     | Female                   | Total                     |
|--------------------------------|--------------------------|--------------------------|---------------------------|
| n                              | 747                      | 665                      | 1412                      |
| Gestational age (weeks)        | 40.0 (39.0–40.0)          | 40.0 (39.0–40.0)          | 40.0 (39.0–40.0)           |
| Graffar Index                  | 27.0 (23.0–33.0)          | 27.0 (23.0–33.0)          | 27.0 (23.0–33.0)           |
| Maternal age (years)           | 26.0 (21.8–30.9)          | 25.5 (21.7–30.3)          | 25.8 (21.8–30.8)           |
| Maternal education (years)     | 10.0 (8.0–12.0)           | 10.0 (8.0–12.0)           | 10.0 (8.0–12.0)            |
Table 2: Sociodemographic predictors and association with weight SuperImposition by Translation and Rotation growth parameter*, stratified by sex of child in the Santiago Longitudinal Study, 1991–1996

| Characteristic       | Unadjusted | Adjusted ‡ | Unadjusted | Adjusted ‡ | Unadjusted | Adjusted ‡ | Unadjusted | Adjusted ‡ | Unadjusted | Adjusted ‡ | Unadjusted | Adjusted ‡ |
|----------------------|------------|------------|------------|------------|------------|------------|------------|------------|------------|------------|------------|------------|
|                      | Males      | Females    | Total      |            |            |            |            |            |            |            |            |            |
|                      | Size Tempo Velocity | Size Tempo Velocity | Size Tempo Velocity | Size Tempo Velocity | Size Tempo Velocity | Size Tempo Velocity | Size Tempo Velocity | Size Tempo Velocity | Size Tempo Velocity | Size Tempo Velocity | Size Tempo Velocity | Size Tempo Velocity |
| Gest age             | 0.59       | -2.28      | -1.96      | 0.76       | -2.38      | -1.58      | 0.64       | -2.35      | -1.96      | -1.06      | 0.53       | -2.01      |
|                      | (-0.12 to 1.31) | (-3.15 to -2.29) | (-1.41) | (-0.02 to 1.54) | (-3.32 to -3.87) | (-0.42 to -3.85) | (0.10 to 1.18) | (-2.98 to -2.70) | (-0.05 to -2.98) | (-2.67 to -2.98) | (-0.01 to -2.15) | (-1.70) |
| Maternal age         | 0.11       | -0.06      | -0.06      | 0.21       | -0.02      | -0.36      | 0.16       | -0.03      | -0.20      | -0.18      | 0.15       | -0.01      |
|                      | (-0.00 to 0.23) | (-0.20 to -0.31) | (-0.11 to -0.33) | (0.07 to 0.15) | (-0.18 to -0.71) | (-0.22 to -0.67) | (0.07 to 0.25) | (-0.14 to -0.78) | (-0.00 to 0.22) | (-0.16 to -0.56) | (-0.01 to 0.34) | (-0.22) |
| Maternal education   | -0.03      | 0.14       | -0.04      | -0.01      | 0.06       | -0.03      | -0.03      | 0.10       | -0.04      | 0.00       | 0.04       | -0.05      |
|                      | (-0.32 to 0.26) | (-0.62 to 0.49) | (-0.21 to 0.06) | (-0.31 to 0.29) | (-0.30 to -0.69) | (-0.30 to -0.95) | (-0.34 to -0.41) | (-0.15 to 0.36) | (-0.50 to 0.41) | (-0.75 to 0.15) | (-0.83 to 4.42) |
| Graffar Index§       | -0.12      | -0.13      | -0.08      | -0.07      | 0.12       | -0.03      | -0.09      | -0.00      | 0.06       | -0.00      | 0.02       | 0.00       |
|                      | (-0.23 to 0.01) | (-0.39 to -0.27) | (-0.22 to -0.41) | (-0.19 to 0.03) | (0.00 to 0.28) | (-0.24 to -0.41) | (-0.18 to to 0.09) | (-0.01 to 0.25) | (-0.00 to 0.04) | (-0.06 to 3.49) | (-0.66 to 3.22) |

*Size units are percentage change in log(weight) from average, tempo units are time (days), velocity units in per cent change from average.
†Bold values indicate significance with Bonferroni correction at alpha level of 0.05.
‡Adjusted linear regression models only include non-zero coefficients from lasso regression models that include all covariates in full model. NA indicates the variable is not included in the adjusted analysis.
§Higher Graffar Index values indicate lower socioeconomic status.
sociodemographic predictors, we found associations for SEP and GA. In the pooled group, the coefficient of association between the Graffar Index and the length parameter (−0.22, 95% CI –0.31 to –0.13; table 3) indicated that for each unit increase in the Graffar Index, lower values indicating higher SEP, there was a −0.22% decline from the average length velocity. Conversely, this association reflects a positive relationship between the average shift in growth relative to the average growth curve as well

### Table 3 Sociodemographic predictors and association with length Superimposition by Translation and Rotation growth parameters, stratified by sex of child in the Santiago Longitudinal Study, 1991–1996

| Characteristic | Tempo | Velocity | Tempo | Velocity | Tempo | Velocity | Tempo | Velocity | Tempo | Velocity |
|---------------|-------|----------|-------|----------|-------|----------|-------|----------|-------|----------|
| Gest age      | −3.33 | 0.99     | −3.05 | NA       | −2.57 | 0.25     | −2.53 | NA       | −2.97 | 0.64     |
|               | (−4.09 to −2.56) | (−4.10 to 1.68) | (−4.36 to −2.55) | (−3.33 to −1.79) | (−3.52 to −2.42) | (−3.13 to −1.51) | (−3.51 to −2.41) | (−3.13 to −1.51) |
| Maternal age  | −0.04 | 0.09     | −0.01 | NA       | −0.17 | 0.01     | −0.15 | NA       | −0.10 | 0.05     |
|               | (−0.18 to −0.10) | (−0.03 to 0.20) | (−0.30 to −0.13) | (−0.29 to 0.14) | (−0.30 to −0.13) | (−0.29 to 0.14) | (−0.30 to −0.13) | (−0.29 to 0.14) |
| Maternal education | 0.06 | 0.12     | NA   | NA       | −0.18 | 0.28     | −0.14 | 0.16     | −0.05 | 0.20     |
|               | (−0.26 to 0.38) | (−0.16 to 0.40) | (−0.49 to 0.13) | (−0.01 to 0.58) | (−0.13 to 0.17) | (−0.01 to 0.52) | (−0.13 to 0.17) | (−0.01 to 0.52) |
| Graffar Index | 0.06  | −0.26    | 0.05  | −0.21    | 0.16  | −0.19    | 0.13  | −0.17    | 0.11  | −0.23    |
|               | (−0.37 to −0.15) | (−0.37 to 0.36) | (−0.25 to −0.14) | (−0.32 to −0.07) | (−0.32 to −0.07) | (−0.32 to −0.07) | (−0.32 to −0.07) | (−0.32 to −0.07) |

*Size units are percentage change in log(length) from average, tempo units are time (days), velocity units in per cent change from average.
†Bold values indicate significance with Bonferroni correction at alpha level of 0.05.
‡Adjusted linear regression models only include non-zero coefficients from lasso regression models that include all covariates in full model. NA indicates the variable is not included in the adjusted analysis.
§Higher Graffar Index values indicate lower socioeconomic status.

WFL trajectories: size and tempo

Evaluations of shifts in WFL size and tempo from the average indicated associations with SEP and GA. Increases in the Graffar Index, equivalent to lower SEP, were associated with a positive shift in the WFL tempo parameter for females (0.25, 95% CI 0.05 to 0.42). This estimate approximates a rightward shift in time (days) relative to the average growth curve indicating later growth timing with lower SEP.

Similar to weight and length trajectory analyses, an increase in GA was inversely associated with a decline in tempo from the average in the pooled sample (−1.99, 95% CI −2.83 to −1.49) (table 4) indicating about a 2-day shift to the left on the time scale from the average growth curve for every 1 week increase in GA. Similar values were found in the sex-stratified analyses, all indicating earlier timing of WFL growth with higher GA.

The post hoc analysis examining the association between odds of exclusive or any breast feeding at 5 months and the continuous SEP measure (the Graffar Index) did not find a substantive or significant association (data not shown).

**DISCUSSION**

In this research, we found that lower SEP, measured by the Graffar Index, was inversely associated with length growth characteristics—but not weight—in the first 5 months. Lower SEP was associated with later timing of WFL growth as reflected by the positive association between the Graffar Index and the WFL tempo parameter. These higher tempo values translate to a rightward shift in growth relative to the average growth curve as well as a later age at peak velocity. This delay in growth can be considered an unfavourable outcome associated with lower SEP.

Maternal age was not associated with any of the three adjusted growth parameters for length, weight or WFL.
GA was inversely associated with the tempo growth parameters for length, weight and WFL indicating that higher GA is associated with earlier timing of these three measures. GA is also positively associated with length velocity in the pooled sample indicating faster length change with increasing GA.

Of three previous studies investigating associations between sociodemographic predictors and infant growth before 6 months, two studies found a significant and fully adjusted positive association between length (linear) growth and maternal education, and used as a proxy for SEP. Only one study found an inverse association with length growth, which was close to null on adjustment. Many studies including age ranges exceeding 6 months of age up to 5 years of age demonstrated a positive association between maternal education and length/height growth. The majority of these studies support the conclusion that lower SEP is associated with slower length (linear) growth in infancy and early childhood.

Several prior studies representing high-income European countries have noted that their findings of no evidence of a relationship between SEP and length (linear) growth prior to 6 months may not generalise to low-income to middle-income countries. Deviations from the Western diet and lifestyle were one of the reasons given for this limitation. Chile, the country from which our data were collected, offers an interesting context in this respect. The recruitment period for this study, 1991–1996, occurred as Chile was transitioning from a low-income to an upper-middle-income country. In 1990, 40% of the Chilean population was below the poverty line; by 2012, WHO classified Chile as an upper-middle-income country. There were nutrition and epidemiological transitions beginning in the 1970s and continuing during the 1990s when study infants were enrolled. Specifically, consumption of high-calorie food, accompanied by a sedentary lifestyle, resulted in rising obesity prevalence across all socioeconomic levels. In the context of an emerging western diet and lifestyle, we found that lower SEP was associated with poorer length (linear) growth in early infancy. Of course, contemporary generations born 20 years ago.

Table 4: Sociodemographic predictors and association with weight-for-length (WFL) Superimposition by Translation and Rotation growth parameters stratified by sex of child in the Santiago Longitudinal Study, 1991–1996

| Characteristic | Males | | | Females | | | Both | | |
|---|---|---|---|---|---|---|---|---|---|
| | Unadjusted | Adjusted ‡ | Unadjusted | Adjusted ‡ | Unadjusted | Adjusted ‡ | Unadjusted | Adjusted ‡ |
| Gest age | 0.09 (-0.55 to 0.73) | -2.03 (−2.91 to −1.15) | NA | -1.58 (−2.90 to −1.11) | 0.05 (-0.58 to 0.69) | -2.34 (−3.35 to −1.32) | NA | -1.33 (−2.04 to −0.52) | 0.07 (-0.38 to 0.52) | -2.17 (−2.84 to −1.51) | NA | -1.99 (−2.83 to −1.49) |
| Maternal age | 0.07 (-0.03 to 0.18) | -0.09 (-0.23 to 0.06) | 0.04 (-0.24 to 0.16) | -0.08 (-0.36 to 0.17) | 0.02 (-0.09 to 0.13) | -0.18 (-0.36 to 0.14) | NA | -0.13 (-0.36 to 0.14) | 0.05 (-0.03 to 0.12) | -0.13 (-0.24 to 0.02) | NA | -0.11 (-0.22 to 0.03) |
| Maternal education | -0.09 (-0.35 to 0.16) | 0.08 (-0.27 to 0.44) | NA | NA | -0.10 (-0.35 to 0.14) | 0.00 (-0.40 to 0.40) | NA | 0.07 (-2.11 to 0.42) | -0.10 (-0.28 to 0.31) | 0.04 (-0.22 to 0.31) | NA | NA |
| Graffar Index§ | -0.08 (-0.18 to 0.02) | -0.07 (-0.21 to 0.07) | -0.05 (-0.24 to 0.15) | -0.08 (-0.24 to 0.17) | 0.08 (-0.02 to 0.19) | 0.26 (0.10 to 0.43) | 0.04 (-0.21 to 0.18) | 0.25 (0.05 to 0.42) | -0.01 (-0.08 to 0.07) | 0.08 (-0.03 to 0.19) | NA | 0.06 (-0.14 to 0.17) |

*Size units are percentage change in log(WFL) from average, tempo units are time (days) and velocity units in cent per change from average.
†Bold values indicate significance with Bonferroni correction at alpha level of 0.05.
‡Adjusted linear regression models only include non-zero coefficients from lasso regression models that include all covariates in full model.
NA indicates the variable is not included in the adjusted analysis.
§Higher Graffar Index values indicate lower socioeconomic status.
growth parameter for weight. This was similarly reported in another cohort from the same geographic area of Santiago, Chile, the Growth and Obesity Cohort Study, which started a decade later and studied ages between birth and 2 years. Our findings add to this work. Through our intense focus on the first 5 postnatal months, our results demonstrate that the association between SEP and weight growth appears earlier in the postnatal period than previously documented.

Other potential mechanisms relating to SEP could include gestational weight gain and maternal nutrient status. Size at birth, considered a proxy for these two factors and represented in these analyses by the size SITAR parameter, was not associated with any of the sociodemographic measures. Further research will be useful in clarifying the biological mechanisms behind the association between SEP and early infant growth.

Strengths of this study include the combination of an analytical approach to growth that better captures the non-linear characteristic of growth in the first 5 months of life with a detailed measure of SEP appropriate to the context of a lower income setting. Another strength is the monthly anthropometric measures collected in the first 5 postnatal months. We also note several limitations. The sample size (n=1412) is smaller than other studies with sample sizes in the thousands or tens of thousands. Our study, therefore, may not have been powered to detect some effects reported in larger studies. Another limitation is that the Graffar Index, developed to assess differences in low-income to middle-income populations, limits the generalisability of our findings to higher income groups.

This investigation examined various growth characteristics from birth to 5 months and their association with sociodemographic factors in a Chilean infancy cohort. We found associations between lower SEP and slower length (linear) growth, which are similar in direction to previous findings for maternal education that span periods of time greater than the first 6 months and up to 5 years of age. The association between maternal age and weight size, in our study, was similar to findings in other studies of growth between birth and 2 years of age. In sum, our results extend findings from previous research by showing that sociodemographic factors affect infant growth even in the first 5 months of growth and in relatively homogenous low-income to middle-income populations.

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Contributors AVH designed the study, conducted the data analysis and wrote the first draft of the paper. KEN and SG supervised and contributed to the study design, interpretation of results and draft revisions. EB helped acquire the data. All authors contributed to revisions of the draft for intellectual content and approved the final version of the manuscript.

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Ethics approval The Santiago Longitudinal Study had been approved by Institutional Review Boards from (1) University of Michigan Medical Center, Ann Arbor, (2) Institute of Nutrition and Food Technology, Chile and (3) University of California, San Diego. The Office of Human Research Ethics at the University of North Carolina, Chapel Hill exempted this current research using existing anonymous data from review under the 45 CFR 46.101(b) regulatory category.

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