

# Nutritional Status at 2 Years in Former Infants with Bronchopulmonary Dysplasia Influences Nutrition and Pulmonary Outcomes During Childhood

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**ABSTRACT:** Improved survival rates for extreme prematurity have been accompanied by an increase in the incidence of bronchopulmonary dysplasia (BPD). The objective of this study was to assess factors associated with long-term nutritional and pulmonary function outcomes. The study was a cross-sectional study of 52 children who had been born prematurely, had experienced BPD, and were 4–8 y old at the time of the study. Undernutrition was defined as a Z score for weight-for-height of  $<-2$  SD. Body composition and lung function were evaluated. Resting energy expenditure (REE) was measured using indirect calorimetry. Stepwise logistic regression was used to test for factors associated with undernutrition and pulmonary function. Eighteen children (35%) with BPD, predominantly girls, were undernourished. Undernutrition occurred within the first months of life and was associated with high REE. Multivariate analysis showed that factors significantly associated with undernutrition were female sex and undernutrition at age 2 y. Thirty-one children (60%) had abnormal lung function tests. Multivariate analysis showed that undernutrition at age 2 y was the only factor associated with the risk of developing distension of the airways. Nutritional status at age 2 y in children who had BPD in infancy influences nutritional and pulmonary outcomes in childhood. (*Pediatr Res* 60: 340–344, 2006)

Northway *et al.* (1) first identified BPD in preterm children with severe respiratory disease treated with mechanical ventilation in 1967. In the past decade, significant advances in the treatment of prematurity have contributed to an increased survival of extremely premature infants (2). The incidence of BPD has decreased in preterm infants born after 28 wk PMA with a birth weight exceeding 1200 g, but still occurs frequently in extremely preterm children (2). Progress in neonatal intensive care has improved the survival rates for extreme prematurity, but has also resulted in a concomitant increase in the incidence of BPD (2). Children with BPD often experience growth failure and chronic lung disease during their first 2 y of life (2). BPD (oxygen requirement at 28 d or 36 wk PMA) is considered a good predictor of rehospitalization for respira-

tory causes and the use of pulmonary medications after discharge during the first years of life (2).

Several studies have reported nutritional or pulmonary complications of BPD early in life, but little is known about these factors during long-term follow-up. To our knowledge, no study has assessed both nutritional and pulmonary outcomes in later childhood in children having BPD in infancy. We hypothesized that early life events during the neonatal period could influence long-term outcomes of children with BPD, in particular the fixed pulmonary sequelae and undernutrition in later childhood. The objective of this study was to identify early risk factors associated with nutritional and respiratory outcomes in later childhood in children who experienced BPD as infants.

## SUBJECTS AND METHODS

Children were selected from a single-center population of children with BPD born between 1994 and 1999. The study was performed from April 2002 to April 2003. Inclusion criteria were prematurity, BPD, and age 4–8 y at the time of the study. Patients included 24–32 wk PMA premature infants. BPD was defined as an oxygen ( $>21\%$ ) requirement for at least 28 d as oxygen therapy in babies born up to 36 wk PMA or babies born past 36 wk PMA discharged to home (2). Exclusion criteria were a recent acute infection lasting for  $<15$  d, parental or patient refusal to participate in the study, and any malformation known to interfere with nutritional status (*e.g.* diaphragmatic hernia, congenital heart disease, and genetic disease). A sample of 52 children (22 girls, 30 boys) was selected out of 195 eligible patients. One hundred and forty-three eligible children were not included: 42 refused to participate, 60 were lost to follow-up, 5 moved to another region, 15 died, and 21 had associated malformations (16 diaphragmatic hernia, 5 cardiologic malformations).

The perinatal characteristics were obtained by retrospective analysis of their neonatal files (gestational age, birth weight, length of the period of ventilation, length of oxygen therapy, patent ductus arteriosus, necrotizing enterocolitis, gastroesophageal reflux, intrauterine growth restriction, chorioamnionitis, duration of hospitalization, weight at discharge and at term, intraventricular hemorrhage, neurologic status at 2 y of age, and tracheal stenosis). Socioeconomic status was taken as the level of education of the mother. Height and weight growth curves from birth to age 4–8 y were also obtained. We carefully checked the medical charts of the 143 nonincluded children to confirm that they did not differ from the 52 children studied on

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**Abbreviations:** BMD, bone mineral density; BPD, bronchopulmonary dysplasia; FFM, fat-free mass; FM, fat mass; FRC, functional residual capacity; PMA, postmenstrual age; R, resistance; REE, resting energy expenditure;  $V_{CO_2}$ , expired carbon dioxide flow;  $V_{O_2}$ , inspired oxygen flow

socioeconomic level, neonatal characteristics (intrauterine growth restriction, gestational age, birth weight, chorioamnionitis), neurologic status (intraventricular hemorrhage and cerebral palsy at 2 y of age), use of corticosteroid in the neonatal period, and initial evolution (oxygen therapy, mechanical ventilation, duration of hospitalization, weight at discharge, patent ductus arteriosus, first oral feeding, gastroesophageal reflux, tracheal stenosis, esophagitis, and neonatal ileus).

Before the study began, the purpose and objectives were carefully explained to the participants (children and parents). Written informed consent was obtained from parents and from children when appropriate. All procedures were performed in accordance with the ethical standards of the Helsinki Declaration of 1975, as revised in 1983, and French Good Clinical Practice (1987). This study was approved by the local ethical committee (Comité de Protection des Personnes pour la Recherche Biomédicale).

Nutritional status was evaluated using auxological characteristics (age, weight, height, head circumference) and dual x-ray absorptiometry (DXA). Body weight (W) and height (H) were measured without shoes and with light indoor clothing. A standard precision hospital scale was used for all measurements of body weight. Height was measured with a wall-mounted stadiometer. Body mass index (BMI) was calculated as weight (in kilograms) divided by height (in meters) squared. Growth measurements were converted to Z scores relative to the French growth references of Sempé *et al.* (3). Z scores were computed by subtracting the predicted value for the child's age (for Z scores reported by age) or for the child's height (for Z scores for W/H) and sex from the child's measurement, and then dividing the difference by the SD of the measurement in the reference group as follows:

$$Z = (\text{observed} - \text{predicted})/SD$$

The expected value for the measurement is 0, and 2 SD above this value estimates the 98th percentile and 2 SD below estimates the 2nd percentile. Height and weight were corrected for gestational age until the age of 2 y. Children were divided in two groups according to their nutritional status. Undernutrition was defined as a Z score for weight for height 2 SD below the mean.

The DXA instrument was a Lunar DPX-IQ "pencil-beam" (Lunar Radiation Corporation, Madison, WI). The whole-body scan time was 560–590 s, and the radiation dose was <0.3  $\mu$ Sv. A tissue bar was scanned as the reference material. Total body image acquisition and analysis were obtained following the manufacturer's instructions. FM and FFM were calculated using the pediatric software provided by the manufacturer. BMD was also calculated, and the reference values used in the calculation of Z scores for BMD were from an American source provided with the software.

REE was measured by indirect calorimetry, using the Deltatrac II open-circuit ventilated hood system (Datex Instrumentation Corporation, Helsinki, Finland). The respiration quotient and flow settings were calibrated by reference to alcohol combustion every 6 mo and with a reference gas mixture (95% O<sub>2</sub>, 5% CO<sub>2</sub>) before each measurement. Vo<sub>2</sub>, Vco<sub>2</sub>, and respiration quotient were calculated. REE was calculated every minute from oxygen consumption (Vo<sub>2</sub> in milliliters per minute) and production of carbon dioxide (Vco<sub>2</sub> in milliliters per minute) using the Weir formula without protein correction (4):

$$REE \text{ (kcal/h)} = (3.9 \times Vo_2 + 1.1 \times Vco_2) \times 1.44/24$$

REE was measured after a fast of 10–12 h. After an adaptation period of 15 min under a transparent canopy, continuous respiratory exchange measurements were initiated (5). REE was measured for 30 min.

Parents performed an alimentary inventory using a 7-d food questionnaire. Data were analyzed with BILNUT 3 software (Nutrisoft, Cerelles, France).

All pulmonary function tests were performed after bronchodilators had been stopped for at least 2 wk. Airway resistance (R) was measured by interruption of the airflow (software Dyn'R, Paris, France); seven measurements were performed. A variation <14% was needed to validate the results. FRC was measured by the dilution of helium according to American Thoracic Society standards (Medisoft, Brussels, Belgium); this technique required two measurements with a variation of <10% to validate the results. The spirometry–flow volume curve, area under the curve, and peak flow were evaluated. Five reproducible spirometry–flow volume curves (<10%) were necessary to validate the data.

Values are expressed as the percentage of the predicted value normalized for height and sex. R < 150%, FRC < 120%, and peak flow > 120% were considered within the normal ranges. Obstruction airflow was defined as R > 150%, and distension was defined as FRC > 120%. We assumed a maximal expiratory flow–volume loop with a marked concavity to demonstrate distal flow limitation during spirometry. Measurements were repeated 20 min after 200 mg of inhaled salbutamol. Reversibility of obstruction and hyperinflation

was defined as a decrease of 10% of the FRC and 30% of the R. In the spirometry tests, reversibility was positive when the concavity disappeared and an increase of >30% was observed in the area under the curve.

Univariate analyses were first performed to check and summarize the data (box plots, means, SD and percentiles for numerical parameters, and frequencies for qualitative parameters). The population was then divided into two groups: undernourished children (Z W/H < -2 SD) and children with normal nutritional status (Z W/H  $\geq$  -2 SD). The population was also classified according to its pulmonary status (obstruction *versus* no obstruction of the airways; hyperinflation *versus* no hyperinflation of the airways). Bivariate analysis was then performed to compare the groups. The Wilcoxon test was used for numerical parameters and the  $\chi^2$  test or Fisher exact test, if needed, was used for qualitative parameters.

Finally, parameters that were significant at the 15% level in the bivariate analysis were entered into a stepwise logistic regression analysis to evaluate the most significant factors associated with undernutrition and abnormal pulmonary function. The evolution of the Z score from term to 4 y of age was assessed using nonparametric analysis of variance for repeated measurements. All statistical analysis was performed with the SAS Software. The significance level was set at 0.05.

## RESULTS

At the time of the study, 18 children with BPD were undernourished (35%). Table 1 details the auxological characteristics and body composition of the population. Girls were more frequently undernourished than were boys (12 girls and 6 boys,  $p = 0.009$ ). FM, FFM, and ZBMD were significantly lower in the undernourished group than in the normally nourished group (Table 1).

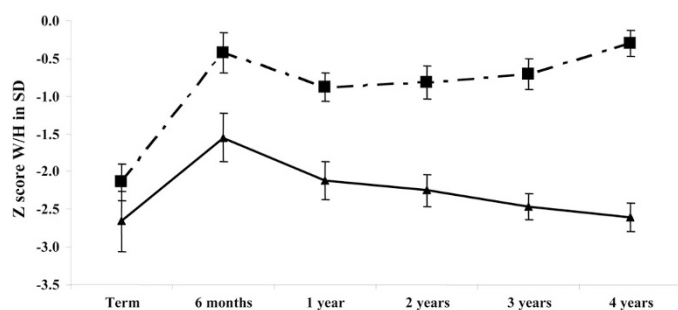
The Z scores of W/H were calculated at term, and at ages 6 mo, 12 mo, 24 mo, 3 y, and 4 y. The nonparametric analysis of variance for repeated measurements showed that Z scores were significantly lower at any age except term in the undernourished children compared with the normally nourished children (Fig. 1).

When expressed as an absolute value, REE was significantly lower in the undernourished group (857  $\pm$  94 kcal/d) than in the normally nourished group (971  $\pm$  90 kcal/d,  $p < 0.01$ ). However, because undernourished children have lower body weight and FFM, the REE expressed per kilogram of body weight or per kilogram of FFM was significantly higher in the undernourished group. REE per kilogram of body weight was 60  $\pm$  8 kcal/kg/d in undernourished children and

**Table 1.** Body weight and body composition of undernourished and normally nourished children at the age of 4–8 y

	Normal nutritional status (n = 34)	Undernutrition (n = 18)	p Value*
Age (y)	6 $\pm$ 1	6 $\pm$ 1	0.9
Weight (kg)	19 $\pm$ 3	15 $\pm$ 2	0.04
Height (m)	112 $\pm$ 7	109 $\pm$ 7	0.4
HC (cm)	51 $\pm$ 2	48 $\pm$ 2	0.05
BMI (kg/m <sup>2</sup> )	15 $\pm$ 2	12 $\pm$ 1	<0.001
Z Weight/age (SD)	-0.2 $\pm$ 1.1	-2.1 $\pm$ 0.8	<0.001
Z Height/age (SD)	-0.2 $\pm$ 1.1	-0.9 $\pm$ 1.3	0.4
Z Weight/height (SD)	-0.3 $\pm$ 0.2	-2.6 $\pm$ 1	<0.001
Z HC	-0.6 $\pm$ 1.2	-1.9 $\pm$ 1.7	0.04
FFM (kg)	15.5 $\pm$ 2	13.4 $\pm$ 2	0.04
FM (% of weight)	16 $\pm$ 8	11 $\pm$ 5	<0.001
Z BMD	0.4 $\pm$ 0.7	-0.3 $\pm$ 1.0	0.01

\* Difference between undernourished and normally nourished children (mean  $\pm$  SD).



**Figure 1.** Differences in Z scores of weight/height (SD) between undernourished (▲) and normally nourished (■) children from term to 4 y ( $n = 52$ ). Time effect:  $p < 0.0001$ ; group effect:  $p < 0.0001$ ; interaction:  $p < 0.0001$ .

49 ± 6 kcal/kg/d in normally nourished children ( $p = 0.01$ ). REE per kilogram of FFM was 78 ± 10 kcal/kg FFM/d in undernourished children and 57 ± 9 kcal/kg FFM/d in normally nourished children ( $p = 0.01$ ). The energy intake and carbohydrate intake were significantly higher in the undernourished group (2033 ± 660 kcal/d) than in the normally nourished group (1800 ± 568,  $p = 0.01$ ). Carbohydrate intake as a percentage of total energy intake was 59% ± 7% in the undernourished children and 49% ± 7% in the normally nourished children ( $p = 0.02$ ). Energy intake remained significantly higher in the undernourished group than in the children with normal nutritional status when expressed relative to weight (140 ± 46 kcal/kg/d versus 104 ± 31 kcal/kg/d,  $p = 0.01$ ) or relative to FFM (155 ± 52 kcal/kg FFM/d versus 103 ± 45 kcal/kg FFM/d,  $p = 0.04$ ).

Thirty-one children (60%) presented with abnormal lung function. At the time of the study, only one child was receiving inhaled steroids (fluticasone, 100 µg/d) on a regular basis. None of the patients used bronchodilators at the time of the study or on a regular basis. Obstruction was present in 16 patients and was reversible in 7 children. Distension was diagnosed in 14 patients and was reversible in 4. Forty-five patients cooperated fully in the spirometry tests: marked concavity was present in 21 of the 45 expiratory flow curves and reversibility was observed in 9 children. Five children had distension associated with obstruction (both distal and proximal), six had distension associated with distal obstruction, and six had both distal and proximal obstruction. Four of 14 patients with hyperinflation exhibited reversibility after salbutamol. Reversibility after salbutamol was also observed in 7 of 16 children with proximal obstruction and in 9 of 21 with distal obstruction. Pulmonary function test results did not differ significantly between the undernourished children and those with normal nutritional status (Table 2). However, the group with distal obstruction had significantly lower FFM (87% versus 93% of FFM,  $p = 0.05$ ) and Z scores for BMD ( $-0.2 ± 0.7$  versus  $0.1 ± 0.9$ ,  $p = 0.03$ ). REE was significantly higher in children with distal obstruction (970 ± 100 kcal/d versus 933 ± 90 kcal/d,  $p = 0.03$ ). In addition, distal obstruction and hyperinflation of the airways were significantly associated with intrauterine growth restriction (11/21 versus 8/21,  $p = 0.04$ ). The Z scores for W/H were significantly lower at the age of 2 y in children who exhibited

**Table 2.** Lung function tests (% of theoretic values)

	Normal nutritional status ( $n = 34$ )	Undernutrition ( $n = 18$ )	$p$ Value*
FRC	113 ± 26	110 ± 17	0.8
FRC (post $\beta_2$ )	102 ± 17	100 ± 15	0.8
R	147 ± 37	130 ± 31	0.1
R (post $\beta_2$ )	124 ± 29	113 ± 33	0.8
Area (1)	1.2 ± 0.6	1.3 ± 0.7	0.1
Area (post $\beta_2$ ) (2)	1.7 ± 0.7	1.6 ± 0.7	0.1
Area 1/2	137 ± 32	132 ± 29	0.7
Peak-flow	84 ± 21	86 ± 24	0.8
Peak-flow (post $\beta_2$ )	96 ± 20	101 ± 19	0.6

R, resistance; Area, area under the spirometric curve ( $L^2/s$ );  $\beta_2$ , inhaled salbutamol.

\* Difference between undernourished and normally nourished groups.

hyperinflation ( $-1.6 ± 1.7$  SD versus  $-1.0 ± 0.8$  SD,  $p = 0.05$ ) or distal obstruction of the airways ( $-1.7 ± 1.4$  SD versus  $-1.0 ± 0.9$  SD,  $p = 0.02$ ).

Lower gestational age was the only factor associated with undernutrition (Table 3). Several neonatal characteristics were significantly associated with abnormal pulmonary function tests: antenatal growth restriction, gestational age, and being a twin. Twins presented a significantly higher risk for developing pulmonary hyperinflation ( $p = 0.01$ ). Antenatal growth restriction was associated with hyperinflation of the airways ( $p = 0.02$ ). Gestational age at birth was associated with a higher risk of developing hyperinflation in later childhood (29 ± 2 wk PMA versus 27 ± 1.6 wk PMA,  $p = 0.006$ ).

Bivariate analysis showed that only four parameters were associated with undernutrition: sex, Z score for head circumference, gestational age at birth, and Z score for W/H at 2 y of age. A stepwise logistic regression performed on those four

**Table 3.** Neonatal characteristics (mean ± SD)

	Normal nutritional status ( $n = 34$ )	Undernutrition ( $n = 18$ )	$p$ Value*
Birth weight (g)	1047 ± 309	993 ± 309	0.5
Gestational age (wk)	28.6 ± 2	27.5 ± 1	0.05
Weight at discharge (g)	2742 ± 629	2889 ± 900	0.1
Weight at term (g)	2422 ± 488	2124 ± 609	0.06
Hospitalization duration (d)	116 ± 103	144 ± 124	0.1
Total ventilation (d)	45 ± 93	45 ± 41	0.8
Tracheal intubation (d)	38 ± 37	37 ± 37	0.7
Continuous positive airway pressure ventilation (d)	8 ± 10	8 ± 11	0.8
Oxygen therapy duration (d)	150 ± 36	169 ± 37	0.9
First oral feeding (d)	24 ± 13	21 ± 13	0.9
Intrauterine growth restriction ( $n$ )	12	7	0.1
Maternal-fetal infection ( $n$ )	4	6	0.7
Twins ( $n$ )	7	3	0.8
Gastroesophageal reflux ( $n$ )	32	15	0.3
Esophagitis ( $n$ )	2	2	0.6
Neonatal ileus ( $n$ )	14	7	0.8
Patent ductus arteriosus ( $n$ )	26	10	0.1
Intraventricular hemorrhage ( $n$ )	18	6	0.4
Neurological abnormalities at 2 y of age ( $n$ )	13	4	0.07

\* Difference between undernourished and normally nourished groups.

parameters showed that only two independent factors were significantly associated with undernutrition: under nutrition at the age of 2 y [ $p < 0.001$ ; odds ratio (OR) = 8.9; 95% confidence interval (CI), 2.2–36.4] and female sex [ $p = 0.03$ ; OR = 4.5 (95% CI, 1.1–18.3)].

Bivariate analysis showed that only four parameters were associated with abnormal pulmonary function test: gestational age at birth, Z score for W/H at 2 y of age, antenatal growth retardation, and twin birth). A stepwise logistic regression performed on those four parameters showed that the only independent factor significantly associated with hyperinflation of the airways was undernutrition at the age of 2 y [ $p = 0.05$ ; OR = 1.2 (95% CI, 1.1–3.0)].

## DISCUSSION

Rapid progress in the management of prematurity has contributed to an increase in survival of extremely premature infants (6,7). These significant advances, especially the use of antenatal steroids and continuous positive airway pressure ventilation, have influenced pulmonary outcome and tracheal sequelae (6,7). Although our study suggests that nutritional outcomes in infancy have not changed, undernutrition remains a frequent feature in the preterm population. Considering the recent advances in neonatal care, it is difficult to compare current results in children with BPD with previously published studies. A few studies have reported nutritional status during long-term follow-up of patients with BPD (8–10), although most were published more than 6 y ago and therefore address historical forms of BPD. In preterm infants without BPD, catch up of height and weight usually occurs before the age of 2 y and only 5–10% present with growth retardation after the age of 2 y (11–13). Conversely, preterm children who develop BPD usually show impaired growth early in life, which persists for years (2,7,9). Our study shows that undernutrition involves different body compartments (*e.g.* FM, FFM, BMD) and is associated with a smaller head circumference compared with normally nourished children. Undernutrition is also associated with elevated REE, despite the high energy intake. Our results demonstrating that long-term undernutrition originates from the first months of life suggest that the neonatal period is crucial for the future nutritional status of infants with BPD.

One surprising result was that girls were undernourished more frequently than were boys. This difference in nutritional status between girls and boys has not been reported previously. However, our results are consistent with two previous randomized trials on nutritional intervention in preterm infants that showed a sex difference in the effect of diet, which was greater in boys than in girls (12,13). Cooke *et al.* (12) examined the changes over the whole body during dietary intervention in preterm infants at six and 12 mo after hospital discharge, and found that only boys gained significant body weight. Carver *et al.* (13) studied a group of 125 preterm infants receiving a nutrient-enriched formula at six and 12 mo after discharge. The nutrient-enriched formula increased growth mainly in children with birth weight <1250 g and in boys (13). These two studies suggest that the sex of

preterm infants influences the postdischarge responsiveness to nutrient supplementation. We have no clear explanation for such a difference.

Lung sequelae are frequent events in children with BPD (14), who usually experience pulmonary infections and hyper-reactivity until the age of 2 y (14). Lung pulmonary tests at school age usually demonstrate obstruction, hyperinflation, and hyperreactivity of the airways (14–16), and 60% of the children in our study demonstrated one or more of these dysfunctions. Hyperinflation seems to be related to a fixed lesion that does not progress with time and reflects gas trapping in the distal airways (14). Hyperinflation of the airways is a more specific sequel of BPD than obstruction of the airways and could be related to asthma (14).

The main objective of our study was to identify early risk factors in infancy for undernutrition or lung sequelae in later childhood. The cross-sectional study design could show only an association of early neonatal events with nutritional and pulmonary status between the ages of 4 and 8 y, and cannot assess any causality underlying this relationship. Multivariate analysis showed that undernutrition at 4–8 y of age was not associated with neonatal events. Our results did not confirm those of previous studies showing that neonatal factors such as patent ductus arteriosus, chorioamnionitis, gastroesophageal reflux, delayed oral nutrition, prolonged hospitalization, or cerebral abnormalities (cerebral palsy or periventricular leukomalacia) contribute to growth failure (15,16). However, the earlier studies were performed only during the first months of life and did not relate to growth velocity in later childhood.

A number of neonatal factors, such as baro- and volutrauma, oxygen toxicity, and prolonged hospitalization can affect the immature pulmonary systems of preterm newborn infants (6), but were not associated with lung sequelae in later childhood in our study. One limitation of our study is that only 25% of the eligible population with BPD was studied because we could not obtain information on neonatal characteristics, neurologic status, steroid use during the neonatal period, and initial evolution in neonatal care unit of all 195 children. Although we cannot exclude the possibility of a selection bias, the children who were not included did not differ from the 52 children studied in regards to the severity of their disease and neonatal characteristics. We tried to limit all other bias by studying a single-center cohort treated by the same experienced team of neonatologists.

Family history of atopy, childhood respiratory illness, passive exposure to cigarette smoke, steroid therapy, and recurrent infection are important confounding factors for lung function or growth in children (15,16). We included only early postnatal systemic steroids, but this was not significantly associated with long-term pulmonary outcome. Socioeconomic status can also influence growth (15). We evaluated socioeconomic status by the level of maternal education, but found no significant difference between the groups.

The most interesting finding is that undernutrition before 2 y of age is an independent factor associated with both hyperinflation of the airways and undernutrition in later childhood, despite the lack of a direct relationship between pulmonary function and nutritional status at age 4–8 y. Our data

suggest that undernutrition in later childhood does not result from low energy intake but from an abnormal energy balance and a persistent increase in REE. REE was also positively associated with abnormal pulmonary function (hyperinflation or obstruction of the airways) at age 4–8 y. REE is associated with respiratory status because of an increased oxygen consumption of respiratory muscles secondary to an increased resistive load in the neonatal period (17). Our data show that this pattern persists at the long-term follow-up and may explain the persistence of undernutrition with time.

Our data also show that undernutrition is not correlated with pulmonary status in later childhood, but that both are associated with nutritional status before the age of 2 y. After this period, undernutrition and hyperinflation of the airways seem to be fixed sequelae and independent features. In children who had BPD in infancy, nutritional status at 2 y of age could influence both nutrition and pulmonary outcomes in childhood. If this hypothesis is confirmed, early randomized nutritional intervention should be performed before the age of 2 y to prevent lung sequelae and undernutrition in later childhood.

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