High beverage sugar as well as high animal protein intake at infancy may increase overweight risk at 8 years: a prospective longitudinal pilot study

Peter JM Weijs, Laura M Kool, Nicolen M van Baar and Saskia C van der Zee

Abstract

Background: Combined effects of early exposure to beverage sugar and animal protein and later life overweight risk have not been studied.

Methods: A prospective longitudinal study was initiated in 2001 with 226 infants between 4 and 13 months of age. Dietary intake was assessed with a 2 day food record. Also information on infant body weight and socio-economic status was obtained at baseline. At 8 year follow-up in 2009, children were surveyed again. Main outcome measure was overweight at 8 years as defined by BMIsds ≥ +1.0. Also maternal BMI, present dietary intake and physical activity, were obtained by questionnaire and 2-day food record.

Results: At the 8 year follow up, 120 children (53%) were surveyed again. Of those, questionnaires and food records were completed for 63 children, for the other 57 children only weight and height at 8 years was available; 20 out of 120 children (17%) were self-reported overweight at 8 years of age. Unadjusted odds ratios (ORs; 95% CI) for overweight at 8 years were 1.10 (1.02, 1.18) for beverage sugar intake per one percent of energy intake and 4.06 (1.50, 11.00) for the highest tertile of animal protein intake at infancy compared to the lowest two tertiles. After adjustment for sex, age, infant weight, breastfed at intake assessment, and socio-economic status, odds ratios were 1.13 (1.03, 1.24) for beverage sugar, and 9.67 (2.56, 36.53) for highest tertile of animal protein intake. In the subgroup with completed questionnaire (n = 63) ORs were also adjusted for current maternal overweight, more than 2 months full breastfeeding, physical activity, and energy intake, but ORs remained significantly associated with overweight at 8 years.

Conclusions: A high intake of sugar containing beverages as well as animal protein in the first year of life may increase the risk of overweight at 8 years. The results of this pilot investigation should be confirmed in a larger cohort.

Introduction

Obesity and childhood obesity are increasing and etiology, epidemiology and treatment should be addressed [1]. After fetal nutritional experiences and early breastfeeding in the first months of life, the introduction of complementary feeding along with either breastfeeding or formula feeding is important for long-term infant development [2-4]. A high infant protein intake has been suggested to be related to childhood overweight [5-8], although not confirmed in hispanic children [9]. The suggested mechanism is an accelerated growth during infancy via stimulation of insulin and IGF-1 [2]. Sugar containing beverage intake in childhood has been shown to be related to overweight in childhood [10], and this effect seems to extend into adulthood [11]. The suggested mechanism is that higher sugar containing beverage intake is not compensated for by lower energy intake from other sources [12]. More specifically it is suggested that beverages from fruit sources or corn syrup contain a high level of fructose [12,13]. Fructose does not increase insulin levels after intake as glucose
does, which results in relatively high levels of plasma tri-
glycerides as a result of unsuppressed lipolysis [13]. How-
ever, there is no conclusive evidence for fructose as the
central cause of obesity [14,15]. Also very early postnatal
high carbohydrate complementary feeding is suggested to
be related to later overweight [3]. These effects might
imprint metabolic changes that last into later years of life
and predispose for obesity and cardiovascular illness [2].

That other factors play a role in the etiology of over-
weight is undisputed, including overweight parents and
inactive lifestyle of child [16-18]. However, we were able
to retrieve only one study on beverage sugar intake dur-
ing infancy and risk of overweight development [19]. The
European Society of Pediatric Gastroenterology, Hepatol-
ogy, and Nutrition (ESPGHAN) states that there are few
data on the effects of specific complementary foods on
growth [20].

The aim of the present pilot study was to determine
whether high beverage sugar intake or high animal protein
intake during infancy is associated with an increased risk
of overweight at the age of 8 years.

Methods
Subjects
In April 2001 226 parents of infants 4-13 months of age
responded to a notice distributed through a publisher of
a magazine aimed at parents of very young children.
Almost equal samples were obtained at 5 different age
groups (4, 6, 8, 10, and 12 months of age) to cover the
larger part of infancy that is not exclusively based on
breastfeeding and/or formula feeding. Parents/caretakers
of infants were invited to complete a questionnaire on
weight, height, and parental education. They also
received a 2 day food record with instructions. Descrip-
tion of calculation of nutrient intake have been published
before [21,22]. Parents/caretakers signed an informed
consent form. All procedures were in accordance with
the ethical standards of the Hogeschool van Amsterdam.

Five of the 226 parents were excluded because they had
not explicitly provided permission to be approached
again after the initial assessment. Therefore, in april
2009, 221 of the 226 parents/caretakers were mailed
again with a letter of invitation, a short questionnaire and
a 2 day food record. To the parents no direct reference
was made to the problem of overweight or obesity, to
avoid bias in the provided information. The response was
63 out of 221 complete questionnaires including the
2 day food record. The non-responders were approached
again by telephone with only the question for current
weight and height of their child. In this way another 57
children were included into this study. Total loss of fol-
low-up (221-120 = 101) was due to migration (49 letters
returned to sender) or failure to trace (52 no contact by
telephone).

Socio-economic status was judged by parental educa-
tion. Mothers education was divided into high and low
level of education, with high level of education defined as
university, higher professional education, or preparatory
education (college) for scientific education. All other edu-
cation was defined as low level of education.

Dietary intake at infancy
By means of a 2 day food record (1 weekday and 1 week-
end day) the parents provided information on actual con-
sumption of food in portions, or weighed if necessary.
Parents were specifically asked to subtract spilled or not
consumed amounts. Portion sizes were translated into
weight by standard portion sizes used in The Netherlands
[23]. Food weights were then multiplied by actual con-
tents of nutrients per 100 g food by using the Dutch
Food Composition Table [24,25]. For foods that could
not be translated to existing Dutch Food Composition
Table codes, mostly meals in a jar, the nutrients were cal-
culated using data on the nutrient levels of ingredients.

Sugar containing beverages were any drinks provided to
the infant that contained mono- and/or disaccharides,
thus both naturally sweetened, sweetened by industry or
sweetened at home; however milk, milk products, mothers
milk, and infant formula were excluded. Animal protein
was defined as protein minus vegetable protein, which are
both available from the Dutch Food Composition Table.

Questionnaire at 8 years
In 2009 parents were asked for weight and height of
father, mother and the 8 year old child. Parents were ret-
rospectively asked for breastfeeding and formula feeding
practices apart from the at baseline assessment of dietary
intake. This retrospective information was complemen-
tary to the prospective information from dietary assess-
ment at infancy. Parents were asked whether their child
was ever breastfed or not, formula fed or not, a combina-
tion of breastfeeding and formula feeding, and for how
long. The 2 day food record was calculated for current
nutritional intake, in order to adjust for current energy
intake. Parents were asked for total hours per week that
the child currently spends on physical activity (4 items:
sports at school, sports at club, free outside play, and
active travel to and from school) and inactivity (3 items:
television watching, computer use, and gaming).

Definition of overweight
Current weight and height were self-reported. Weight and
height were used to calculate body mass index (kg/m²).
Since BMI is sex and age specific for children, the BMI
standard deviation score (BMIsds) was used, as calculated
with the Growth Analyzer http://www.growthanalyser.org.
WHO BMIsds cut-off point of +1 and +2 was used to
define overweight and obesity, equivalent to BMI 25 and
30 kg/m² at 19 years (http://www.who.int/growthref/who2007_bmi_for_age/en/index.html at 19-06-2009). Overweight at 8 years includes children with obesity in the present analysis. Current maternal overweight was defined as BMI > 25.

Statistical analyses

Differences in continuous variables between infant age groups were assessed with an independent samples t-test or analysis of variance (ANOVA) and dichotomous variables with the Chi square test.

The relation between animal protein and outcome was nonlinear and therefore included as tertile in the model (highest tertile = 1, lower two tertiles = 0). Tertiles for animal protein intake were generated per age group (4, 6, 8, 10, 12 months), since protein intake at 4 months is very different from protein intake at 12 months. Beverage sugar intake was used as continuous variable. Analyses were performed on dichotomous outcome overweight at 8 years with logistic regression analysis. We performed separate analyses for the group with information about weight and height at 8 years (n = 120) and the subgroup with complete questionnaires at 8 years (n = 63). Model 1 is the unadjusted effect of beverage sugar or animal protein intake, both as percentage of energy intake. Model 2a and 2b are adjusted for sex, infant age, body weight, and socio-economic status based on highest level of maternal education [2]. Since animal protein intake for breastfed children is different from non-breastfed children, we also adjusted for breastfeeding at the time of intake assessment. Model 2 is adjusted for each other, thus both beverage sugar and animal protein. Model 3 is adjusted for both beverage sugar and animal protein, and for adjustments from model 2a/2b. Model 4 is also adjusted for confounders at age 8 years, namely current maternal overweight, current physical activity in hours per week, and current energy intake in kcal per day, as well as duration of full breastfeeding of more than two months [16].

In addition, linear regression analysis was performed with BMIsds as dependent variable and beverage sugar and animal protein intake as independent variable to quantify the possible effect on BMIsds.

Statistical analysis was performed by SPSS 17 (SPSS Inc. Chicago, USA), and a P-value less than 0.05 was considered statistically significant. Means in the text are presented with SD.

Results

A total of 120 (64 boys, 56 girls) infant BMI were used for analysis. Mean (sd) ages of responders at the time of completing the questionnaire were 4.8 (0.3), 6.9 (0.3), 8.8 (0.3), 10.9 (0.3), and 12.4 (0.2) months (see also Additional file 1, Table S1). Mean age at infancy was 8.7 (2.8; range 4.4-13.0) months and at follow-up 8.7 (0.3; range 8.3-9.1) years. Mean BMI and BMIsds at 8 years was 16.3 (2.1) and -0.12 (1.17). For the whole group with complete data at 8 years (n = 63) 48 (80%) of the children had ever been breastfed, and 22 (37%) still received breastfeeding at the age of 6 months. Further infant characteristics are presented in Table 1. Table 2 shows food, energy, protein, and sugar intakes in infancy for children being overweight or not at age 8 years [26]. Both beverage sugar and animal protein intake from complementary foods like meat, milk and meals from a jar were higher in overweight children when adjusted for total energy intake. Overweight children consumed more drinks and less infant formula, and therefore also less protein from formula. The intake of beverage sugar and animal protein adjusted for total energy intake was fairly constant across age groups (see supplementay data file 1). Mean beverage sugar intake of users was not statistically different between age groups (one-way anova, with post-hoc Bonferroni). Protein from mothers milk is included in animal protein, but contributes a mean of about 10% to total animal protein in the three younger groups and only about 2.5% in the two older age groups. Formula milk contributes a mean of 47% (sd 37) to animal protein intake.

Table 1 Subject characteristics (in percentage or as mean ± SD)

| Total group (n = 120) | Subgroup (n = 63) |
|----------------------|------------------|
| Not overweight N = 100 | Overweight N = 20 | P | Not overweight N = 52 | Overweight N = 11 | P |
| Infant age, months | 85 ± 2.9 | 94 ± 2.6 | 0.18 | 83.2 ± 2.9 | 96.2 ± 2.3 | 0.21 |
| Infant weight, kg | 84 ± 1.5 | 90 ± 1.3 | 0.10 | 82.2 ± 1.4 | 92.2 ± 0.9 | 0.03 |
| Gender, % boys | 50 | 70 | 0.10 | 50 | 73 | 0.17 |
| High socio-economic status, % | 31 | 35 | 0.73 | 33 | 36 | 0.81 |
| Breastfed at time of intake assessment, % | 21 | 25 | 0.69 | 21 | 27 | 0.66 |
| Maternal overweight, % | - | - | - | 39 | 46 | 0.67 |
| BMI mother, kg/m² | - | - | - | 25.4 ± 5.1 | 25.6 ± 2.6 | 0.94 |
| Physical activity, h/week | - | - | - | 14.4 ± 5.3 | 12.1 ± 5.8 | 0.20 |
| Duration of breastfeeding, weeks | - | - | - | 35 ± 36 | 25 ± 24 | 0.32 |
Interactions between beverage sugar and animal protein intake and age, gender, socio-economic status, and maternal overweight were tested, and all were found to be not significant.

A total of 20 out of 120 children (17%) were considered overweight (BMI<sup>sd</sup> > +1.0) and in the subsample 9 out of 63 children were overweight (14%), which is not significantly different (p = 0.675). Table 3 shows that the unadjusted odds ratio (95% CI) was 1.10 (1.02, 1.18) for beverage sugar intake, and 4.06 (1.50, 11.00) for the highest tertile of animal protein intake compared to the lowest two tertiles. Adjusted odds ratios (model 3) were 1.13 (1.03, 1.24) for beverage sugar and 9.67 (2.56, 36.53) for animal protein, thus both were still independently associated with overweight at 8 years. In the subgroup analysis (n = 63) model 4 showed that after addition of current maternal overweight, full breastfeeding for more than two months, current physical activity in hours per week, and current energy intake in kcal per day, the ORs for beverage sugar and animal protein were still significantly associated with overweight at 8 years.

When protein from mothers milk was excluded from animal protein, this did not change the OR’s. The OR’s for becoming overweight for animal protein become slightly lower but demonstrate very significant independent effects (beverage sugar: model 2, OR 1.11 (1.03, 1.19) p = 0.006; model 3, 1.12 (1.02, 1.22) p = 0.014; animal protein: model 2, 3.27 (1.16, 9.19) p = 0.025; model 3, 4.66 (1.47, 14.8) p = 0.009).

Figure 1 shows that for infants in the lower two tertiles for both beverage sugar and animal protein, only 3

Table 2 Food, energy and nutrient intake of infants who became overweight or not at 8 years

| Food, energy and nutrient intake of infants who became overweight or not at 8 years | Not overweight | Overweight | Difference: overweight vs not overweight |
|--------------------------------|----------------|------------|------------------------------------------|
|                               | n = 100        | n = 20     | Mean | sd  | Mean | sd  | Mean | 95% CI | P         |
| Formula, g                    | 487            | 318        | -169 | -327 | -12  | 0.04 |
| Mothers milk<sup>1</sup>, g    | 85             | 122        | 37   | -62  | 135  | 0.46 |
| Drinks<sup>2</sup>, g         | 73             | 162        | 89   | 11   | 169  | 0.03 |
| Milk and milk products<sup>2</sup>, g | 117          | 181        | 64   | -42  | 170  | 0.23 |
| Meat, meat products and fowl<sup>2</sup>, g | 10.9       | 14.5       | 3.6  | -4.6 | 11.9  | 0.39 |
| Energy, kcal                  | 836            | 859        | 23   | -73  | 119  | 0.64 |
| Energy, kcal/kg               | 99             | 96         | -3   | -12  | 5    | 0.38 |
| Protein, g                    | 264            | 29.9       | 3.5  | -2.5 | 9.5   | 0.25 |
| Protein, g/kg                 | 3.0            | 3.2        | 0.2  | -0.3 | 0.7   | 0.46 |
| Protein, animal<sup>3</sup>, g | 19.3           | 21.7       | 2.4  | -1.8 | 6.6   | 0.26 |
| Protein, animal, en%          | 9.0            | 9.9        | 0.9  | -0.4 | 2.3   | 0.19 |
| Protein, from formula only, g | 8.4            | 5.6        | -2.8 | -5.5 | -0.9  | 0.04 |
| Protein, from formula only, en% | 4.2           | 3.0        | -1.4 | -2.9 | 0.1   | 0.06 |
| Protein, animal, complementary foods, g | 11.0       | 15.8       | 4.8  | -0.9 | 10.6  | 0.10 |
| Protein, animal, complementary foods<sup>4</sup>, en% | 4.6         | 6.9        | 2.3  | 0.0  | 46    | 0.05 |
| Fat, g                        | 30.4           | 30.6       | 0.2  | -3.2 | 3.5   | 0.93 |
| Fat, en%                      | 33.8           | 32.6       | -1.2 | -4.9 | 2.4   | 0.50 |
| Carbohydrate, g               | 114            | 116        | 2    | -14  | 17    | 0.82 |
| Carbohydrate, en%             | 54.1           | 53.8       | -0.3 | -2.9 | 2.4   | 0.86 |
| Sugar, g                      | 74.0           | 77.4       | 3.4  | 5.4  | 12.2  | 0.45 |
| Sugar, en%                    | 36.1           | 36.3       | 0.2  | -3.5 | 3.9   | 0.91 |
| Beverage sugar<sup>5</sup>, g | 10.6           | 19.7       | 9.1  | -0.7 | 18.9  | 0.07 |
| Beverage sugar, en%           | 4.5            | 8.8        | 4.3  | 0.2  | 8.4   | 0.04 |

1 Mothers milk in gram is assessed by frequency and duration of breastfeeding, not weighing.
2 These food groups are predefined by the Dutch Food Composition Table.
3 Protein not from animal sources is predefined by the Dutch Food Composition Table, protein from animal sources is derived by difference between protein and protein not from animal sources.
4 Complementary foods with protein from animal sources are a sum of 6 predefined food groups from the Dutch Food Composition Table: milk and milk products, cheese, eggs, meat/meat products and fowl, fish, and precomposed foods (like meals in a jar; this last group may also contain some protein not from animal sources).
5 For Drinks, Beverage sugar in gram and Beverage sugar in en% were equal variances not assumed, resulting in a non significant differene for Beverage sugar in gram.
out of 58 infants (5%) were categorised as overweight. Infants in the highest tertile for either beverage sugar or animal protein had 5/23 (22%) or 7/27 (26%) infants with overweight at 8 years, respectively. Infants that were in the highest tertile for both beverage sugar and animal protein had 5/12 (42%) overweight at 8 years. The percentage overweight between the four groups of Figure 1 was significantly different (Chi square test, \( p = 0.005 \)). Table 4 shows that linear regression analysis provided an effect size of beverage sugar on BMIsds of 0.05 per energy percent of increase in beverage sugar intake, which was not altered by adjustment for confounders. A change in beverage sugar intake of 10 percent of energy intake, might therefore be responsible for an increase in BMIsds of 0.5. For the highest tertile of animal protein intake the increase was 0.5-1.0 BMIsds compared to the lower two tertiles.

**Discussion**

To our knowledge this is the first study to show two independent effects of both beverage sugar and animal protein intake during infancy and overweight at age 8 years. Beverage sugar intake during infancy increases the risk of overweight at 8 years by more than 10% extra risk per energy percent beverage sugar increase. For animal protein infants with the highest tertile of intake had a more than 9 times higher risk of becoming overweight at 8 years.

In 1995 Rolland-Cachera suggested the early protein hypothesis; a high protein intake in excess of metabolic requirements enhances weight gain in infancy and increases the risk of obesity later in life [5]. This hypothesis has been confirmed in similar small studies [6-8], although not in hispanic youth [9]. The period of dietary exposure, the timing of overweight assessment and the definition of overweight might in part explain different outcomes. Gunther et al. suggested that particularly

| Model  | Beverage Sugar | Animal Protein |
|--------|----------------|---------------|
| Model 1a | 1.10 (1.02, 1.18) | 4.06 (1.50, 11.00) |
| Model 1b | 1.10 (1.02, 1.20) | 7.38 (2.17, 25.10) |
| Model 2a | 1.11 (1.03, 1.20) | 4.78 (1.65, 13.87) |
| Model 2b | 1.13 (1.03, 1.24) | 4.67 (2.56, 36.53) |
| Model 3a | 1.13 (1.03, 1.24) | 9.67 (2.56, 36.53) |
| Model 3b | - | - |
| Model 4a | - | - |
| Model 4b | - | - |

Model 1a, 1b: unadjusted effects (a and b indicate two separate analyses).
Model 2a, 2b: adjusted for sex, infant age, infant body weight, breastfed at time of assessment, SES (a and b indicate two separate analyses).
Model 2: adjusted for each other (beverage sugar and animal protein).
Model 3: adjusted for each other, plus sex, infant age, infant body weight, breastfed at time of assessment, SES.
Model 4: adjusted as model 3 plus maternal overweight, physical activity in hours per week, and energy intake at 8 years in kcal per day.

Figure 1 The prevalence of overweight (%) according to the highest and combined lower tertiles of beverage sugar (BS) intake as well as for animal protein (AP) intake.
animal protein might be responsible for the association [27]. When the present analysis was performed with two dummies for second and third tertile of animal protein intake (data not shown), the second tertile did not show a significant relationship. In line with this observation is the hypothesis that animal protein above a certain level of intake might deregulate insulin and IGF-1 with an impact on preadipocyte differentiation and multiplication [2,27]. High protein intakes may also decrease human growth hormone secretion and reduce lipolysis, which might aid fat accumulation [2]. Lower protein levels in infant formula seems to reduce weight up to age 2 years [28], and might therefore contribute to reduction of later overweight.

A recent review by Gibson [14] showed an overwhelming amount of studies on sugar sweetened soft drinks and obesity. Gibson included all cold beverages containing added sugars, carbonated or not. The evidence seems to be inconclusive, although a small effect of sugar sweetened soft drinks on BMI is suggested. A recent study by Herbst et al. [19] is the single study that prospectively investigated infant beverage sugar intake during the first year of life. Herbst et al. [19] show no or possibly a protective effect of sugar from beverages and sweets, however the level of beverage sugar intake is very low. At 1 year of age the beverage sugar intake is 0 percent of total energy intake in the Donald study [19] compared to more than 5 percent in the present study. A reasonable explanation for this difference is that the Donald study has selected infants from a higher socio-economic level: > 80% of infants was breastfed for more than 4 months versus 41% in our study, > 60% of parents had a high socio-economic status versus one third in our study, and maternal overweight was only 25% versus > 40% in our study). In line with our study, the authors conclude that when added sugar intake increases to higher levels this might be detrimental to BMI development.

Beverage sugar was consumed through apple and orange juice, most often from apple concentrate, and lemonade also from sugar concentrate. Apple juice has a high fructose content. Fructose consumption results in decreased circulating levels of insulin and leptin when compared with glucose [13]. Because insulin and leptin function as key signals to the central nervous system in the regulation of energy balance, it has been assumed that over a longer period of time high fructose intake could lead to increased caloric intake or decreased caloric expenditure and weight gain [15]. However, at this moment and from this study it is not possible to draw any firm conclusions about the mechanism involved.

Limitations of the study are the total number of infants, although similar studies were of similar size [6-8]. The low response rate to the initial study is also a limitation, but each child serves as its own control. Confounders were not all available for the whole sample of 120 children, but for the subsample of 63. However, the primary effect of beverage sugar and animal protein did not seem to change when tested in the subsample (from model 3 to model 4). Therefore we assume that the effects we found are very robust and would not be different when

| Model 1a beverage sugar | 0.049 (0.017, 0.081) | 0.003 | 0.050 (0.001, 0.099) | 0.047 |
| Model 1b animal protein | 0.47 (0.03, 0.92) | 0.037 | 0.69 (-0.01, 1.37) | 0.051 |
| Model 2a beverage sugar | 0.043 (0.006, 0.079) | 0.023 | 0.044 (-0.017, 0.104) | 0.152 |
| Model 2b animal protein | 0.57 (0.11, 1.02) | 0.015 | 0.82 (0.12, 1.52) | 0.022 |
| Model 3 beverage sugar | 0.049 (0.017, 0.081) | 0.003 | 0.049 (0.001, 0.097) | 0.045 |
| Model 3 animal protein | 0.48 (0.05, 0.91) | 0.029 | 0.68 (0.01, 1.35) | 0.049 |
| Model 4 beverage sugar | 0.044 (0.008, 0.080) | 0.016 | 0.049 (-0.009, 0.106) | 0.097 |
| Model 4 animal protein | 0.59 (0.14, 1.03) | 0.011 | 0.86 (0.17, 1.55) | 0.015 |
| Model 4a beverage sugar | - | - | 0.055 (-0.005, 0.115) | 0.070 |
| Model 4b animal protein | - | - | 0.97 (0.23, 1.71) | 0.011 |

Model 1a, 1b: unadjusted effects (a and b indicate two separate analyses).
Model 2a, 2b: adjusted for sex, infant age, infant body weight, breastfed at time of assessment, SES (a and b indicate two separate analyses).
Model 2: adjusted for other (beverage sugar and animal protein).
Model 3: adjusted for each other, plus sex, infant age, infant body weight, breastfed at time of assessment, SES.
Model 4: adjusted as model 3 plus maternal overweight, physical activity in hours per week, and energy intake at 8 years in kcal per day.
the whole sample would have provided all information on confounders. Although a large random sample was the basis of the study, socio-economic status (maternal education) was slightly higher compared to Dutch average http://statline.cbs.nl/statweb/. The educational level of parents was very similar to the other Dutch investigation into dietary intake of infants, the Nutrient Intake Research (VIO) study [29]. This study used a representative sample of 300 infants for both the 9 and the 12 month old infants. We have evaluated the quality of the self reported dietary intake data in an earlier study as good [21], also based on the similarity with the VIO study. Two day food records at one time point is a limited assessment of longer-term exposure to food constituents. However, providing high levels of beverage sugar and/or food products high in animal protein are most likely related to long established food habits of (primarily) the mother. It should also be stressed that both beverage sugar and animal protein were not related to poor food habits. Beverage sugar is largely from fruit juices and from apple concentrate, which the parents judge as ‘fruit’ and therefore as ‘good’. Animal protein is derived from normal healthy food groups and infant formula. The level of intake of percentage energy beverage sugar or animal protein is not significantly different between high and low socio-economic status groups. The relationship between high animal protein intake (highest tertile) with overweight at 8 years is even stronger in high compared to low socio-economic status group (p = 0.006 vs p = 0.102 respectively). This may have resulted in low underreporting as well as high consistency in food ‘habits’. Comparing responders with non-responders, no differences appeared in socio-economic status, but parents with breastfed infants at the time of investigation in 2001 had responded to the follow-up call in 2009 significantly more often (26/38 vs 93/188, p = 0.009). Information on maternal smoking is missing, however it is not likely that smoking is associated with beverage sugar or animal protein intake at infancy.

Also, the intake of mother’s milk may be more inaccurately estimated than the intake of formula milk. However, considering the low contribution of mother’s milk to the total consumption, it is unlikely to play a key role in effect size and/or significance.

The use of BMIsds as definition of overweight is not ideal but acceptable as an outcome measure; no definition of obesity is ideal at present [16]. Both weight and height are self reported which might introduce some bias. However, recently it has been shown that BMI misclassification was almost absent in an adult population when self reported weight and height were evaluated [30]. Children are more often subject to measurement of weight and height, therefore BMI misclassification is assumed to be limited. Although some BMI misclassification may have occurred it is not likely that the amount of BMI misclassification is related to protein and beverage sugar intake in infancy. It is unlikely this would have caused the significant effect found in this study since the effect of non-differential misclassification on estimates of relative risk will be to always reduce the true relative risk [31]. Physical activity is also self reported, however self reporting is probably less important than selection of a method with a high internal and external validity [32]. Another limitation is the observational nature of the design, which makes it difficult to conclude a causal nature [2].

Provision of sugar containing beverages and a high animal protein diet in the first year of life appears to increase the risk of overweight at 8 years. Although childhood overweight is clearly multi-factorial, each significant and clinically meaningful contributing factor that can be influenced by the parents should be addressed accordingly. Sugar containing beverages provided during infancy are much less difficult to address than lifestyle changes during later stages of life [32,33].

Future studies with a larger sample size should confirm these observations.

Acknowledgements
Tanja Braak and Brenda de Groot are acknowledged for their contribution in the original study. Special thanks to Rob Korver for assistance in the first part of the study and development of our own nutrient intake calculating program. Financial support was obtained from the Ministry of Health, Welfare and Sports (grantnumber 3989 83-01). The sponsor only received a report of the study.

Author details
1Department of Nutrition and Dietetics, School of Sports and Nutrition, Hogeschool van Amsterdam, University of Applied Sciences, Amsterdam, The Netherlands. 2Department of Nutrition and Dietetics, Internal Medicine, VU University Medical Center, Amsterdam, The Netherlands. 3EMGO Institute for Health and Care Research, VU University Medical Center, Amsterdam, The Netherlands. 4Department of Environmental Health, Public Health Service Amsterdam, Amsterdam, The Netherlands (address: Nieuwe Achtergracht 100, 1018 WT Amsterdam.

Authors’ contributions
PW designed the study, LK, NvB, PW conducted literature research, PW, LK, NvB, SvdZ took part in food calculations and statistical analysis, all authors contributed significantly to writing of the manuscript and all authors read and approved the final manuscript.

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

Received: 20 April 2011 Accepted: 23 September 2011 Published: 23 September 2011
References

1. Ogden CL, Carroll MD, Curtin LR, McDowell MA, Tabak CJ, Flegal KM: Prevalence of overweight and obesity in the United States, 1999-2004. JAMA 2006, 295:1549-55.

2. Koletzko B, von Kries R, Closa Monasterolo R, et al: Can infant feeding choices modulate later obesity risk? Am J Clin Nutr 2009, 89:1502S-8.

3. Patel MS, Srinivasan M, Laychock SG: Metabolic programming: Role of nutrition in the immediate postnatal life. J Inherit Metab Dis 2009, 32:218-28.

4. Brown CM, Dulloo AG, Montani JP: Sugary drinks in the pathogenesis of obesity and cardiovascular diseases. Int J Obes 2008, 32:529-34.

5. Rolland-Cachera MF, Deheeger M, Akrout M, Bellisle F: Influence of macronutrients on adiposity development: a follow up study of nutrition and growth from 10 months to 8 years of age. Int J Obes Relat Metab Disord 1995, 19:573-8.

6. Scaglioni S, Agostoni C, Notaris RD, et al: Early macronutrient intake and overweight at five years of age. Int J Obes Relat Metab Disord 2000, 24:770-71.

7. Hoppe C, Mølgaard C, Thomsen BL, Juul A, Michaelsen KF: Influence of added sugar intake in early childhood and body mass index at age 7 y of age. Am J Clin Nutr 2007, 85:1626-33.

8. Butte NF: Impact of Infant Feeding Practices on Childhood Obesity. J Nutr 2009, 139:412S-416S.

9. Ludwig DS, Peterson KE, Gottsmaker SL: Relation between consumption of sugar-sweetened drinks and childhood obesity: a prospective, observational analysis. Lancet 2001, 357:505-08.

10. Nissen E, Mikkila V, Mannisto S, et al: Sweets and sugar-sweetened soft drink intake in children in relation to adult BMI and overweight. The Cardiovascular Risk in Young Finns Study. Public Health Nutr 2009, 28:1-9.

11. Malik VS, Schulze MB, Hu FB: Intake of sugar-sweetened beverages and weight gain: a systematic review. Am J Clin Nutr 2006, 84:274-88.

12. Teff KL, Grudzka J, Townsend RR, et al: Endocrine and metabolic effects of consuming fructose- and glucose-sweetened beverages with meals in obese men and women: influence of insulin resistance on plasma triglyceride responses. J Clin Endocrinol Metab 2009, 94:1562-9.

13. Gibson S: Sugar-sweetened soft drinks and obesity: a systematic review of the evidence from observational studies and interventions. Nutr Res Rev 2008, 21:134-47.

14. Stanhope KL, Havel PJ: Fructose consumption: considerations for future research on its effects on adipose lipid metabolism, and insulin sensitivity in humans. J Nutr 2009, 139:1236S-1241S.

15. Reilly JJ, Armstrong J, Dorosti AR, et al: Early life risk factors for obesity in childhood: cohort study. BMJ 2005, 330:1357.

16. Miesman J, Rosboom TJ, Bonsel GJ, et al: Maternal pre-pregnancy body mass index explains infant’s weight and BMI at 14 months: results from a multi-ethnic birth cohort study. Arch Dis Child 2009.

17. Baird J, Fisher D, Lucas P, Kleijnjen J, Roberts H, Law C: Being big or growing fast: systematic review of size and growth in infancy and later obesity. BMJ 2005, 331:329.

18. Herbst A, Diethelm K, Cheng G, Alexy U, Icks A, Buyken AE: Direction of Associations between added sugar intake in early childhood and body mass index at age 7 years may depend on intake levels. J Nutr 2011, 141(7):1348-54.

19. ESPGHAN Committee on Nutrition. Complementary Feeding: A Commentary by the ESPGHAN Committee on Nutrition. J Pediatr Gastroenterol Nutr 2008, 46(1):99-110.

20. Weijs PJ, Bakker MI, Korver KR, van Goor Ghanavi-Ztch K, van Wijnen JH: Dioxin and dioxin-like PCB exposure of non-breastfed Dutch infants. Chemosphere 2006, 64:1521-5.

21. Weijs PJ, de Groot B, Braak T, Korver R: Protein intake is very high in non-breastfed Dutch infants 4-12 months of age. Clin Nutr 2005, 24:377.

22. van Donders-Eijgelen MR, van der Heijden L, Huikko KFAM, Maten, gewichten en codenummers. Landbouwuniversiteit Wageningen, Wageningen 1997.

23. NEVO Foundation: Dutch Food Composition Table. Netherlands Nutrition Centre, The Hague 2001.

24. NEVO Foundation: Dutch Food Composition Table. Netherlands Nutrition Centre, The Hague 2006.

25. Health Council of The Netherlands: Dietary Reference Intakes: energy, proteins, fats and digestible carbohydrates. Publication number 2001/19. Health Council of The Netherlands, The Hague 2001.

26. Gunther ALB, Buyken AE, Kroke A: Early protein intake and later obesity risk: which protein sources at which time points throughout infancy and childhood are important for body mass index and body fat percentage at 7 y of age? Am J Clin Nutr 2007, 86:1765-72.

27. Koletzko B, von Kries R, Closa R, et al: Lower protein in infant formula is associated with lower weight up to age 2 y: a randomized clinical trial. Am J Clin Nutr 2009, 89:1836-46.