Cross-sectional and longitudinal associations between energy intake and BMI z-score in European children

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Abstract

Background: Evidence for the effect of dietary energy on BMI z-scores in young children is limited. We aim to investigate cross-sectional and longitudinal effects of daily energy intake (EI) on BMI z-scores of European boys and girls considering growth-related height dependencies of EI using residual EI.

Methods: To investigate cross-sectional and longitudinal effects of daily energy intake (EI) on BMI z-scores of European boys and girls considering growth-related height dependencies of EI using residual EI.

Methods: Subjects were children aged 2-<10 y old (N = 2753, 48.2% girls) participating in the IDEFICS (Identification and prevention of Dietary- and lifestyle-induced health EfFects In Children and infantS) baseline and follow-up examination. Usual EI (kcal/day) was calculated based on the National Cancer Institute-method excluding subjects with implausible reported EI. Effect of age, height and sex-adjusted residuals of EI on BMI z-score was investigated stratified by baseline age-group (2-<4 y, 4-<6 y, 6-<8 y and 8-<10 y) cross-sectionally using linear regression models adjusted for relevant confounders (crude model: age, sex, country; fully adjusted model: plus parental ISCED level, parental BMI, screen time; subgroup analysis: plus objectively measured physical activity). Longitudinal associations were estimated between changes in (Δ) residual EI per year and ΔBMI z-score per year with adjustments analogously to the cross-sectional models but with additional adjustment for residual EI at baseline.

Results: Cross-sectionally, positive associations were observed between residual EI and BMI z-score for the full study sample, for boys and in older (≥6 years) but not in younger children in the crude and fully adjusted model. Longitudinally, small positive associations were observed between Δresidual EI per y on ΔBMI z-score per y for the full study sample and in 4-<6 y olds in the crude and fully adjusted model.

Conclusion: In conclusion, EI above the average intakes for a certain sex, age and height are weakly associated with BMI z-scores in European children. Residual EI may be considered as a useful exposure measure in children as it accounts for growth-related changes in usual EI during childhood.

Keywords: Residual energy intake, Europe, Cohort, BMI z-score, Children

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Background
The worldwide prevalence of overweight and obesity in pre-school children and school-aged children has become a major public health concern [1]. Recently the IDEFICS (Identification and Prevention of Dietary- and Lifestyle-induced Health Effects In Children and Infants) study reported an obesity prevalence of 7% and an overweight prevalence of 13% among European children under 10 y of age [2]. Childhood overweight is a likely consequence of the modern environment and lifestyle, characterized by increasing sedentary activities and less physical activity (PA) as well as higher energy intakes (EI) [3, 4]. Cross-sectional studies in children and adolescents found associations between EI and Body Mass Index (BMI) z-score [5, 6].

Data on longitudinal associations between energy intake and weight development in pre-school children and school children are scarce and inconsistent. Total EI (kcal) was found to be associated with weight gain in 3 to 5 y old U.S. children [7], whereas EI was unrelated to BMI in two other U.S. studies among 8 to 14 y olds [8] and ~11 y old adolescents [9]. Associations between EI and weight change were observed in Italian children aged 8 y at baseline and 14 y at follow-up, but associations disappeared after adjusting for parental obesity [10]. In a Danish study, EI was related to weight gain in 8 - 10 y old overweight boys, but not in girls [11]. In adolescents EI was found to be influenced by PA, whereas PA was associated with lower body fat mass/BMI. This indicates that energy expenditure through PA counteracts greater fat mass/BMI while EI is high [8, 12]. Studying associations between reported energy intake and health outcomes is challenging, especially in children. Measurement errors resulting e.g. portion size estimation, erroneous food composition tables, incomplete recalls, misreporting or daily variations in intake pose a great problem to nutritional epidemiologists [13]. In children, dietary assessment is even more complicated as data typically rely on proxy-reporters. In addition, EI in childhood and youth is influenced by energy demands and varies over time depending on growth [14] such that absolute EI values cannot be directly compared between children of different ages.

In the light of the above and knowing that dietary habits may track from childhood into adulthood [15, 16] it is important to understand to which extent early EI is associated with changes in BMI among pre-school children. The present study aims to investigate cross-sectional and longitudinal associations between usual daily EI and (changes in) BMI z-scores of European boys and girls, accounting for growth-related changes in usual EI during childhood and trying to address the abovementioned methodological challenges.

Methods
Data for the present cross-sectional and longitudinal study were obtained from the IDEFICS (Identification and prevention of Dietary- and lifestyle-induced health Effects In Children and infantS) Study baseline examination (at time T0) in 2007/2008, and from the follow-up survey (T1) in 2009/2010.

Study participants
The IDEFICS study was designed as a prospective cohort study. Children from Sweden, Germany, Hungary, Italy, Cyprus, Spain, Belgium, Estonia aged 2 to <10 y who attended selected pre-schools or kindergartens and primary schools (grades 1 and 2) were eligible for participation [17]. The children were approached via schools and kindergartens to facilitate equal enrollment of all social groups. In addition to the signed informed consent given by parents, each child was asked to give verbal assent immediately before examination. Each participating center obtained ethical approval from the local responsible authorities in accordance with the ethical standards laid down in the 1964 Declaration of Helsinki and its later amendments. All 16,228 children in the defined age categories who fulfilled the inclusion criteria at baseline were invited for follow-up examination.

Questionnaires and anthropometric measurements
Questionnaires were developed in English, translated into local languages, and then back-translated to check for translation errors. Parents completed a questionnaire to assess - among others – parental BMI, behavioral factors and their determinants, socio-demographic characteristics and information on children’s total daily screen time (time of computer and/or TV use). For the present investigation, the highest educational level of the parents according to the International Standard Classification of Education (ISCED) [18] was used as a proxy indicator for socio-economic status (SES) of the family.

The field methods comprised anthropometric measurements of standing height (cm) using a Seca 225 stadiometer (Seca GmbH & KG, Birmingham, UK) in accordance with international standards for anthropometric assessment and weight (kg) [19]. Body weight was assessed in fasting children using a prototype of the TANITA BC 420 SMA digital scale (TANITA Europe GmbH, Sindelfingen, Germany) specifically adapted for children’s feet. All measurements were performed in light clothing (e.g. underwear). All anthropometric measurements followed detailed standard operation procedures [20].

BMI z-scores
BMI was calculated by dividing body weight in kilograms by squared body height in meters. BMI was transformed to an age- and sex-specific z-score according to Cole et
al. [21]. Weight groups (thin/normal and overweight/obese) were categorized using age- and sex-specific cut-off values based on the extended IOTF criteria [22]. The annual change in BMI z-score (∆BMI z-score) was calculated as BMI z-score at follow-up minus BMI z-score at baseline divided by the time span between baseline and follow-up:

\[
\Delta \text{BMI z-score} = \frac{(\text{BMI z-score T1} - \text{BMI z-score T0})}{(\text{age T1} - \text{age T0})}.
\]

Assessment of physical activity
Free-living PA was objectively assessed using Actigraph uniaxial accelerometers (either ActiTrainer or GT1M; Actigraph, LLC, Pensacola, FL, USA), containing both identical sensor units. The monitor was set to record PA in a 15-second sampling interval ("epoch"). Accelerometer measurements were included from children who wore the accelerometer for at least three days, with at least 6 hours per day, including one weekend day. Accelerometers were mounted on the right hip of each child secured by an elastic belt. The duration of moderate-to-vigorous physical activity (MVPA) was determined according to the cut-offs of Evenson [23] as described earlier [13]. We excluded 1221 subjects classified as under-reporters (656 at T0 and 565 at T1) and 462 over-reporters (261 at T0 and 201 at T1) from the analysis. After exclusion of misreports, individual daily energy intakes (EI) were estimated based on the National Cancer Institute Method [29] accounting for the daily variation in diet, and adjusting for weekend days/weekdays, interview sequence, age and sex. To correct for the daily variation of EI, all repeated 24HDR that were available for T0 and T1 examinations in a subgroup of children were used. Based on all IDEFICS children providing at least one plausible 24HDR, 2068 children at T0 and 903 children at T1, provided more than one 24HDR (up to 6).

Dietary information
Dietary intake of the previous 24 hours was assessed using the computer-assisted 24-hour dietary recalls (24HDR), called 'Self-Administered Children and Infant Nutrition Assessment’ (SACINA) [25]. The software was based on the YANA-C (‘Young adolescents’ nutrition assessment on computer’) system [26] developed within the HELENA Study (http://www.helenastudy.com). Proxy respondents were asked to recall the children’s diet and enter type and amount of all drinks and foods consumed during the previous day, starting with the first intake after waking up in the morning. Proxies were requested to complete repeated 24HDR, including two working days and one weekend day. Standardized photographs were used to assist accurate estimation of portion size and proxies could ask for assistance of the survey dietitian when completing the 24HDR. During the recall, impossible and implausible values (e.g. energy, amount (portion size), quantity (numbers of portions), missing meals) were checked for single food items and for single recalls and were reconfirmed with the participant directly. Further, the system checked for frequent and easily forgettable foods and food combinations. Meals, drinks and snacks consumed in school or kindergarten the day prior to the 24HDR were recorded using a standardized observer sheet, completed by trained personnel. Country-specific food composition tables (FCT) were used to match simple foods or European homogeneous multi-ingredient food items [25]. All nutrients and energy values were expressed per 100 g edible portion according to McCance and Widdowson [27]. The metabolizable energy values of foods and beverages were given in kilocalories (kcal).

The validity of proxy-reported energy intakes from the 24HDR was investigated based on comparison with total energy expenditure measured using the doubly labelled water (DLW) technique. The instrument was found to be valid to assess EI on group level rather than on individual level [28].

Dietary data analysis
Missing quantities or implausible values (above median + 2.5 standard deviation for single food items) that could not be corrected were imputed by country, food group and age-specific median intakes (0.01 % of the entries). Incomplete 24HDR and those with four or more imputed values were excluded from the analysis. The amount of all consumed foods and beverages was recorded in grams (g).

Age and sex-specific Goldberg cut-offs adapted for use in children were applied to classify subjects with under-reported, plausible reported and over reported EI as described earlier [13]. We excluded 1221 subjects classified as under-reporters (656 at T0 and 565 at T1) and 462 over-reporters (261 at T0 and 201 at T1) from the analysis.

After exclusion of misreports, individual daily energy intakes (EI) were estimated based on the National Cancer Institute Method [29] accounting for the daily variation in diet, and adjusting for weekend days/weekdays, interview sequence, age and sex. To correct for the daily variation of EI, all repeated 24HDR that were available for T0 and T1 examinations in a subgroup of children were used. Based on all IDEFICS children providing at least one plausible 24HDR, 2068 children at T0 and 903 children at T1, provided more than one 24HDR (up to 6).

Covariate information
Age, height, sex, country of residence and - according to previous literature - total screen time per week and physical activity [10, 30, 31], parental ISCED level [32], and parental BMI [33] and MVPA were considered being potential confounders in the present study. Total television (TV) and computer use time will hereafter be termed screen time. Where baseline covariate information was missing, but follow-up information available, the baseline value was imputed by the follow-up value so as not to further reduce the study sample.

Inclusion criteria for study sample
From 16228 children who participated in the IDEFICS baseline examinations 9590 children provided at least
one complete T0 24HDR. Among these, 3456 individuals also contributed at least one complete T1 24HDR, and 2785 had plausible reported EI values at T0 and T1. A total of 2753 children with complete co-variate information were eligible for the final analysis (study sample, see Fig. 1).

No substantial differences were observed when comparing main characteristics such as weight group, age and sex distribution or ISCED level of the baseline sample and the sample presented here (results not shown).

Statistical analysis

Usual energy requirements and hence energy intakes differ depending on age, height and sex. To facilitate comparability of EI values of children of different ages, sex, and heights, EI values need to be adjusted. For this purpose, a growth model for EI during childhood dependent on age, height and sex was estimated. Typically, EI increases strongly during early childhood, to a smaller extent in later childhood and settles down in adulthood. To account for this non-linearity, the following fractional polynomial model for repeated measures (using T0 and T1 data simultaneously) was selected based on the Akaike information criterion to model EI growth during childhood:

\[
EI_{ij} = \beta_0 + \beta_1 \text{age}_{ij} + \beta_2 \text{height}_{ij}^1 + \beta_3 \text{height}_{ij}^2
\]

\[
+ \beta_4 \text{height}_{ij}^3 + \beta_5 \text{sex}_i + \beta_6 \text{height}_{ij}^1 \times \text{sex}_i
\]

\[
+ \beta_7 \text{height}_{ij}^2 \times \text{sex}_i + \beta_8 \text{height}_{ij}^3 \times \text{sex}_i + \epsilon_{ij}
\]

(for individual i, i = 1,...,2757, at measurement occasion j, j = 1,2).

The residual EI \(\epsilon_{ij}\) indicates the subject-specific deviation from the average EI for a given age, height and sex as displayed in Fig. 2. For instance, a residual EI of 100 would indicate that this child consumed 100 kcal more than the average study population child with the same age, height and sex at this measurement occasion.

The estimated residuals for energy intake at T0 and T1 were used as exposure measures in the subsequent analyses. Residuals were rescaled such that one unit refers to 100 kcal in the regression models to achieve meaningful interpretations of effect estimates (1 unit ~ 100 kcal). In the longitudinal models, the annual change in residual EI (\(\Delta\text{residual EI}\)) between T0 and T1 (Fig. 2) was calculated as exposure measure, i.e. \(\Delta\text{residual EI} = (\text{residual EI T1} - \text{residual EI T0}) / (\text{age T1} - \text{age T0})\).

In the cross-sectional analysis, the association between residuals of EI and BMI z-score at baseline was estimated using linear regression analysis. All models were adjusted for baseline age, sex and country (model 1). In a second model, baseline values of highest parental ISCED level, parental BMI and average screen time per week were additionally included (fully adjusted model 2). In a subgroup analysis (N = 1933, model 3), objectively measured PA as duration of MVPA (min/day) was added to model 2. To evaluate the longitudinal effect of \(\Delta\text{residual EI}\) per year on \(\Delta\text{BMI z-score}\) per year, linear regression models were fitted analogue to the models 1, 2 and 3 with additional adjustment for BMI z-score and residual EI at baseline.

As we found significant interactions between age as well as sex and residual EI in the above models, all models were fitted for the total study group as well as stratified by sex and by baseline age group (2-<4 y, 4-<6 y, 6-<8 y and 8-<10 y), e.g. children who were 2-<4 y at baseline examination were assigned to the 2 to <4 y age group etc.

All analyses were performed using SAS® statistical software version 9.3 (SAS Institute, Inc., Cary, NC) using the procedure PROC MIXED; 99% confidence intervals are presented for \(\beta_{EI}\) estimates in all models.
Results

Study sample

The study sample consisted of 48.2% female participants; 67.2% were normal weight with the highest prevalence in 2-<4 y old children (75.1%) compared to other age groups. Children from the study sample were more likely to be Italian (37.5%), and less likely Belgian (3.3%, Table 1).

Baseline and follow-up characteristics

On average, boys had higher usual intakes of energy (kcal/day) than girls and older children had higher usual intakes than younger children at baseline and at follow-up (Table 2). For pre-school children and especially for pre-school boys (2-<4 y, 4-<6 y), change in daily EI was higher than for school age children (6-<8 y, 8-<10 y). Body height (cm) was slightly higher for girls than for boys at T0 and follow-up examination. Across age-groups, change in height between T0 and T1 was highest in children aged 2-<4 y at baseline and lowest in the oldest age group. Both boys and girls had a slight increase in relative BMI z-scores between baseline and follow-up examination, with boys having a higher increase than girls (+0.08 kg/m² and +0.05 kg/m² respectively). Screen time per week was higher for boys than for girls. Overall, mean parental BMI was similar for parents of boys and girls (24.20 kg/m²). Duration of MVPA per day was generally higher for boys than for girls. Baseline and follow-up characteristics of exposure and outcome variates differ for the subgroup of children (Additional file 1).

Cross-sectional effect of residuals of energy intake at baseline on baseline BMI z-score

For the total study sample, for boys as well as for 6-<8 y and 8-<10 y olds positive associations were observed between residuals of EI and baseline BMI z-score in the crude and adjusted models (models 1 and 2) and except for the 6-<8 y olds also in model 3 with additional adjustment for MVPA (Table 3). Association was strongest in 8-<10 y olds (ßEI 0.14, 99%CI 0.23 – 0.26, model 3). This indicates that a child consuming 100 kcal/day more than the average study population child with the same age, height and sex at baseline had a 0.14 kg/m² higher BMI z-score at baseline. No association was found for girls, and in the younger age groups (2-<4 y olds and 4-<6 y olds).

Further, adjusting for residing in the intervention vs. control region did not change findings (data not shown).

Longitudinal effect of residual change in energy intake per year on ΔBMI z-score per year

For the total study sample and for 4-<6 y olds positive associations were observed between Δresidual EI per y on ΔBMI z-score per y in the crude model 1 and the fully adjusted model 2 (Table 4). Associations were strongest in 4-<6 y olds. No association was found for girls, 2-<4 y olds, 6-<8 and 8-<10 y olds. Overall, size of the effect estimate was smaller longitudinally, whereas association was stronger cross-sectionally. Adjusting for residing in the intervention vs. control region did not change findings (data not shown).

In sensitivity analyses, the longitudinal associations were additionally estimated relating the (1) change in EI...
per cm increase in height (ΔEI/Δheight) and (2) change in usual EI (ΔEI) to ΔBMI z-score per y adjusting for baseline values of EI and BMI z-score. These exposure measures are easier to interpret, but less precise with respect to the age- and height dependencies of energy intake such that we finally decided to present the estimates based on residual changes. However, using ΔEI/Δheight or ΔEI as exposures, results were similar to those described above: Again significant positive associations were found for the total study group, in boys and in the group of 4- < 6 y olds.

**Discussion**

Cross-sectionally, we observed positive associations between residual EI and BMI z-score for the full study sample and in older (≥6 years) but not in younger children. Longitudinally, small positive associations were observed between Δresidual EI per y on ΔBMI z-score per y for the full study sample and in 4- < 6 y old children. Previous studies have assessed longitudinal changes in diet assuming a linear change between ages [10, 11, 30]. Anderson [34] for the first time modelled a piecewise linear relationship between age and energy intake to account for greater increases of EI in earlier childhood (from 3 - 7 y) compared to later childhood (7 -13 y). Findings from the present study confirmed an overall non-linear increase in daily EI during childhood depending on changes in height, sex and age. Changes in daily EI (from baseline to follow-up) decline non-linearly across age groups, especially for boys who at the same time show a decrease in change of height growth (Table 2). Using residual EI we accounted for non-linearity in changes in energy intake during childhood in this sample of pre-school children and school aged children.

The European Society for Paediatric Gastroenterology, Hepatology, and Nutrition (ESPGHAN) Committee on Nutrition supports the prevention of childhood obesity

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**Table 1 Characteristics of the study population (total group and stratified by age; total numbers and percentages)**

|                  | 2- < 4 years | 4- < 6 years | 6- < 8 years | 8- < 10 years | All |
|------------------|-------------|-------------|-------------|--------------|-----|
| N                | 481         | 729         | 1005        | 1048         | 2753|
| %                | 100         | 100         | 100         | 100          | 100 |

**Country**

|        | 2- < 4 years | 4- < 6 years | 6- < 8 years | 8- < 10 years | All |
|--------|-------------|-------------|-------------|--------------|-----|
| Italy  | 174 36.2    | 284 39      | 378 35.8    | 196 40.3     | 1032 37.5 |
| Estonia| 48 10       | 81 11.1     | 125 11.8    | 98 20.2      | 352 12.8  |
| Cyprus | 12 2.5      | 28 3.8      | 83 7.9      | 14 2.9       | 137 5     |
| Belgium| 26 5.4      | 33 4.5      | 30 2.8      | 2 0.4        | 91 3.3    |
| Sweden | 96 20       | 93 12.8     | 134 12.7    | 68 14        | 391 14.2  |
| Germany| 57 11.9     | 111 15.2    | 143 13.5    | 46 9.5       | 357 13    |
| Hungary| 28 5.8      | 41 5.6      | 89 8.4      | 51 10.5      | 209 7.6   |
| Spain  | 40 8.3      | 58 8        | 75 7.1      | 11 2.3       | 184 6.7   |

**ISCED-Level of parents**

|            | 2- < 4 years | 4- < 6 years | 6- < 8 years | 8- < 10 years | All |
|------------|-------------|-------------|-------------|--------------|-----|
| Low Education | 44 9.1     | 69 9.5      | 86 8.1      | 52 10.7      | 251 9.1 |
| Medium Education | 250 52     | 421 57.8    | 582 55.1    | 284 58.4     | 1537 55.8 |
| High Education  | 187 38.9   | 239 32.8    | 389 36.8    | 150 30.9     | 965 35.1 |

**Sex of the child**

|        | 2- < 4 years | 4- < 6 years | 6- < 8 years | 8- < 10 years | All |
|--------|-------------|-------------|-------------|--------------|-----|
| Boys   | 259 53.8    | 403 55.3    | 544 51.5    | 221 45.5     | 1427 51.8 |
| Girls  | 222 46.2    | 326 44.7    | 513 48.5    | 265 54.5     | 1326 48.2 |

**Weight group**

|          | 2- < 4 years | 4- < 6 years | 6- < 8 years | 8- < 10 years | All |
|----------|-------------|-------------|-------------|--------------|-----|
| Underweight | 55 11.4    | 85 11.7     | 92 8.7      | 40 8.2       | 272 9.9  |
| Normal weight | 361 75.1   | 491 67.4    | 699 66.1    | 300 61.7     | 1851 67.2 |
| Overweight | 52 10.8     | 98 13.4     | 171 16.2    | 92 18.9      | 413 15.0 |
| Obese    | 13 2.7      | 55 7.5      | 95 9.0      | 54 11.1      | 217 7   |

**Study region**

|              | 2- < 4 years | 4- < 6 years | 6- < 8 years | 8- < 10 years | All |
|--------------|-------------|-------------|-------------|--------------|-----|
| Intervention region | 273 56.8   | 416 57.1    | 548 51.8    | 254 52.3     | 1491 54.2 |
| Control region   | 208 43.2    | 313 42.9    | 509 48.2    | 232 47.7     | 1262 45.8 |
| All            | 481 100     | 729 100     | 1057 100    | 1048 100     | 2753 100 |

a International Standard Classification of Education Maximum (ISCED); maximum of both parents (0, 1, 2 = low education; 3, 4 = medium education; 5, 6 = high education)
and recommends the individual determination of energy intake, taking energy expenditure and growth into account [35]. In our study, results from the cross-sectional analyses revealed associations between residual EI at baseline and baseline BMI z-score for the whole study sample, boys and 6- < 8 y and 8- < 10 y olds. These findings support previous associations between BMI z-score and total EI in school children from other cross-sectional studies [5, 6, 36]. Cross-sectional observations provide no insight into temporality of the relationship between EI and BMI z-score, and reverse causation is considered a major concern. Our longitudinal findings provide better evidence to support causality between EI and subsequent changes in BMI z-score, even though size of the effect estimate was larger cross-sectionally. Results from the present longitudinal analyses showed that Δresidual EI was associated with subsequent increase in annual BMI z-score for the full study sample and for 4- < 6 y olds. These findings are both, in agreement [14, 30, 37] and disagreement [9, 11] with results from other longitudinal studies investigating the association between EI with subsequent change in BMI z-score. In our study, reported EI was slightly lower than recommended by the Scientific Committee on Food for boys and girls in the European Community [38]. We did not find associations between residual EI and BMI z-score for girls, neither cross-
sectionally nor longitudinally. For girls, changes in EI and relative ΔBMI z-score were generally lower than for boys, especially for the 4-<6 y olds. Higher EI for boys than for girls has been observed in previous longitudinal European and U.S. investigations [11, 30, 39]. The observed higher consumption of energy-dense foods among growing children may arise from the necessity to fulfil increased energy demands [40]. In contrast to earlier longitudinal studies [14, 39], changes in EI among 4-<6 y boys of our sample was higher while their change in height was lower (compared to girls of the same age with lower change in EI, Table 2) and may explain the subsequent increase of change in BMI z-scores. Even though cross-sectional data suggest that for boys, an EI above the average EI for the respective age and height may be an important determinant of increases in BMI z-score two years later this association was not observed in the longitudinal analysis.

Longitudinal findings from earlier research describe the protective role of physical activity on weight gain, implying an important role of physical activity in the development of obesity [41–43]. In U.S. children and adolescents, MVPA was inversely associated with fat mass and with BMI [8]. In our study, longitudinally Δresidual EI was no longer associated with ΔBMI z-score after adjustment for daily minutes of MVPA in a subsample of the children. However, the size of the effect estimate remained almost unchanged such that the insignificant association might be a result of the smaller power resulting from the smaller sample size. Our finding underlines the importance of studying sex- and age-related energy intakes, as they are influenced by growths and most likely by PA in seeking to understand the complex relationships between utilization (growth, expenditure or storage) of dietary energy and BMI z-score in childhood.

### Table 3 Cross-sectional effect of age, height and sex-adjusted residuals of energy intake at baseline (1 unit ~ 100 kcal) on BMI z-score at baseline stratified by sex and age group at baseline

|                  | Boys     | Girls    | 2-<4 years | 4-<6 years | 6-<8 years | 8-<10 years | All         |
|------------------|----------|----------|------------|------------|------------|-------------|-------------|
|                  | (N = 1427) | (N = 1326) | (N = 481)  | (N = 729)  | (N = 1057) | (N = 486)   | (N = 2753)  |
| βEI SE           | 0.11     | 0.03     | 0.06       | 0.03       | 0.02       | 0.05        | 0.11        |
| 99 % CI          | 0.02 - 0.18 | -0.02 - 0.13 | -0.14 - 0.10 | -0.04 - 0.18 | 0.03 - 0.19 | 0.04 - 0.25 | 0.04 - 0.14 |
|                  | 0.11     | 0.03     | 0.06       | 0.03       | 0.01       | 0.05        | 0.10        |
| 99 % CI          | 0.02 - 0.14 | -0.01 - 0.14 | -0.13 - 0.10 | -0.02 - 0.19 | 0.02 - 0.18 | 0.02 - 0.22 | 0.04 - 0.14 |
|                  | 0.13     | 0.03     | 0.04       | 0.03       | 0.02       | 0.06        | 0.08        |
| 99 % CI          | 0.04 - 0.21 | -0.05 - 0.12 | -0.18 - 0.13 | -0.04 - 0.21 | -0.02 - 0.17 | 0.02 - 0.26 | 0.03 - 0.15 |

Model 1) is the crude model adjusted for age, sex, country, if not stratified by age/sex, respectively
Model 2) moreover includes maximum parental ISCED level, parental BMI and screen time
Model 3) moreover includes MVPA (subgroup analysis; N = 1, 933)
(Models stratified by sex are not adjusted for sex)
βEI Estimate for EI, SE Standard Error, 99 % CI 99 % Confidence Interval

### Table 4 Longitudinal effect of change in residual energy intake per year (Δresiduals EI, 1 unit ~ 100 kcal) on ΔBMI z-score per year stratified by sex and baseline age group

|                  | Boys     | Girls    | 2-<4 years | 4-<6 years | 6-<8 years | 8-<10 years | All         |
|------------------|----------|----------|------------|------------|------------|-------------|-------------|
|                  | (N = 1427) | (N = 1326) | (N = 481)  | (N = 729)  | (N = 1057) | (N = 486)   | (N = 2753)  |
| βEI SE           | 0.017    | 0.007    | 0.011      | 0.007      | 0.013      | 0.017       | 0.029       |
| 99 % CI          | 0.006 - 0.029 | -0.031 - 0.057 | 0.003 - 0.057 | -0.013 - 0.025 | -0.020 - 0.023 | 0.001 - 0.008 | 0.014 - 0.005 |
|                  | 0.0178   | 0.007    | 0.011      | 0.007      | 0.021      | 0.017       | 0.032       |
| 99 % CI          | 0.005 - 0.025 | -0.023 - 0.065 | 0.005 - 0.058 | -0.014 - 0.024 | -0.021 - 0.021 | 0.000 - 0.008 | 0.015 - 0.005 |
|                  | 0.015    | 0.008    | 0.006      | 0.008      | 0.024      | 0.020       | 0.027       |
| 99 % CI          | 0.003 - 0.009 | -0.003 - 0.008 | 0.003 - 0.009 | -0.003 - 0.007 | -0.003 - 0.006 | 0.011 - 0.005 | 0.001 - 0.005 |
|                  | -0.006   | 0.037    | -0.014     | -0.025     | -0.029     | -0.078     | -0.003     |
| 99 % CI          | -0.002 - 0.006 | -0.002 - 0.006 | 0.000 - 0.006 | -0.003 - 0.006 | -0.002 - 0.006 | -0.003 - 0.006 | -0.003 - 0.006 |

Model 1) is the crude model adjusted for BMI z-score and residual EI at baseline, country, age, sex, if not stratified by age/sex, respectively
Model 2) moreover includes maximum parental ISCED level, parental BMI and screen time
Model 3) moreover includes MVPA (subgroup analysis; N = 1, 933)
(Models stratified by sex are not adjusted for sex)
βEI Estimate for EI, SE Standard Error, 99 % CI 99 % Confidence Interval
Strengths and limitations
In the IDEFICS study dietary information was given by proxy-respondents. Proxy-reporting relates to the number of meals under parental control; incomplete reporting of dietary intakes may contribute to reporting bias [44]. In order to reduce errors due to portion size estimation, erroneous food composition tables, incomplete recalls, misreporting or daily variations in intake we followed a rigorous approach. Firstly, the development of a computer-assisted assessment tool with standardized photographs, with multiple plausibility checks and reminding questions facilitated reporting of accurate portion sizes and complete recalls. School/kindergarten meal assessment through observers helped to complete all foods and beverages consumed in the setting. Secondly, the exclusion of incomplete recalls and recalls with implausible energy reporting corrected substantially for reporting bias. Plausible reporters (mean age 8.4 y) were found to be older than over-reporters (7.3 y) and younger than under-reporters (8.7 y); the mean BMI z-score (0.72 kg/m²) of plausible reporters was higher than that of over-reporters (+0.12 kg/m²) and lower than that of under-reporters (1.44 kg/m²) as observed in earlier studies [13]. Hence, a small selection bias that may have led to an attenuation of effect estimates cannot completely be precluded. However, inclusion of misreports may obscure or even inverse relationships with weight status as recently reported [45]. Finally, deriving the usual EI based on the NCI method [29] and accounting for day-to-day variation in EI is a clear strength of this study. Even though proxy-reported dietary data has limitations, we still believe our data is valuable for population research.

The IDEFICS study allows a deeper insight into the effect of deviations from usual age-, sex- and height-specific EI on BMI z-scores in 2 to 10 y old children across Europe. The calculation of residual EI as exposure variable instead of merely using EI as done in previous studies, helped to account for growth-related changes in usual EI when investigating the effect of EI on BMI z-scores in a population of children covering a wide age span. However, using residuals from a growth model as exposure induces uncertainty in the effect estimates and may hence increase the risk for type 2 error. In addition, no objective measurements of individual energy expenditure were available for the two examination time points (and PA only in a sub-group). Hence, residual EI estimates did account for differences in EI resulting not only from differences of body height, height growth but also from different PA levels between children. In a “perfect” data situation, the best option for the estimation of residual EI might be to regress EI on measured total energy expenditure, but unfortunately measuring total energy expenditure using doubly labeled water (DLW) was not feasible in this large-scale multi-centre children cohort for cost, logistic and ethical reasons.

In the present study, no substantial differences were observed when comparing main characteristics of the baseline sample and the sample presented here. As only 17% of the initial baseline cohort provided follow-up data and complete co-variate information, the IDEFICS Study - like many cohort studies - may therefore suffer from volunteer bias; we acknowledge this as a possible study limitation. However, the large sample size comprises data from eight European countries; the strictly standardized data assessment, documentation and data cleaning processing guarantee the highest possible data quality. Further, availability of relevant confounders, such as parental BMI and educational level, screen time and objectively measured MVPA is an additional strength of this study.

Conclusion
In conclusion, age-, sex-, and height-specific deviations from population average EI were weakly associated cross-sectional and longitudinal with changes in BMI z-scores in European children aged 2 to 10 y old. Longitudinal associations between changes in EI and BMI were particularly observed for the full study sample and for 4-<6 y old children. Our results suggest that the promotion of appropriate energy intakes may be most effective when considering sex- and age-related energy demands due to different growth and PA levels according to the recommendation of the ESPGHAN Committee on Nutrition. Residual EI seems to be a useful exposure measures when studying effects of dietary exposures in children with a wide age span as they accommodate the growth-related changes in dietary intake during childhood.

Additional file

Additional file 1: Baseline and follow-up characteristics (mean and SD) of exposure and outcome variates given by sex and age group at baseline for MVPA-subgroup. (DOCX 21 kb)

Abbreviations
BMI: body mass index; DLW: doubly labeled water; EI: energy intake; ESPGHAN: European Society for Paediatric Gastroenterology, Hepatology, and Nutrition; FCT: food composition tables; IDEFICS Study: Identification and prevention of Dietary- and lifestyle-induced health Effects In Children and infants Study; ISCED: International Standard Classification of Education; MVPA: moderate to vigorous physical activity; NCI: National Cancer Institute (USA); PA: physical activity; SACINA: Self-Administered Children and Infant Nutrition Assessment; SES: Socio economic status; T0/T1: Baseline examinations/follow-up; TV: television; 24HDR: 24-hr dietary recalls.

Competing interests
The authors declare that they have no competing interests.

Authors’ contributions
This manuscript represents original work that has not been published previously and is currently not considered by another journal. The authors’ responsibilities were as follows – AH and CB had the idea of the analysis; AH wrote the paper and had primary responsibility for final content; CB...
conducted data analysis; MW cleaned dietary data; AH, SDH, GE, CH, EK, VK, LAM, VP, TV and GB conducted research. All authors were responsible for critical revisions and final approval of the manuscript.

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Study subjects and their parents could consent to single components of the study while abstaining from others. Parents had given consent for examinations, collection of samples, subsequent analysis and storage of personal data and collected samples. Study subjects and their parents could consent to single components of the study while abstaining from others.

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