

Prospective association between consumption frequency of organic food and body 2 weight change, risk of overweight or obesity: Results from the NutriNet-Santé Study

Emmanuelle Kesse-Guyot, Julia Baudry, Karen Assmann, Pilar Galan, Denis

Lairon

▶ To cite this version:

Emmanuelle Kesse-Guyot, Julia Baudry, Karen Assmann, Pilar Galan, Denis Lairon. Prospective association between consumption frequency of organic food and body 2 weight change, risk of overweight or obesity: Results from the NutriNet-Santé Study. British Journal of Nutrition, 2017, 117 (2), 10.1017/S0007114517000058. hal-01794028

HAL Id: hal-01794028 https://amu.hal.science/hal-01794028

Submitted on $17~\mathrm{May}~2018$

HAL is a multi-disciplinary open access archive for the deposit and dissemination of scientific research documents, whether they are published or not. The documents may come from teaching and research institutions in France or abroad, or from public or private research centers. L'archive ouverte pluridisciplinaire **HAL**, est destinée au dépôt et à la diffusion de documents scientifiques de niveau recherche, publiés ou non, émanant des établissements d'enseignement et de recherche français ou étrangers, des laboratoires publics ou privés.

1	Prospective association between consumption frequency of organic food and body
2	weight change, risk of overweight or obesity: Results from the NutriNet-Santé Study
3	Emmanuelle Kesse-Guyot ¹ *, Julia Baudry ¹ , Karen E Assmann ¹ , Pilar Galan ¹ , Serge
4	Hercberg ^{1, 2} , Denis Lairon ³
5	¹ Sorbonne Paris Cité Epidemiology and Statistics Research Center (CRESS), Inserm U1153,
6	Inra U1125, Cnam, Paris 13 University, Nutritional Epidemiology Research Team (EREN), F-
7	93017 Bobigny, France ;
8	² Département de Santé Publique, Hôpital Avicenne, F-93017 Bobigny, France ;
9	³ Aix Marseille Université, NORT ; Insem, UMR S 1062 ; Inra 1260, F-13385 Marseille,
10	France
11	*Correspondence: Email: <u>e.kesse@eren.smbh.univ-paris13.fr</u>
12	Equipe de Recherche en Epidémiologie Nutritionnelle (EREN)
13	SMBH Université Paris 13
14	74 rue Marcel Cachin, 93017 Bobigny, France
15	Running title: Organic food consumption and corpulence
16	Keywords: organic food, obesity, prospective cohort study, weight gain
	Abbreviations: BMI: body mass index, OS: organic score, mPNNS-GS, PCA: Principal
	component analysis

17 Abstract

A lower body mass index (BMI) has been reported among consumers of organic foods but 18 this relationship has never been examined in a prospective design. Our aim was to 19 prospectively investigate the association between the frequency of organic foods and weight 20 21 change. We analyzed data from 62,224 participants of the NutriNet-Santé cohort (78% women, mean age=45y) with information on consumption frequency of organic foods, dietary 22 intake, and repeated anthropometric data. For 16 products, participants reported their 23 consumption frequency of labeled organic foods (never, occasionally, most of the time). An 24 organic score (OS) with a maximum of 32 points was computed. The associations of the OS 25 (modeled as quartiles (Q)) with change in BMI during follow-up (on average 3.1 y), and with 26 27 the risk of overweight and obesity were estimated ANCOVA and multivariable logistic 28 regression. A lower BMI increase was observed across quartiles of the OS (mean difference (95%CI) Q4 versus Q1= - 0.16 (-0.32, -0.01). An increase in OS was associated with a lower 29 risk of overweight and obesity (among non-overweight and non-obese participants at 30 inclusion): odds ratios (95%CI) Q4 versus Q1 were 0.77 (0.68, 0.86) and 0.69 (0.58, 0.82), 31 respectively. Concerning obesity risk, the association was stronger among participants with 32 higher adherence to nutritional guidelines. This study supports a strong protective role of 33 34 consumption frequency of organic foods concerning the risk of overweight and obesity that depends on overall dietary quality. Upon confirmation, these results may contribute to fine-35 36 tune nutritional guidelines by accounting for farming practices in food production.

37 Introduction

38 The global obesity epidemic, involving multifaceted origins, is a major public health issue ⁽¹⁾.

39 Overweight and obesity contribute to the global burden of chronic diseases ⁽²⁾ - especially

40 type 2 diabetes and ischemic cardiovascular diseases $^{(3)}$, along with cancers $^{(4,5)}$.

Among the factors involved in the etiology of obesity, excess caloric consumption, sedentary lifestyle and genetic susceptibility are well-recognized, but there is a growing concern for the identification of novel factors involved such as gut microbiota ⁽⁶⁾ or environmental chemicals

Among the different dietary factors, organic food is of major interest since it presents multiple 45 features that could potentially protect from weight gain and obesity. Notably, compared to 46 47 conventional food, organic food has been suggested to present better nutritional values concerning fatty acids profiles and specific micronutrients ⁽¹¹⁻¹³⁾. Nonetheless, the possible 48 49 implications at an individual level in terms of daily nutrient intake are unknown due to the lack of food composition tables accounting for farming practices. A small number of clinical 50 studies have been conducted that compared specific nutritional biomarkers according to the 51 type of diet (organic or conventional). However, due to short study durations, these studies 52 were not well-equipped to provide reliable results, and findings were inconsistent ⁽¹⁴⁾. 53 54 Organic foods are also characterized by a markedly low level or an absence of pesticide residues, as repeatedly reported in food residue analyses ^(11,14,15) and in experimental studies 55 showing that adopting an organic diet leads to a drastic reduction in pesticide residues and 56 urine metabolites in children and adults (16-20). For most pesticide families (organochlorines 57 (now banned in the EU but still persistent), organophosphates, and pyrethrynoïds), a large 58 number of molecules have been recognized as endocrine disruptors ⁽²¹⁾, leading to possible 59 metabolic disorders ⁽²²⁾. Indeed, a higher exposure to some of these compounds has been 60 associated with a higher risk for obesity or type 2 diabetes in humans $^{(9)}$. 61

62 In this context, consumption of organic foods might contribute to the management of weight gain and of obesity risk. In a recent cross-sectional analysis based on the NutriNet-Santé 63 cohort, we showed that participants identified as regular consumers of organic food, 64 compared to non-consumers, presented reduced odds of being overweight or obese (-36% and 65 -62% in men and -42% and -48% in women, respectively)⁽²³⁾. Comparable findings were 66 reported in the German National Nutrition Survey II (NVS II), a nationwide food 67 consumption study conducted among 13,074 adults: German buyers of organic food exhibited 68 healthier lifestyles compared with non-buyers, and presented lower body weight ⁽²⁴⁾. 69 To the best of our knowledge, no epidemiologic study has yet investigated the prospective 70 relation between consumption frequency of organic foods and the risk of overweight and 71 72 obesity. The main objective of the present study was thus to investigate the longitudinal 73 association between the frequency consumption of organic foods and change in body mass 74 index, the risk of overweight, and the risk of obesity in a very large adult cohort. Given our previous finding that organic food consumers show a higher level of adherence to nutritional 75 recommendations ⁽²³⁾ - especially a higher consumption of fruit and vegetables and a lower 76 consumption of animal products -a secondary objective was to explore a potential modifying 77 effect of the nutritional quality of the diet on the investigated associations. 78

79 Material and methods

80 Study population

The NutriNet-Santé is a web-based prospective observational cohort that was launched in France in May 2009. The objectives, design and methodology have been described elsewhere (25). The study was conducted according to the guidelines laid down in the Declaration of Helsinki and was approved by the Institutional Review Board of the French Institute for Health and Medical Research (IRB Inserm no. 0000388FWA00005831) and the "Commission Nationale de l'Informatique et des Libertés" (CNIL no. 908450 and no. 909216). All subjects
signed an electronic informed consent.

88 Data collection and computation

Volunteers filled in self-administrated questionnaires using a dedicated website at baseline and during follow-up on an approximately monthly basis. The baseline questionnaires, inquired sociodemographic data and lifestyles, health status, physical activity, anthropometrics and diet. These questionnaires were first pilot-tested and then compared to traditional assessment methods or objectively validated ⁽²⁶⁻²⁸⁾.

94 Consumption frequency of organic products data

95 Two months after enrollment, volunteers were asked to provide information on their 96 consumption frequency of 16 labeled organic products (fruit, vegetables, soya, dairy products, meat and fish, eggs, grains and legumes, bread and cereals, flour, vegetable oils and 97 98 condiments, ready-to-eat meals, coffee/tea/herbal wine, tea, biscuits/chocolate/sugar/marmalade, other foods, dietary supplements). Initially, the 99 collection of these data was related to research questions focused on reasons for non-100 consumptions. Consumption frequencies were presented in 8 modalities: (1) most of the time; 101 102 2) occasionally; 3) never ("too expensive"); 4) never ("product not available"); 5) never ("I'm 103 not interested in organic products"); 6) never ("I avoid such products"); 7) never ("for no specific reason"); and 8) "I don't know"). For each product, we allocated two and one points 104 to the "most of the time" and "occasionally" modalities, respectively (and 0 otherwise) - since 105 106 the objective of the present study was to focus on the level of frequency, and not on reasons for non-consumption. The 16 dietary components were summed up to provide an organic 107 score (ranging from 0 to 32). 108

109 Anthropometric data

At enrollment and yearly after, participants were asked to report weight and height assessed during a medical or occupational health examination by a physician, or from selfmeasurement guided by standardized procedures (on flat surface, lightly dressed, and without shoes). Self-reported anthropometric data have been shown to present an elevated concordance with clinical assessment ⁽²⁶⁾.

Body mass index (kg/m2) was calculated as weight divided by the square of height. Subjects

116 were classified as underweight or normal weight (BMI<25), overweight (including obesity;

BMI≥25), or obese (BMI≥30) according to the World Health Organization (WHO) reference
 values ⁽¹⁾.

119 Dietary data and physical activity

At baseline, quantitative dietary intakes were assessed using three 24-hour records (24HR), 120 randomly allocated over a two-week period, including two week-days and one weekend day, 121 using a validated method ^(27,28). Participants reported all foods and beverages consumed at 122 123 each eating occasion. Portion sizes were estimated with the help of photographs, derived from a previously validated picture booklet ⁽²⁹⁾ or directly entered as grams, volumes or purchased 124 units. Since alcohol is only episodically consumed by most individuals, alcohol intake was 125 126 calculated using either the 24HR or a frequency questionnaire for those identified as abstainers in the three 24HR days. Moreover, since fish and seafood are infrequently 127 consumed by many individuals, the weekly consumption of this food group was assessed by a 128 129 specific frequency question. Individual daily mean food consumption was calculated from the three 24HR, weighted for the type of day (week or weekend day). Nutrient intakes were 130 calculated using the NutriNet-Santé composition table ⁽³⁰⁾. Under-reporters were identified 131 and excluded using the validated method developed by Black ⁽³¹⁾. To assess nutritional diet 132 quality, a modified version of the validated PNNS-GS (without physical activity) was 133 computed. This modified score, the mPNNS-GS, reflects adherence to the official French 134

nutritional recommendations ⁽³²⁾. The score includes 12 components: eight refer to foodserving recommendations (fruit and vegetables; starchy foods; whole grain products; dairy
products; meat, eggs and fish; fish and seafood; vegetable fat; water vs soda) and four refer to
moderation in consumption (added fat; salt; sweets; alcohol). Moreover, points are deducted
for overconsumption of salt, added sugars, or when energy intake exceeds the estimated
energy needs by more than 5%.

In order to also account for *a posteriori* dietary patterns, we performed a principal component 141 analysis (PCA), on 31 aggregated food groups. Dietary patterns obtained by PCA are 142 independent linear combinations of the 31 food group consumptions, maximizing the 143 144 explained variance. Two dietary patterns were retained based on Cattel's Scree plots and the 145 interpretability of the factors. Supplementary table S1 presents all factor loading coefficients (corresponding to the correlations between the different food groups and the two dietary 146 patterns) > 0.3. For each participant, the individual pattern score was calculated by summing 147 the intake of the 31 food groups, weighted by their factor loading. 148

149 Covariates

150 At baseline, self-administered questionnaires were used to collect data including age, gender, 151 formal education (\leq high school diploma, high school, post-secondary graduate), occupation (managerial staff, intermediate profession, employee/manual worker, retired, unemployed, 152 153 never employed/homemaker and self-employed), marital status (cohabiting or single), income, number of children and smoking status (never, former and current). Income per 154 155 household unit was calculated by dividing the household's total monthly income by the number of consumption units (CU), using the following coefficients: 1 CU for the first adult 156 in the household, 0.5 CU for all other household members aged 14 or older and 0.3 CU for 157 children under 14⁽³³⁾. The following categories of monthly income were used: <1,200, 1,200-158 159 1,800, 1,800-2,700 and >2,700 euros per household unit.

160 Physical activity was assessed by the International Physical Activity Questionnaire (IPAQ)

161 ⁽³⁴⁾. Metabolic equivalents (MET) measured in minutes per week were computed. The

162 recommended IPAQ categories of physical activity were used: low (<30 min of brisk

163 walking/day), moderate (30 - <60 min/day of brisk walking /day or equivalent) and high (≥ 60

164 min of brisk walking /day or equivalent).

165 The inquired baseline health data included use of medication and self-reported history of 166 diseases (cancer, cardiovascular diseases, hypertension, dyslipidemia and diabetes).

167 Statistical analysis

168 For the present study, we used data from volunteers who were included before June 2014 and

169 initially aged 18-74 y. Among them, we selected those 1) who completed the organic

170 questionnaire, 2) with baseline anthropometric data and at least one measurement during

171 follow-up, 3) with available data to compute PNNS-GS and 4) who were not identified as

172 energy underreporters (Figure 1). Follow-up anthropometric data were collected until June

173 2015.

The participants included (N=62,224) into our analyses were compared to those excluded,
using Mann-Whitney-U tests and Chi square tests.

Baseline characteristics were presented by quartiles of the organic score. Values represent means (±SD) or percentages, and P-values were calculated using linear contrast tests (for continuous variables) or Chi square trend tests (for categorical variables).

179 The association of the organic score with BMI change was assessed by covariance analysis

180 (ANCOVA), modeling change in BMI as a percentage of the baseline value. Mean differences

181 (95% confidence intervals) were presented across quartiles of the organic score. Three

182 different models were performed. The first model was adjusted for baseline age and gender.

183 The second model was further adjusted for year and month of inclusion, follow-up duration,

184 occupation, marital status, education, monthly income per household unit, baseline use of

185 dietary supplements, the mPNNS-GS, PCA-extracted dietary patterns scores, energy intake, physical activity and smoking status. The final model was further adjusted for history of 186 diseases (cancer, cardiovascular diseases, diabetes, hypertension and dyslipidemia). 187 In a second set of analyses, we estimated odds ratios (OR) and 95% confidence intervals (CI) 188 for becoming overweight or obese after exclusion of overweight and obese subjects at 189 baseline, respectively (leading to new study samples of n = 43,301 and n = 56,806). Three 190 191 multivariate logistic regression models were computed with similar covariables to those used 192 in the covariance analysis.

A set of supplementary analyses was performed for the obesity risk outcome. First, stratified 193 194 analyses were conducted according to physical activity level/day brisk walking versus ≥ 30 195 min/day brisk walking or equivalent), use of dietary supplements (yes versus no), tobacco status (never and former smokers versus current smokers), education level, and level of 196 197 adherence to nutritional guidelines (using tertiles of the mPNNS-GS). Second, we used an 198 alternative method of accounting for potential confounder bias: adjustment for a 'propensity' score' that contains information on potential confounders in a combined manner ⁽³⁵⁾. To 199 obtain the propensity score, a multinomial logistic regression model was used to estimate the 200 201 predicted probability of organic food consumption (using quartiles of the 16-point organic score) as a function of a wide range of factors (sociodemographic, health characteristics, food 202 group consumptions). Finally, we used inverse probability weighting to correct the estimates 203 for potential selection bias ⁽³⁶⁾. All tests of statistical significance were two-sided and the type 204 I error was set at 5%. Statistical analyses were performed using SAS® software (version 9.3, 205 206 SAS Institute Inc, Cary, NC, USA).

207 **Results**

208 Comparison of included and excluded participants

Compared with excluded subjects, included subjects (N=62,224) were older, had a higher
education level and income, were less often smokers and less physically active. They also
presented a lower BMI and a higher nutritional quality of the diet (all P-values<0.05, data not
tabulated).

213 Baseline Characteristics of the sample

Baseline characteristics across quartiles of organic score are shown in **Table 1**. Higher levels of the organic score were related to higher proportions of women, participants who were cohabiting, former smokers, physically active participants, individuals reaching a postsecondary educational level, and participants with a high income or occupational level. A positive association was also observed with age, follow-up duration and the mPNNS-GS (reflecting the nutritional quality of the diet), while there was a negative association with energy intake, alcohol consumption and BMI.

Components of the organic score across quartiles of the organic score are shown in Table 2.
In the first quartile (Q1), participants mostly reported no consumption of any organic
products. In the 4th quartile (Q4), participants reported more frequently consuming organic
products, especially products of the following food groups: eggs, starchy food, vegetables,
vegetable oil, fruits and flour.

Then mean follow-up time in our study sample was $3 \cdot 12$ (SD=1 $\cdot 37$) years. Results on the prospective association between the organic score and change in BMI over time are presented in **Table 3**. In the second model, higher organic score levels were related to a substantially lower increase in BMI over time (mean difference Q4 versus Q1= -0.15; confidence interval= -0.31, -0.01, P for trend =0.05). After further adjustment for history of chronic diseases (third model), an even stronger association was observed (mean difference Q4 versus Q1= -0.16; confidence interval= -0.32, -0.01, P for trend =0.04). Results on the prospective association between the organic score and the risk of overweight
and obesity are presented in **Table 4**. In the fully-adjusted model, accounting for
sociodemographic data, lifestyles and history of chronic diseases, a linear decrease in the risk
of overweight was observed across quartiles of the organic score, with risk reduction of 23%
in Q4 compared to Q1. Findings concerning the risk of obesity were similar, with a risk
reduction of 31% in Q4 compared to Q1.

239 Stratified analyses are presented in Figure 2 and in Supplemental Table S2. We observed that the association between the organic score and the risk of obesity was stronger among 240 participants with a higher nutritional quality of the diet. Overall, in the different stratified 241 242 analyses, the association between the organic score and the risk of obesity was observed in 243 each sub-group, except for dietary supplement users and participants with an intermediate education level. Accounting for selection bias via inverse probability weighting did not 244 substantially modify our findings (data not shown). Models with additional adjustment for a 245 propensity score are presented in Supplemental Table S3. Here, the investigated associations 246 were attenuated but remained statistically significant. 247

248 **Discussion**

The present results show, for the first time, a strong negative association between the consumption frequency of organic foods and BMI change over time, as well as a marked reduction of the risk of overweight and obesity.

In stratified analyses, significant associations were observed in almost all investigated subgroups, except for dietary supplement users and participants with an intermediate level of education. Importantly, both participants with a low level of physical activity and participants with a low education level presented a significantly lower risk of obesity with increasing organic food consumption. It is noteworthy that the nutritional quality of the diet, estimated by using an a priori dietary index reflecting adherence to the French nutritional guidelines ⁽³²⁾, appears to be a key effect modifier. Indeed, the strongest associations were observed amongparticipants presenting a high nutritional quality of the diet.

260 Consumption frequency of organic foods and the risk of obesity or overweight

261 We prospectively observed a markedly lower risk of obesity among subjects with a high

262 consumption frequency of organic foods. Previous investigations of data from the NutriNet-

263 Santé cohort have revealed that regular consumers of organic foods presented a healthier diet

and healthier lifestyle characteristics (related to physical activity and tobacco use) as

265 compared to irregular consumers or non-consumers. Moreover, regular consumers of organic

266 foods in the NutriNet-Santé study presented specificities with respect to chronic disease

267 history ^(23,37). After accounting for these potential confounders, the findings of the present

study were partially attenuated but the association remained strong and highly significant,

with a reduction in the risk of obesity of 37% after a 3.1 y follow-up.

270 A similar association was observed for overweight, although the strength of the association

was smaller.

272 Comparison of our results with the findings of other studies

No previous longitudinal study has investigated the association between organic food 273 consumption frequency and weight change or the risk of overweight and obesity, but a 274 275 potential beneficial link between body weight and organic food consumption or purchase has been documented in several cross-sectional studies ^(23,24,38). For instance, a previous 276 investigation of NutriNet-Santé data ⁽²³⁾ has shown that regular organic food consumers 277 showed a markedly lower probability of overweight (excluding obesity) and obesity 278 compared to non-consumers (-36% and -62% in men and -42% and -48% in women, 279 respectively). In addition, in the German National Nutrition Survey II⁽²⁴⁾, as compared to 280 non-buyers of organic food, buyers of organic food presented lower proportions of 281 overweight (35.5% versus 39.2%) and obesity (17.9% versus 22.5%). Our findings are also 282

283 concordant with a small Italian clinical study (including 100 healthy males and 50 males suffering from chronic kidney disease, CDK) that reported a statistically significant reduction 284 in weight among CDK patients after introducing an organic diet for a 2-week period 285 $(85\cdot17\pm13\cdot97 \text{ kg at baseline versus } 79\cdot52\pm10\cdot41 \text{ kg after the 2-week intervention, } p<0\cdot05)^{(39)}$. 286 Overall, the currently available cross-sectional or longitudinal surveys consistently reported 287 an association between higher organic food consumption and a lower BMI. The association 288 with a lower increase in BMI over time that we observed in the present study is of particular 289 interest as it supports a possible role of the organic-based diet in weight management among 290 all subjects, beyond the risk of overweight or obesity among initially normal weight (or 291 292 underweight) individuals.

293 Modulating effect of the nutritional quality of the diet

There is growing evidence supporting the observation that consumers of organic food present 294 a nutritionally healthier diet as well as other beneficial lifestyles such as lower alcohol 295 consumption, no smoking and a higher physical activity level $(^{23,24,40,41)}$. Thus, we 296 hypothesized that the link between consumption of organic foods and body weight might be 297 modulated by the overall nutritional quality of the diet. Indeed, we found that the strongest 298 reduction in obesity risk (related to a higher consumption frequency of organic foods) was 299 300 observed among participants with a healthier diet. Conversely, among participants with a less 301 healthy diet (susceptible to promote weight gain), the association between consumption frequency of organic food and obesity risk was of smaller magnitude. It is well known that 302 unhealthy eating habits play a key role in the etiology of obesity ⁽³⁾. Therefore, the mode of 303 production of the foods consumed may be of secondary importance in these subjects. It is also 304 305 possible that their dietary patterns, which are low in fruit and vegetables, lead to less contaminated diets. 306

Several hypotheses can be proposed to explain the differences in the risk of obesity between organic food consumers and non-consumers who present a healthy diet. Because we adjusted and then stratified for the global nutritional quality of the diet (using the mPNNS-GS score) in our models, it is not probable that our results were biased by the fact that organic food consumers present healthier diets. Other possible explanations include differences in organic food and conventional food with respect to various nutritional compounds.

First, the results of a number of studies argue for a higher concentration of polyunsaturated 313 fatty acids (especially n-3 fatty acids) in organic dairy and meat products; and of antioxidants 314 (especially vitamin C and phenolic compounds) in organic plant foods ⁽¹¹⁻¹³⁾. While observed 315 316 differences in nutrient content can vary by about 10-68% at the food level (not accounted for in our study), it is possible that the overall variations of nutrient intake in an organic diet is 317 sufficient to affect weight management ^(42,43). However, this remains to be further evaluated. 318 319 Another hypothesis is related to the fact that individuals with a higher adherence to the French nutritional guidelines tend to consume more plant-based foods. It is well known that plant 320 foods are frequently contaminated by various pesticide residues (about 45% of the tested 321 samples in Europe)⁽⁴⁴⁾ because they are heavily sprayed with pesticides during conventional 322 agricultural production and storage. This hypothesis is in line with findings of human surveys 323 that have related obesity and type 2 diabetes to pesticide exposure (7,9,10,45,46). Thus, unlike the 324 consumption of pesticide-free or only slightly contaminated plant products ^(11,14,15), a high 325 consumption of conventionally grown plant foods may be related to adverse health effects 326 related to higher pesticide exposure. 327

Indeed, replacing conventional food by organic food has repeatedly been shown to drastically reduce the level of organophosphate residues in human urines ⁽¹⁶⁻²⁰⁾. A specific example of a potential adverse health effect of contaminated fruit and vegetables is that high consumers of conventional or contaminated fruits and vegetables presented a particularly low semen quality ^(47,48). This reinforces the concept that different dietary profiles (with various degrees of intake
 of contaminated food) lead to different levels of exposure to "obesogen" chemicals ⁽⁹⁾, but this
 hypothesis needs to be investigated in future biomonitoring-based studies that compare
 organic and conventional diets with various dietary profiles.

336 Potential mechanistic pathways explaining the association between organic food consumption
337 and body weight

338 Our findings may be interpreted in light of the reduced exposure to pesticides among organic

food consumers. Pesticides (prohibited in organic farming when they are synthetic) often

340 present endocrine-disrupting properties that cause developmental and reproductive

341 abnormalities via the modification of signaling processes ⁽⁴⁶⁾. Besides, new scientific research

342 argues for a role of pesticides in metabolic disruption, ⁽²²⁾ leading to obesity and type 2

343 diabetes ⁽⁹⁾. Mechanistic pathways depend on the type of pesticides. Previous studies have

344 shown an alteration of glucose and lipid metabolism by organochlorines ⁽⁴⁹⁾. Moreover,

345 organochlorines have been shown to affect the control of adipogenesis through an alteration

of glucose transport and glycolysis, an alteration of mitochondrial activity, and of fatty acid

347 oxidation ⁽⁴⁹⁾. Organophosphates have been shown to alter carbohydrate and lipid metabolism

348 by disrupting glucose homeostasis ⁽⁴⁹⁾. In addition, some pesticides have been shown to affect

the regulation of eating behavior and the differentiation of adipocytes ⁽⁵⁰⁾.

350 Further studies are necessary to better evaluate metabolic disruption and the "obesogen"

351 capacity of endocrine-disrupting chemicals contained in conventional foods.

352 Strengths and limitations

The main limitation of our study is that it is based on self-reported weight and height data.However, these data have been shown to have a good concordance with data from clinical

355 assessments in a validation study ⁽²⁶⁾. This validation study showed high intraclass correlation

356 coefficients, ranging from 0.94 for height to 0.99 for weight, and the concordance for BMI

classification was 93% (sensibility=88% and specificity=99%). Second, the generalizability 357 of our findings is limited since participants were volunteers involved in a long-term cohort 358 focused on nutrition and health. The individuals included in our study sample are thus likely 359 to be particularly health conscious. A final limitation pertains to the difficulty to disentangle 360 361 the role of overall dietary patterns from the role of organic food consumption, despite of the extensive adjustment and stratification made. Since the design of our study is observational, 362 residual confounding cannot be ruled out. In particular, it is likely that unmeasured or only 363 indirectly measured factors, including genetic factors, ethnicity, environmental (e.g. food or 364 built environment) or psychological factors (e.g. occupational stress) may modify the 365 366 association between organic food consumption and obesity.

Our study also presents important strengths. First, the rich and accurately collected data permitted to account for a broad range of potential confounders, including lifestyles and health outcomes. Moreover, the very large sample size of our study enabled us to conduct statistically powerful stratified analyses. Another important strength is the prospective design of our analysis that implies a high level of evidence. Finally, the availability of accurate dietary data allowed us to adjust for the nutritional quality of the diet, using a validated dietary index.

374 Conclusion

This study, based on data collected in a very large prospective cohort, is the first to support a prospective relation between consumption frequency of organic foods and body weight change, as well as a strong negative association with the risk of overweight and obesity. The overall nutritional quality of the diet may exert a modulating effect in these relationships, with a stronger effect observed among those presenting a healthy plant-based diet. Further studies, especially studies based on quantitative organic consumption data taking into account a diversity of dietary profiles (plant based, western etc.), are needed to confirm these results. If

382	confirmed, these findings are of major interest from a public health point of view, as they
383	reinforce the need to fine-tune nutritional guidelines by accounting for the mode of food
384	production.

385 Acknowledgment

- 386 The authors Nathalie Arnault, Stephen Besseau, Laurent Bourhis, Julien Allègre, Than Duong Van,
- 387 Younes Esseddik, Cédric Agaesse, Claudia Chahine, Paul Flanzy, Mac Rakotondrazafy and Fabien
- 388 Szabo for their technical contribution to the NutriNet-Santé study.

Financial support

- 390 The NutriNet-Santé study is supported by the French Ministry of Health (DGS), the French
- 391 Institute for Health Surveillance (InVS), the National Institute for Prevention and Health
- 392 Education (INPES), the Foundation for Medical Research (FRM), the National Institute for
- 393 Health and Medical Research (INSERM), the National Institute for Agricultural Research
- 394 (INRA), the National Conservatory of Arts and Crafts (CNAM), and the University of Paris
- **395 13**.

396 Conflict of Interest

397 None of the authors declares any conflict of interest.

398 Authorship

- 399 EKG: conducted the literature review and drafted the manuscript; EKG: performed analyses;
- 400 EKG, JB, KEA, PG, SH and DL: were involved in the interpretation of results and critically
- 401 reviewed the manuscript; and SH, PG and EKG: were responsible for the development of the
- 402 design and the protocol of the study. All authors read and approved the final manuscript.

403		Reference List
404		
405	1.	WHO (2000) Obesity: preventing and managing the global epidemic., vol. 894: World Health
406		Organization Technical Report Series.
407	2.	WHO (2009) Global Health Risks. Geneva: WHO Technical Report.
408	3.	WHO Europe (2007) The challenge of obesity in the WHO European Region.
409	4	Latino-Martel P. Cottet V. Druesne-Pecollo N <i>et al.</i> (2016) Alcoholic beverages obesity
410		physical activity and other nutritional factors and cancer risk. A review of the evidence
411		Crit Rev Oncol Hematol 99 , 308-323.
412	5.	American Institute for Cancer Research / World Cancer Research Fund (2007) Food, Nutrition,
413		<i>Physical activity, and the Prevention of Cancer: a global perspective.</i> Washington, DC:
414		American Institute for Cancer Research.
415	6.	Hansen TH, Gobel RJ, Hansen T <i>et al.</i> (2015) The gut microbiome in cardio-metabolic health.
416		Genome Med 7, 33.
417	7	Lee DH Porta M Jacobs DR Jr. et al. (2014) Chlorinated persistent organic pollutants obesity
418	7.	and type 2 diabetes. <i>Endocr Rev</i> 35 , 557-601.
419	8	Snedeker SM & Hay AG (2012) Do interactions between out ecology and environmental
420	0.	chemicals contribute to obesity and diabetes? <i>Environ Health Perspect</i> 120 , 332-339.
421	9.	Thayer KA, Heindel JJ, Bucher JR et al. (2012) Role of environmental chemicals in diabetes
422		and obesity: a National Toxicology Program workshop review. Environ Health Perspect
423		120 , 779-789.
424	10.	Lubrano C, Genovesi G, Specchia P et al. (2013) Obesity and metabolic comorbidities:
425		environmental diseases? Oxid Med Cell Longev 2013, 640673.
426	11.	Baranski M, Srednicka-Tober D, Volakakis N et al. (2014) Higher antioxidant and lower
427		cadmium concentrations and lower incidence of pesticide residues in organically grown
428		crops: a systematic literature review and meta-analyses. Br J Nutr 112, 794-811.
429	12.	Srednicka-Tober D, Baranski M, Seal C et al. (2016) Composition differences between organic
430		and conventional meat: a systematic literature review and meta-analysis. Br J Nutr, 1-
431		18.
432	13.	Srednicka-Tober D, Baranski M, Seal CJ et al. (2016) Higher PUFA and n-3 PUFA, conjugated
433		linoleic acid, alpha-tocopherol and iron, but lower iodine and selenium concentrations in
434		organic milk: a systematic literature review and meta- and redundancy analyses. Br J
435		Nutr, 1-18.
436	14	Smith-Spangler C. Brandeau ML, Hunter GE et al. (2012) Are organic foods safer or healthier
437		than conventional alternatives?: a systematic review. Ann Intern Med 157, 348-366.
438	15	Lairon D (2010) Nutritional quality and safety of organic food: a review Agron Sustain Dev 30
439	10.	33-41.

440 441 442	16.	Curl CL, Fenske RA & Elgethun K (2003) Organophosphorus pesticide exposure of urban and suburban preschool children with organic and conventional diets. <i>Environ Health</i> <i>Perspect</i> 111, 377-382.
443 444	17.	Lu C, Toepel K, Irish R <i>et al.</i> (2006) Organic diets significantly lower children's dietary exposure to organophosphorus pesticides. <i>Environ Health Perspect</i> 114 , 260-263.
445 446 447	18.	Bradman A, Quiros-Alcala L, Castorina R et al. (2015) Effect of Organic Diet Intervention on Pesticide Exposures in Young Children Living in Low-Income Urban and Agricultural Communities. Environ Health Perspect 123, 1086-1093.
448 449	19.	Oates L, Cohen M, Braun L <i>et al.</i> (2014) Reduction in urinary organophosphate pesticide metabolites in adults after a week-long organic diet. <i>Environ Res</i> 132 , 105-111.
450 451 452	20.	Curl CL, Beresford SA, Fenske RA <i>et al.</i> (2015) Estimating pesticide exposure from dietary intake and organic food choices: the Multi-Ethnic Study of Atherosclerosis (MESA). <i>Environ Health Perspect</i> 123 , 475-483.
453 454	21.	Mnif W, Hassine AI, Bouaziz A et al. (2011) Effect of endocrine disruptor pesticides: a review. Int J Environ Res Public Health 8, 2265-2303.
455 456	22.	Casals-Casas C & Desvergne B (2011) Endocrine disruptors: from endocrine to metabolic disruption. <i>Annu Rev Physiol</i> 73 , 135-162.
457 458 459	23.	Kesse-Guyot E, Peneau S, Mejean C <i>et al.</i> (2013) Profiles of organic food consumers in a large sample of French adults: results from the Nutrinet-Sante cohort study. <i>PLoS One</i> 8 , e76998.
460 461 462	24.	Eisinger-Watzl M, Wittig F, Heuer T <i>et al.</i> (2015) Customers Purchasing Organic Food - Do They Live Healthier? Results of the German National Nutrition Survey II. <i>Eur J Nutr</i> <i>Food Saf</i> 5 , 59-71.
463 464 465	25.	Hercberg S, Castetbon K, Czernichow S <i>et al.</i> (2010) The Nutrinet-Sante Study: a web-based prospective study on the relationship between nutrition and health and determinants of dietary patterns and nutritional status. <i>BMC Public Health</i> 10 , 242.
466 467	26.	Lassale C, Peneau S, Touvier M et al. (2013) Validity of web-based self-reported weight and height: results of the Nutrinet-Sante study. J Med Internet Res 15, e152.
468 469 470	27.	Lassale C, Castetbon K, Laporte F <i>et al.</i> (2015) Validation of a Web-based, self-administered, non-consecutive-day dietary record tool against urinary biomarkers. <i>Br J Nutr</i> 113 , 953-962.
471 472 473	28.	Lassale C, Castetbon K, Laporte F <i>et al.</i> (2015) Correlations between Fruit, Vegetables, Fish, Vitamins, and Fatty Acids Estimated by Web-Based Nonconsecutive Dietary Records and Respective Biomarkers of Nutritional Status. <i>J Acad Nutr Diet</i> .
474 475 476	29.	Le Moullec N, Deheeger M, Preziosi P <i>et al.</i> (1996) Validation du manuel photos utilisé pour l'enquête alimentaire de l'étude SU.VI.MAX. <i>Cahier de Nutrition et de Diététique</i> 31 , 158-164.
477 478	30.	NutriNet-Santé coordination (2013) <i>Table de composition des aliments - Etude NutriNet-Santé</i> . Paris: Economica.

- 479 31. Black AE (2000) Critical evaluation of energy intake using the Goldberg cut-off for energy
 480 intake:basal metabolic rate. A practical guide to its calculation, use and limitations. *Int J* 481 *Obes Relat Metab Disord* 24, 1119-1130.
- 482 32. Estaquio C, Kesse-Guyot E, Deschamps V *et al.* (2009) Adherence to the French Programme
 483 National Nutrition Sante Guideline Score is associated with better nutrient intake and
 484 nutritional status. *J Am Diet Assoc* 109, 1031-1041.
- 485 33. INSEE (Institut National de la Statistique et des Etudes Economiques) website (2012).
 486 <u>http://www.insee.fr/en/methodes/.</u>
- 487 34. Hagstromer M, Oja P & Sjostrom M (2006) The International Physical Activity Questionnaire
 488 (IPAQ): a study of concurrent and construct validity. *Public Health Nutr* 9, 755-762.
- 489 35. Ali MS, Groenwold RH & Klungel OH (2016) Best (but oft-forgotten) practices: propensity
 490 score methods in clinical nutrition research. *Am J Clin Nutr* 104, 247-258.
- 491 36. Seaman SR & White IR (2011) Review of inverse probability weighting for dealing with
 492 missing data. *Stat Methods Med Res.*
- 493 37. Baudry J, Mejean C, Peneau S *et al.* (2015) Health and dietary traits of organic food consumers:
 494 results from the NutriNet-Sante study. *Br J Nutr*, 1-10.
- 495 38. Torjusen H, Brantsaeter AL, Haugen M *et al.* (2010) Characteristics associated with organic
 496 food consumption during pregnancy; data from a large cohort of pregnant women in
 497 Norway. *BMC Public Health* 10, 775.
- 498 39. De Lorenzo A, Noce A, Bigioni M *et al.* (2010) The effects of Italian Mediterranean organic
 499 diet (IMOD) on health status. *Curr Pharm Des* 16, 814-824.
- 40. Oates L, Cohen M & Braun L (2012) Characteristics and consumption patterns of Australian
 organic consumers. J Sci Food Agric 92, 2782-2787.
- 41. Petersen SB, Rasmussen MA, Strom M *et al.* (2013) Sociodemographic characteristics and food
 habits of organic consumers--a study from the Danish National Birth Cohort. *Public Health Nutr* 16, 1810-1819.
- Lorente-Cebrian S, Costa AG, Navas-Carretero S *et al.* (2013) Role of omega-3 fatty acids in
 obesity, metabolic syndrome, and cardiovascular diseases: a review of the evidence. J
 Physiol Biochem 69, 633-651.
- 508 43. Meydani M & Hasan ST (2010) Dietary polyphenols and obesity. *Nutrients* 2, 737-751.
- 509 44. EFSA (European Food Safety Authority) (2015) The 2013 European Union Report on Pesticide
 510 Residues in Food. *EFSA J* 13.
- 511 45. Lee DH, Steffes MW, Sjodin A *et al.* (2011) Low dose organochlorine pesticides and
 512 polychlorinated biphenyls predict obesity, dyslipidemia, and insulin resistance among
 513 people free of diabetes. *PLoS One* 6, e15977.
- 46. Heindel JJ, Newbold R & Schug TT (2015) Endocrine disruptors and obesity. *Nat Rev Endocrinol* 11, 653-661.
- 516 47. Chiu YH, Afeiche MC, Gaskins AJ *et al.* (2015) Fruit and vegetable intake and their pesticide
 517 residues in relation to semen quality among men from a fertility clinic. *Hum Reprod* 30, 1342-1351.

519 48 520	Juhler RK, Larsen SB, Meyer O <i>et al.</i> (1999) Human semen quality in relation to dietary pesticide exposure and organic diet. <i>Arch Environ Contam Toxicol</i> 37 , 415-423.
521 49 522 523	Androutsopoulos VP, Hernandez AF, Liesivuori J <i>et al.</i> (2013) A mechanistic overview of health associated effects of low levels of organochlorine and organophosphorous pesticides. <i>Toxicology</i> 307 , 89-94.
524 50 525 526 527	Mostafalou S & Abdollahi M (2013) Pesticides and human chronic diseases: evidences, mechanisms, and perspectives. <i>Toxicol Appl Pharmacol</i> 268 , 157-177.

500		6.41 .	•	
٦ 4 4	I able 2 Components (nt the arganic scare	across organic score	auartues Nutrinet-Nante
555	Table 2 Components	of the of game score	across of game score	qual mes, run mer Sante

534 study, N=62,224^{*}

	Q1	Q2	Q3	Q4	P [†]
Fruits	0.06 (0.25)	0.57 (0.52)	0.95 (0.41)	1.49 (0.53)	<0.0001
Vegetables	0.07 (0.27)	0.59 (0.55)	0.97 (0.48)	1.51 (0.54)	<0.0001
Rice, pasta, other cereals	0.06 (0.23)	0.42 (0.51)	0.90 (0.50)	1.62 (0.52)	<0.0001
Breads	0.05 (0.22)	0.36 (0.51)	0.82 (0.53)	1.44 (0.59)	<0.0001
Flour	0.04 (0.20)	0.23 (0.45)	0.67 (0.62)	1.47 (0.65)	<0.0001
Vegetable oils	0.04 (0.22)	0.27 (0.51)	0.74 (0.65)	1.58 (0.58)	<0.0001
Dairy products	0.05 (0.23)	0.40 (0.54)	0.86 (0.58)	1.42 (0.65)	<0.0001
Meats & seafoods	0.02 (0.14)	0.19 (0.41)	0.50 (0.54)	0.92 (0.66)	<0.0001
Eggs	0.13 (0.36)	0.70 (0.70)	1.19 (0.70)	1.72 (0.54)	<0.0001
Soja products	0.06 (0.24)	0.20 (0.44)	0.40 (0.59)	0.88 (0.76)	<0.0001
Sweet products	0.03 (0.19)	0.33 (0.48)	0.76 (0.50)	1.38 (0.58)	<0.0001
Tea, coffee, herbal tea	0.04 (0.19)	0.27 (0.47)	0.66 (0.59)	1.33 (0.67)	<0.0001
Wines	0.03 (0.17)	0.14 (0.35)	0.29 (0.47)	0.65 (0.64)	<0.0001
Ready-to-use dishes	0.01 (0.08)	0.06 (0.24)	0.19 (0.39)	0.39 (0.54)	<0.0001
Dietary supplements	0.03 (0.18)	0.13 (0.36)	0.25 (0.49)	0.56 (0.68)	<0.0001
Other dietary items	0.01 (0.12)	0.09 (0.30)	0.24 (0.45)	0.80 (0.80)	<0.0001

535 Q ; Quartile

⁵³⁶ ^{*}Values are means (SD) of sub-score given that non-consumption, occasional, regular

537 consumption are coded 0, 1, 2, respectively.

⁵³⁸ [†]P for linear trend

539 Table 3 Association between organic score in quartile and BMI change over time,

				trend
·00 (ref)	0.01 (-0.13, 0.15)	-0.06 (-0.20, 0.08)	-0.34 (-0.49, -0.20)	<0.0001
·00 (ref)	0.03 (-0.11, 0.17)	0.01 (-0.13, 0.16)	-0.15 (-0.31, -0.00)	0.05
·00 (ref)	0.03 (-0.12, 0.17)	0.01 (-0.14, 0.15)	-0.16 (-0.32, -0.01)	0.04
	·00 (ref) ·00 (ref) ·00 (ref)	$\cdot 00 \text{ (ref)}$ $0 \cdot 01 \text{ (-}0 \cdot 13, 0 \cdot 15)$ $\cdot 00 \text{ (ref)}$ $0 \cdot 03 \text{ (-}0 \cdot 11, 0 \cdot 17)$ $\cdot 00 \text{ (ref)}$ $0 \cdot 03 \text{ (-}0 \cdot 12, 0 \cdot 17)$	$\cdot 00 \text{ (ref)}$ $0 \cdot 01 (-0 \cdot 13, 0 \cdot 15)$ $-0 \cdot 06 (-0 \cdot 20, 0 \cdot 08)$ $\cdot 00 \text{ (ref)}$ $0 \cdot 03 (-0 \cdot 11, 0 \cdot 17)$ $0 \cdot 01 (-0 \cdot 13, 0 \cdot 16)$ $\cdot 00 \text{ (ref)}$ $0 \cdot 03 (-0 \cdot 12, 0 \cdot 17)$ $0 \cdot 01 (-0 \cdot 14, 0 \cdot 15)$	$\cdot 00 \text{ (ref)}$ $0 \cdot 01 (-0 \cdot 13, 0 \cdot 15)$ $-0 \cdot 06 (-0 \cdot 20, 0 \cdot 08)$ $-0 \cdot 34 (-0 \cdot 49, -0 \cdot 20)$ $\cdot 00 \text{ (ref)}$ $0 \cdot 03 (-0 \cdot 11, 0 \cdot 17)$ $0 \cdot 01 (-0 \cdot 13, 0 \cdot 16)$ $-0 \cdot 15 (-0 \cdot 31, -0 \cdot 00)$ $\cdot 00 \text{ (ref)}$ $0 \cdot 03 (-0 \cdot 12, 0 \cdot 17)$ $0 \cdot 01 (-0 \cdot 14, 0 \cdot 15)$ $-0 \cdot 16 (-0 \cdot 32, -0 \cdot 01)$

540 Nutrinet-Santé study, 2009-2015, N=62,224^{*}

541 **Q; Quartile**

⁵⁴² ^{*}Values are mean differences (95% confident intervals). A negative value (-0·xx) indicates

that the observed increase (expressed as a percentage of the initial anthropometric marker)

544 was lower of $0 \cdot xx$ in the respective quartile than in quartile 1 (ref).

[†]Model 1 is adjusted for age and gender.

^{*}Model 2 is model 1 further adjusted for month and year of inclusion, duration of follow-up,

547 occupation, marital status, education, monthly income per unit, dietary supplement use,

548 mPNNS-GS, PCA-extracted dietary patterns scores, energy intake, physical activity and

549 tobacco status.

[§]Model 3 is model 2 further adjusted for the history of chronic diseases.

551 Table 4 Prospective association between quartiles of the organic score and overweight or

	Model	Q1	Q2	Q3	Q4	P for
						trend
Overweight	(N=3,259/40,042)					
	Model 1 [†]	1.00	0.94 (0.86-1.04)	0.83 (0.75-0.91)	0.62 (0.56-0.69)	<.0001
	Model 2 [‡]	1.00	1.00 (0.91, 1.10)	0.93 (0.84, 1.02)	0.75 (0.67, 0.84)	<.0001
	Model 3 [§]	1.00	1.00 (0.91, 1.10)	0.93 (0.84, 1.03)	0.77 (0.68, 0.86)	<-0001
Obesity	(1,337/55,469)					
	Model 1 [†]	1.00	0.87 (0.75, 1.01)	0.79 (0.68, 0.91)	0.52 (0.45, 0.61)	<.0001
	Model 2 [‡]	1.00	0.93 (0.80, 1.08)	0.90 (0.78, 1.04)	0.66 (0.55, 0.78)	<.0001
	Model 3 [§]	1.00	0.94 (0.81, 1.09)	0.92 (0.79, 1.06)	0.69 (0.58, 0.82)	0.0001

552 obesity risk, Nutrinet-Santé study, 2009-2015^{*}.

⁵⁵³ [•]Overweight (including obesity) and obesity analyses were performed among participants who ⁵⁵⁴ were not overweight or obese at inclusion, respectively. Values are odds ratios (95% confident

555 intervals), using 1^{st} quartile Q1 as the reference.

[†]Model 1 is adjusted for age and gender.

⁵⁵⁷ [‡]Model 2 is model 1 further adjusted for month and year of inclusion, delay of follow-up,

occupation, marital status, education, monthly income per unit, dietary supplement use,

559 mPNNS-GS, PCA-extracted dietary patterns scores, energy intake physical activity and

560 tobacco status.

[§]Model 3 is model 2 further adjusted for the history of chronic diseases.

Variable	Factor1	Factor2
Vegetables	0.65	0.14
Fruit	0.45	0.06
Broth	0.43	0.17
Coffee, tea, tisane	0.43	0.01
Soda	-0.42	0.06
Croissants	-0.37	0.03
Nuts	0.36	0.04
Fat and sweet products	-0.32	0.07
Dried fruits	0.33	-0.03
Snacks	-0.28	0.23
Soya products	0.28	-0.16
Meat and meat products	-0.28	0.39
Seafood	0.27	-0.02
Dairy desserts	-0.22	0.04
Bread and breakfast cereals	0.22	0.54
Cakes and pastries	-0.18	0.15
Alcoholic beverages	-0.18	0.34
Margarine	0.16	0.12
Flour	0.13	0.06
Eggs	0.12	0.02
Animal Fat	0.09	0.41
Dairy products	0.07	-0.09
Cheese	-0.02	0.53
Potatoes	0.05	0.41
Vegetable oils	0.03	0.28
Starchy food	0.03	0.07
Meal substitutes	0.02	-0.16
Sauces	-0.01	0.23
Sweet products	-0.01	0.27
Water and non-sweet bevarages	0.01	0.14
Wine	0.01	0.42

*Absolute values < 0.3 are not displayed in the table.

564 Supplemental Table S2 Prospective association between quartiles of the organic score

	Q1	Q2	Q3	Q4	P for
					trend
Education					
< high school diploma	$1 \cdot 00$	1.21 (0.91, 1.61)	0.88 (0.65, 1.19)	0.73 (0.52, 1.04)	0.04
High school diploma	$1 \cdot 00$	0.80 (0.56, 1.14)	0.97 (0.69, 1.35)	0.82 (0.55, 1.20)	0.48
Post-secondary graduate	$1 \cdot 00$	0.95 (0.78, 1.15)	0.92 (0.75, 1.12)	0.69 (0.55, 0.87)	0.003
Physical activity					
<30 min/day brisk walking	$1 \cdot 00$	0.68 (0.51, 0.90)	0.76 (0.58, 1.01)	0.53 (0.38, 0.74)	0.001
\geq 30 min/day brisk walking or equivalent	$1 \cdot 00$	0.97 (0.80, 1.18)	0.90 (0.73, 1.10)	0.74 (0.59, 0.92)	0.01
Smoking status					
Never or former smokers	$1 \cdot 00$	0.95 (0.81, 1.11)	0.90 (0.76, 1.06)	0.72 (0.59, 0.86)	0.001
Current smokers	$1 \cdot 00$	0.74 (0.51, 1.06)	0.95 (0.65, 1.38)	0.55 (0.36, 0.86)	0.04
Dietary supplement users					
yes	$1 \cdot 00$	1.16 (0.94, 1.45)	1.05 (0.84, 1.32)	0.79 (0.61, 1.03)	0.11
no	1.00	0.92 (0.75, 1.12)	0.75 (0.60, 0.95)	0.67 (0.52, 0.85)	0.0003

⁵⁶⁵ and obesity risk, stratified analyses, Nutrinet-Santé study, 2009-2015^{*,†}.

^{*}Obesity analyses were performed among participants who were not obese at inclusion

567 (N=56,806). Values are odds ratios (95% confident intervals), using 1st quartile Q1 as the 568 reference.

⁵⁶⁹ [†]Models are adjusted for age, gender, month and year of inclusion, delay of follow-up,

570 occupation, marital status, education, monthly income per unit, dietary supplement use,

571 mPNNS-GS, PCA-extracted dietary patterns scores, energy intake physical activity, tobacco

572 status and for the history of chronic diseases.

573 Supplemental Table S3 Prospective association between quartiles of the organic score

Model	Q1	Q2	Q3	Q4	P for
					trend
+	1 00				0001
Model 1 ⁺	1.00	0.87(0.81, 0.94)	0.79(0.73, 0.85)	0.52(0.48, 0.57)	<.0001
Model 2 [‡]	$1 \cdot 00$	0.93 (0.87, 1.01)	0.91 (0.84, 0.98)	0.67 (0.61, 0.72)	<.0001
Model 3 [§]	$1 \cdot 00$	0.95 (0.88, 1.02)	0.92 (0.86, 0.99)	0.70 (0.64, 0.76)	<.0001

574 and obesity risk, sensitive analysis, Nutrinet-Santé study, 2009-2015^{*}.

^{*}Obesity analyses were performed among participants who were not obese at inclusion

576 (N=56,806). Values are odds ratios (95% confident intervals), using 1st quartile Q1 as the 577 reference.

⁵⁷⁸ [†]Model 1 is adjusted for age and gender and propensity score.

⁵⁷⁹ [‡]Model 2 is model 1 further adjusted for month and year of inclusion, delay of follow-up,

580 occupation, marital status, education, monthly income per unit, dietary supplement use,

581 mPNNS-GS, PCA-extracted dietary patterns scores, energy intake physical activity and

582 tobacco status.

[§]Model 3 is model 2 further adjusted for the history of chronic diseases.

584 Figure 1. Participants of the NutriNet-Santé selected for the present analyses, 2009–2015



587 Figure 2. Prospective association between organic score in quartile and risk of obesity

588 stratified according the nutritional quality of the diet, Nutrinet-Santé, 2009-2015^{*}



Figure 2. Prospective association between organic score in quartile and risk of obesity stratified according the nutritional quality of the diet, Nutrinet-Santé, 2009-2015* *Values are odds ratios (95% confident intervals) using the 1st quartile (Q1) as the reference, adjusted for age, gender, month and year of inclusion, delay of follow-up, occupation, marital status, education, monthly income per unit, dietary supplement use, mPNNS-GS, PCA-extracted dietary patterns scores energy intake, physical activity, tobacco status and history of chronic diseases.

589

- ^{*}Values are odds ratios (95% confident intervals) using the 1st quartile (Q1) as the reference, adjusted for age,
- gender, month and year of inclusion, delay of follow-up, occupation, marital status, education, monthly income
 per unit, dietary supplement use, mPNNS-GS, PCA-extracted dietary patterns scores energy intake, physical
- activity, tobacco status and history of chronic diseases.

	Q1	Q2	Q3	Q4	\mathbf{P}^{\dagger}
N	15,245	16,249	15,807	14,923	
Organic score max=32	0.73 (0.82)	4.96 (1.41)	10.36 (1.68)	19.15 (4.21)	<0.0001
Female (%)	73.45	77.94	78.71	81.63	<0.0001
Age (years)	44.16 (15.34)	44.36 (14.83)	45.62 (14.29)	46.72 (13.29)	<0.0001
Follow-up duration (days)	1140.34 (504.64)	1157.01 (500.06)	1144.88 (501.80)	1118.12 (499.69)	<0.0001
Education (%)					<0.0001
Unidentified	0.68	0.66	0.58	0.82	
< high school diploma	23.68	20.09	17.21	15.22	
High school diploma	19.12	17.01	15-44	13.99	
Post-secondary graduate	56.52	62.24	66.76	69.97	
Monthly income per unit household unit in \in (%)					<0.0001
Missing	12.06	11.50	10.62	9.35	
900-1200	19.48	16.13	13.26	12.36	
1200-1800	26.76	25.15	22.67	23.09	
1800-2700	21.66	23.72	24.89	26.20	
>2700	20.02	23.50	28.56	29.00	

528 Table 1 Baseline characteristics of the sample across organic score quartiles, Nutrinet-Santé study, 2009-2014, N=62,224^{*}

Occupational categories (%)

23

Non employed	5.60	5.35	5.18	5.84	
Retired	21.66	20.35	21.40	20.31	
Employee/Manual worker	23.22	19.82	16.67	14.29	
Intermediate profession	16.20	17.44	17.42	18.53	
Managerial staff	17.82	21.47	25.48	28.26	
Never employed	13.93	13.92	12.17	10.43	
Craftsman, shopkeeper, business owner,	1.58	1.66	1.68	2.34	
farmer					
Dietary supplement use (%)	35.52	45.55	51.58	62.13	<0.0001
Cohabiting (%)	80.89	81.88	83.22	86.32	<0.0001
Tobacco status (%)					<0.0001
Former smoker	33.05	33.09	35.51	37.69	
Current smoker	15.29	15.31	14.46	13.12	
Never smoker	51.66	51.60	50.03	49.19	
Physical activity (%)					<0.0001
Missing	15.12	14.36	12.85	11.96	
Low	15.12	14.36	12.85	11.96	
Medium	27.65	27.92	30.04	31.9	

High	33.67	37.15	38.33	39.5	
Energy intake (kcal/d)	1942.90 (514.75)	1914.76 (487.92)	1908.50 (491.42)	1894-24 (474-50)	<0.0001
% Carbohydrates	42.59 (6.89)	42.93 (6.81)	43.05 (6.94)	43.43 (7.11)	<0.0001
% Lipids	39.22 (6.61)	38.95 (6.46)	38.96 (6.58)	39.45 (6.72)	0.0017
% Proteins	17.87 (3.95)	17.80 (3.92)	17.67 (4.01)	16.81 (3.92)	<0.0001
Alcohol consumption (g/d)	8.65 (14.11)	8.27 (13.16)	8.22 (12.23)	7.63 (11.20)	<0.0001
mPNNS-GS	7.60 (1.62)	7.87 (1.62)	8.15 (1.60)	8.44 (1.58)	<0.0001
Body mass index (kg/m ²) at baseline	24.50 (4.87)	23.96 (4.56)	23.71 (4.31)	23.00 (3.90)	<0.0001
Obesity [‡] (%) at baseline	11.79	9.42	7.96	5.58	<0.0001
Body mass index (kg/m ²) at follow-up	24.72 (4.92)	24.17 (4.62)	23.91 (4.40)	23.13 (4.00)	<0.0001
Obesity [‡] (%) at follow-up	12.62	10.04	8.78	6.06	<0.0001

529 Abbreviations: Q; Quartile

^{*}All variables were assessed at baseline, except when listed as "at follow-up"

- ⁵³¹ [†]P for linear contrast
- 532 ^{*}Body mass index \geq 30 kg/m²