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# Dietary behaviors influence inflammatory markers: results from the CoLaus study

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1 **DIETARY BEHAVIOURS INFLUENCE INFLAMMATORY MARKERS: RESULTS FROM THE COLAUS STUDY.**

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28

29 **Abstract**

30 **Background:** The effect of different dietary markers on inflammatory markers has seldom been  
31 assessed in a general population setting. We assessed the effect of single foods, nutrients, dietary  
32 patterns and scores on inflammatory markers (CRP, IL-6, TNF- $\alpha$  and leucocyte count).

33 **Methods:** Cross-sectional study including 4027 participants (46.5% men, 57.2 $\pm$ 10.2 years) in Lausanne,  
34 Switzerland (CoLaus study). Dietary intake was collected between 2009-2012 using a semi-quantitative  
35 food frequency questionnaire. Besides single foods and nutrients, three naive (using principal  
36 components analysis - PCA) and four oriented (Mediterranean, Alternative Healthy Eating Index - AHEI)  
37 dietary scores were used.

38 **Results:** Three dietary patterns explaining 20% of total variance were obtained and named "Meat &  
39 chips" (positive loadings for meat and French fries); "Pastries & fat" (positive loadings for hard fats,  
40 pastries and sugar) and "Fruits & vegetables" (positive loadings for fruits & vegetables). After  
41 multivariate adjustment on total energy intake, gender and other socio-demographic factors, fruit  
42 intake, the "Fruits & vegetables" pattern, the Mediterranean and the AHEI scores were negatively  
43 associated with CRP levels (standardized regression scores: -0.043, -0.054, -0.043 and -0.067,  
44 respectively, all  $p < 0.01$ ). The "Fruits & vegetables" pattern was also negatively associated with  
45 leucocyte count (standardized regression score: -0.057,  $p < 0.01$ ). Conversely, no association between  
46 nutrients and inflammatory markers and between all dietary markers and IL-6 or TNF- $\alpha$  was found.

47 **Conclusion:** global dietary behaviours have a small but significant impact on inflammatory markers in  
48 the general population. The effect of individual nutrients or foods (fruits excepted) is of less clinical  
49 importance.

50 **Keywords:** dietary scores; inflammation; epidemiology; dietary patterns; nutrients

51

## 52 Introduction

53 Systemic low-grade inflammation has been linked to several chronic conditions such as  
54 cardiovascular disease (CVD)<sup>1</sup>. Moreover, chronically elevated rates of inflammatory markers such  
55 as C-reactive protein (CRP) and interleukin 6 (IL-6) could be predictors of CVD risk<sup>2</sup>.

56 The already well known association between diet and CVD may partly be linked through  
57 these inflammatory markers<sup>3</sup>. Indeed , several epidemiological studies have identified negative  
58 associations between inflammatory levels and specific nutrients such as polyphenols<sup>4 5</sup>,  
59 polyunsaturated fatty acids<sup>6</sup>, histidine<sup>7</sup> and branched-chain amino acids<sup>8</sup>.

60 Besides specific nutrients, dietary scores based on the consumption of specific food items  
61 have also been linked with inflammatory levels. For example, hypothesis-oriented scores such as the  
62 Mediterranean-diet score<sup>9</sup> has been linked with lower platelet and white blood cell counts<sup>10</sup>.  
63 Similarly, the Southern European Atlantic diet<sup>11</sup> and the Baltic Sea diet<sup>12</sup> have been associated with  
64 lower levels of CRP. Similarly, the Alternate Healthy Eating Index (AHEI) of McCullough and coll.<sup>13</sup>  
65 was negatively associated with IL-6 levels<sup>14 3</sup>. Conversely, the traditional Inuit diet, based on sea  
66 mammals' meat rich in n-3 fatty acids and vitamin D, tended to increase CRP and YKL-40 levels,  
67 possibly due to the presence of organic pollutants in marine mammals<sup>15</sup>.

68 Also, naïve-oriented scores, such as dietary patterns obtained through principal component  
69 analysis (PCA), have been shown to be associated with inflammatory levels. For example, "prudent"<sup>16</sup>  
70 or "health-aware"<sup>17</sup> patterns were negatively associated with CRP levels. Ozawa and coll. used  
71 reduced rank regression analysis and identified a "dietary inflammatory pattern" rich in red and  
72 processed meat, peas, legumes and fried food, and low in whole grains, which was positively  
73 associated with IL-6 levels and other inflammatory markers<sup>18</sup>. However, most studies assessed only  
74 a limited number of nutrients or foods, or even focused on a single dietary score or pattern, while  
75 studies comparing simultaneously the effect of different dietary parameters on inflammatory  
76 markers are scarce.

77           Hence, our study aimed at assessing the associations between a wide range of dietary  
78 parameters (macro- and micronutrients, single foods, dietary patterns and scores) and several  
79 inflammatory markers to find the most accurate indicator of an anti-or pro-inflammatory diet.

## 80 **Participants and methods**

### 81 *Sampling procedure*

82           The CoLaus study is a population-based study assessing the clinical, biological and genetic  
83 determinants of cardiovascular disease in the city of Lausanne, Switzerland<sup>19</sup>. Recruitment began in  
84 June 2003 and ended in May 2006, enrolling 6733 participants who underwent an interview, a physical  
85 exam, and a blood analysis. The first follow-up was performed between April 2009 and September  
86 2012, 5.5 years on average after baseline. We only consider data from the follow-up examination as  
87 dietary intake assessment was first introduced here.

### 88 *Blood samples*

89           Venous blood samples (50 mL) were drawn in the fasting state. Most biological assays were  
90 performed at the clinical laboratory of the Lausanne university hospital (CHUV) within 2 hours of blood  
91 collection on fresh samples. Supplementary samples were stored at -80°C.<sup>20</sup>

### 92 *Dietary intake*

93           Dietary intake was assessed using a validated self-administered, semi quantitative Food  
94 Frequency Questionnaire (FFQ) which also included portion size<sup>21</sup>. Briefly, this FFQ assesses the  
95 dietary intake of the previous 4 weeks and consists of 97 different food items accounting for more  
96 than 90% of the intake of calories, proteins, fat, carbohydrates, alcohol, cholesterol, vitamin D and  
97 retinol, and 85% of fibre, carotene and iron. For each item, consumption frequencies ranging from  
98 “less than once during the last 4 weeks” to “2 or more times per day” were provided, and the  
99 participants also indicated the average serving size (smaller, equal or bigger) compared to a  
100 reference size. Daily consumption of the different food items was computed based on frequency and  
101 portion size and expressed in mL (for drinks) or in grams (for other foods). Conversion into nutrients



102 was performed based on the French CIQUAL food composition table<sup>22</sup> taking into account portion  
103 size.

#### 104 *Dietary scores*

105 Three dietary scores were computed, two based on the Mediterranean diet, the third on a  
106 modification of the alternative healthy eating index (AHEI). The first Mediterranean dietary score  
107 (hereby designated as “Mediterranean score 1”) was derived from Trichopoulou et al.<sup>9</sup>, the score  
108 ranges between zero and eight. The second Mediterranean dietary score (hereby designated as  
109 “Mediterranean score 2”) is adapted to the Swiss population and was computed according to  
110 Vormund et al.<sup>23</sup>. Contrary to the score from Trichopoulou et al, dairy products are considered as  
111 beneficial. The score thus ranges between zero and nine. The AHEI was adapted from McCullough et  
112 al.<sup>13</sup>. In our study, the amount of trans fat could not be assessed, and we considered all participants  
113 taking multivitamins as taking them for a duration  $\geq 5$  years. Thus, the modified AHEI score ranged  
114 between 2.5 and 77.5 instead of 2.5 and 87.5 for the original AHEI score<sup>13</sup>. For all three scores, higher  
115 values represented a healthier diet.

#### 116 *Dietary patterns*

117 Prior to assessment of dietary patterns, the 97 items of the FFQ were grouped into 40 food  
118 and nutrient groups, including vitamin and food supplements (**supplementary table 1**) as previously  
119 reported<sup>24</sup>.

120 Dietary patterns were assessed by PCA with varimax rotation as done in previous studies<sup>25 26</sup>  
121 <sup>27 28</sup>. The Kaiser-Meyer-Olkin (KMO) and the Bartlett test for sphericity were applied to assess the  
122 appropriateness of applying PCA to the dataset. The Bartlett test compares the correlation matrix  
123 between the different items to be included in the PCA to the identity matrix. A non-significant Bartlett  
124 test indicates that the variables are highly correlated and that information compression using PCA is  
125 not useful. The KMO was 0.754, which was above the suggested minimum of 0.5<sup>29</sup> and comparable to

126 values reported in the literature<sup>25 30</sup>. The Bartlett test for sphericity yielded a p-value of <0.0001.  
127 Hence, both KMO and the Bartlett tests indicated that the data were suitable for PCA.

128 The number of dietary patterns to be retained was calculated based on the same criteria as in  
129 other studies<sup>31 26</sup>, namely: 1) analysis of the scree plot; 2) an eigenvalue higher than one and 3) the  
130 interpretability of the dietary pattern. For interpretation purposes, varimax rotation was performed.  
131 Items with absolute factor loading >0.30 were considered to characterize the dietary patterns<sup>24</sup>,  
132 although all items were used to calculate dietary pattern scores. Dietary patterns were categorized  
133 into quintiles and the dietary intake of the highest was reported.

#### 134 *Exclusion criteria*

135 Participants with the following characteristics were excluded: 1) No dietary data; 2) No socio-  
136 clinical data; 3) No inflammatory data; 4) Anti-inflammatory drugs; 5) Inflammation (CRP>20 mg/L); 6)  
137 Total energy intake (TEI) <850 or >4500 kcal/day.

#### 138 *Statistical analysis*

139 Statistical analysis was performed using Stata version 14.1 for Windows (Stata Corp, College  
140 Station, Texas, USA). Participants characteristics were expressed as number (percentage) for  
141 categorical variables or as average  $\pm$  standard deviation for continuous variables.

142 Bivariate associations were assessed using Spearman nonparametric rank correlation. Dietary  
143 markers significantly associated on bivariate analysis with inflammatory markers were further  
144 explored using multivariable analysis. Multivariable analysis was performed using linear regression  
145 adjusting for age (continuous), body mass index (continuous), gender, smoking (never, former,  
146 current), educational level (high, middle, low), physical activity, diabetes (yes/no) and total caloric  
147 intake (continuous). Results were expressed as standardized coefficients; standardized coefficients can  
148 be interpreted as multivariable-adjusted correlations. For multivariable analyses, inflammatory  
149 markers were log-transformed.

150 The importance of dietary scores and patterns relative to single foods was further addressed  
151 by entering simultaneously in each model one dietary pattern and the foods significantly associated  
152 with inflammatory markers. Statistical significance was considered for a two-sided test with  $p < 0.01$ .

## 153 **Results**

### 154 *Characteristics of the participants*

155 Of the initial 5064 participants, 1037 (20.5%) were excluded. The reasons for exclusion are  
156 indicated in **figure 1** and the characteristics of excluded and included participants are summarized in  
157 **supplementary table 2**. Excluded participants were older, had a higher BMI, a lower education, and  
158 were prone to smoke, be sedentary and have diabetes.

### 159 *Dietary patterns*

160 The results of the PCA are summarized in **supplementary table 3**. The KMO was 0.754,  
161 indicating that the data were suitable for PCA. Two unhealthy and one healthy dietary patterns were  
162 identified, explaining 20% of the overall variance. The three patterns were named “Meat & chips”  
163 (unhealthy: positive loadings for all kinds of meat and French fries); “Pastries & fat” (unhealthy:  
164 positive loadings for hard fats, pastries and sugar) and “Fruits & vegetables” (healthy: positive loadings  
165 for fruits & vegetables).

166 Participants in the highest quintile of the “Meat & chips” and the “Pastries & fat” patterns  
167 showed a higher intake of total fat (expressed as percentage of total energy intake) and of saturated  
168 fat (expressed as percentage of total fat). Participants in the highest quintile of the “Meat & chips”  
169 pattern also showed a higher intake of protein while participants in the highest quintile of “Fruits &  
170 vegetables” pattern showed a higher intake of carbohydrates and fiber (**supplementary table 3**).

171 *Associations of individual nutrients with inflammatory markers*

172 On bivariate analysis, total fat was positively associated with IL-6, and carotene was negatively  
173 associated with CRP and leucocyte count (**table 1**). However, these relationships were no longer  
174 significant when adjusting for confounders (**table 2**).

175 *Associations of single foods with inflammatory markers*

176 On bivariate analysis, fruits, carrots and tofu were negatively associated with CRP and  
177 leucocyte count; green salad, bananas, apples and kiwis were negatively associated with leucocyte  
178 count, while vegetables showed a positive association with IL-6 levels (**table 1**). After multivariable  
179 adjustment, only the negative association between tofu and CRP levels remained (**table 2**).

180 *Associations of dietary patterns and scores with inflammatory markers*

181 On bivariate analysis, the “Meat & Chips” pattern was positively associated with CRP levels  
182 and leucocyte count; the “Pastries & Fat” pattern was negatively associated with leucocyte count, and  
183 the “Fruits & Vegetables” pattern was negatively associated with CRP levels and leucocyte count (**table**  
184 **1**). After multivariable adjustment, only the negative associations between the “Fruits & Vegetables”  
185 pattern and CRP levels and leucocyte counts remained (**table 2**).

186 On bivariate analysis, both Mediterranean scores and the AHEI were negatively associated  
187 with CRP levels; the AHEI was also negatively associated with leucocyte count (**table 1**). After  
188 multivariable adjustment, only the negative associations between Trichopoulou’s Mediterranean  
189 score and the AHEI with CRP levels remained (**table 2**).

190 When entered simultaneously with fruit intake, the “Fruits & Vegetables” pattern, the  
191 Trichopoulou’s Mediterranean score and the AHEI tended to remain significantly and negatively  
192 associated with CRP levels, while the association with fruit intake was no longer significant  
193 (**supplementary table 4**).

194

## 195 **Discussion**

196           This study is one of the few that compared the associations between different dietary  
197 parameters and inflammatory markers in the general population. Our results show that (un)healthy  
198 dietary behaviours have a small but significant impact on inflammatory markers, while individual  
199 nutrients or foods have a lower clinical importance.

### 200 *Associations of individual nutrients with inflammatory markers*

201           Only a limited number of macro- and micronutrients were associated with inflammatory  
202 markers on bivariate analysis, and no significant association remained after adjustment for  
203 confounders. These findings are in agreement with Scottish study<sup>17</sup>, which also failed to find any  
204 significant association between several micronutrients (flavonoids and antioxidants) and inflammatory  
205 markers. Also, the lack of association between vitamin D intake and inflammatory markers matches an  
206 earlier study on Inuits, whose diet is known to be rich in this component<sup>32</sup>. Overall, our results suggest  
207 that most macro and micro-nutrients are not associated with inflammatory markers. Still, as no  
208 information was available regarding polyphenol intake, it was not possible to confirm or infirm the  
209 existing information on the anti-inflammatory properties of polyphenols<sup>4 5</sup>.

### 210 *Associations of single foods with inflammatory markers*

211           Fruit intake was negatively associated with CRP levels, but not with IL-6, TNF- $\alpha$  or leucocyte  
212 count. The association with CRP remained after multivariate analysis, a finding in agreement with the  
213 literature<sup>33</sup>. A possible explanation is the high polyphenol content of fruits<sup>34</sup>, which has been linked  
214 with a decrease in inflammation levels<sup>4 5</sup>. Our results thus stress the need of an adequate consumption  
215 of fruit to decrease inflammatory levels.

### 216 *Associations of dietary patterns and scores with inflammatory markers*

217           The Mediterranean diet score was negatively associated with CRP levels, and this association  
218 persisted after multivariate adjustment. These findings are in agreement with previous studies<sup>10 35 36</sup>,

219 suggesting that the beneficial effect of the Mediterranean diet on CVD might partly be linked to a  
220 decreased inflammatory status.

221 The AHEI was negatively associated with CRP levels. This finding is in agreement with some  
222 studies where the AHEI was inversely correlated with CRP<sup>37</sup> or IL-6 levels<sup>3,14</sup>, but not with other studies  
223 which failed to find any association<sup>38,39</sup>. As the associations were rather weak, a possible explanation  
224 is the relatively small sample size (<1000 participants) of the studies that failed to find a significant  
225 association.

226 Of the three dietary patterns obtained, only the “Fruits & Vegetables” one retained its negative  
227 association with CRP and IL-6 after multivariable adjustment. These findings are partly in agreement  
228 with other studies which also assessed dietary patterns using PCA. Indeed, both the “health-aware”  
229 pattern from the Lothian Birth Cohort study<sup>17</sup> and the “prudent” pattern from the Aberdeen  
230 Prospective Osteoporosis Screening Study cohort<sup>16</sup>, which scored high in fruits and vegetables, were  
231 negatively associated with CRP levels.

232 Interestingly, the “Fruits & Vegetables” pattern, the Trichopoulou’s Mediterranean score and  
233 the AHEI showed stronger negative associations with CRP levels than single fruit intake, indicating that  
234 their effects were not only due to a higher fruit intake, but that other components might also  
235 intervene. Overall, our results suggest that a diet rich in fruits (but not only) is paramount for reducing  
236 inflammatory levels.

### 237 *Importance for clinical practice*

238 Many studies have focused on the associations between single nutrients or foods and  
239 inflammatory markers. Still, increasing or decreasing the consumption of specific nutrients or even of  
240 selected foods might be difficult to achieve in general practice. Our results indicate that dietary  
241 recommendations focused on the consumption of several food groups are more important than  
242 recommendations focused on specific foods or nutrients<sup>40</sup>. Hence, in clinical practice, generic

243 recommendations could be provided, instead of focusing on specific foods or nutrients, which are  
244 difficult to identify and to integrate in a normal diet. This would facilitate dietary counselling by general  
245 practitioners, whose nutritional knowledge is usually low<sup>41</sup>.

246 Similarly, from a public health perspective, simple messages aimed at a healthier eating and  
247 increased consumption of fruits and vegetables<sup>42 43</sup> could be delivered. The impact of such measures  
248 in the general population could then be monitored by any of the scores (AHEI, Mediterranean or “Fruits  
249 and vegetables”) rather than by complex nutrient assessment.

#### 250 *Strengths and limitations*

251 Our study has several strengths: firstly, it is one of the very few simultaneously comparing the  
252 effect of different dietary parameters on inflammatory markers. Secondly, it relied on a rather large  
253 sample size, thus allowing a relatively adequate statistical power. Indeed, only five studies assessing  
254 the association between diet and inflammatory markers had a larger sample size<sup>4 11 44 10 18</sup>. Finally, due  
255 to the population-based setting, our results can be transposed to other populations and practical  
256 recommendations can be used in public health and clinical practice.

257 This study also has several limitations. First, and as already indicated, no information regarding  
258 polyphenols was available. Hence, it was not possible to confirm previous findings<sup>5 4</sup>. Future studies  
259 should rely either on an extensive food composition database or on the direct measurement of  
260 polyphenols in serum or urine. Second, the Mediterranean-diet score is based on a Greek population’s  
261 food consumption, and the scores obtained cannot be directly transposed to a Swiss population.  
262 Finally, we could not calculate the Dietary Inflammatory Index (DII), which is composed by a list of 45  
263 items, mainly specific nutrients and spices<sup>45</sup> and is often used in recent literature on CVD and  
264 inflammation, showing decreased CRP levels with high DII scores<sup>46</sup>. However, this score is based on  
265 food items<sup>45</sup> that are rarely collected in epidemiological studies and are dependent on the food  
266 composition table. Hence, its interest for public health or clinical practice is limited.

267 *Conclusion*

268 Our results show that healthy dietary behaviours have a small but significant impact on  
269 inflammatory markers in the general population. The effect of individual nutrients or foods appears to  
270 be of less clinical importance.

### 271 **Conflict of interest**

272 The authors report no conflict of interest.

### 273 **Ethical statement**

274 The institutional Ethics Committee of the University of Lausanne, which afterwards became  
275 the Ethics Commission of Canton Vaud ([www.cer-vd.ch](http://www.cer-vd.ch)) approved the baseline CoLaus study  
276 (reference 16/03, decisions of 13th January and 10th February 2003); the approval was renewed for  
277 the first (reference 33/09, decision of 23rd February 2009) and the second (reference 26/14, decision  
278 of 11th March 2014) follow-up. The full decisions of the CER-VD can be obtained from the authors  
279 upon request. The study was performed in agreement with the Helsinki declaration and its former  
280 amendments, and in accordance with the applicable Swiss legislation. All participants gave their signed  
281 informed consent before entering the study.

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288



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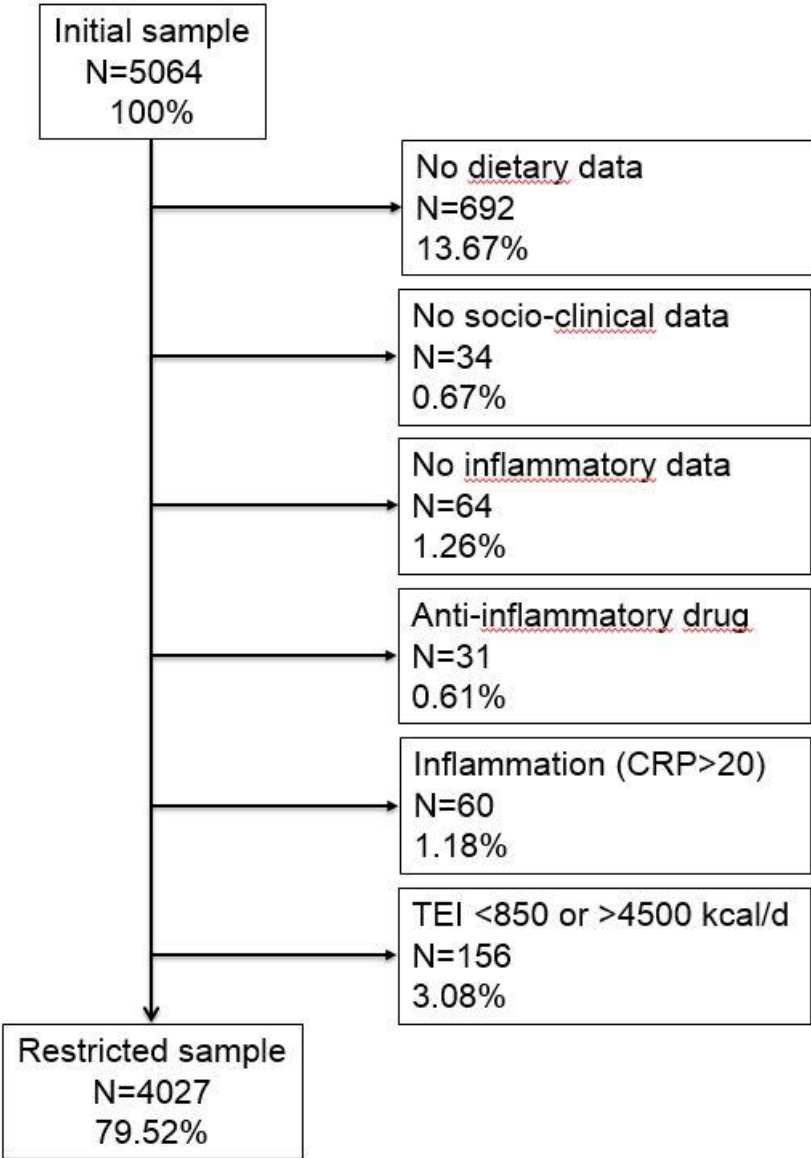
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422

# Figure legends

Figure 1 : selection procedure



## Tables

**Table 1:** bivariate associations between inflammatory and dietary markers CoLaus study, Lausanne, Switzerland, 2009-2012.

	CRP (log)	IL-6 (log)	TNF- $\alpha$ (log)	Leucocytes
Total energy intake	-0.006	<b>0.057</b>	0.016	0.012
Macronutrients				
Total protein	-0.003	0.046	0.017	0.027
Vegetable protein	-0.043	0.041	0.000	-0.030
Animal protein	0.019	0.034	0.022	0.050
Total carbohydrates	-0.034	0.037	0.012	-0.008
Monosaccharides	-0.038	0.018	0.004	-0.011
Polysaccharides	-0.020	0.038	0.012	-0.007
Total fat	0.001	<b>0.054</b>	0.016	0.019
Saturated	0.007	0.039	0.028	0.021
Monounsaturated	-0.007	<b>0.057</b>	0.010	0.017
Polyunsaturated	0.033	<b>0.065</b>	0.007	0.047
Fiber	-0.052	0.028	0.002	<b>-0.056</b>
Cholesterol	-0.010	0.025	-0.005	0.010
Alcohol	-0.022	0.030	0.015	0.011
Micronutrients				
Calcium	-0.014	0.022	0.020	-0.032
Iron	-0.024	0.051	0.005	-0.012
Retinol	0.015	0.013	0.005	0.015
Carotene	<b>-0.061</b>	0.031	-0.008	<b>-0.081</b>
Vitamin D	-0.025	0.032	0.002	-0.019
Vitamin A	-0.021	0.028	-0.009	-0.025
Food Items				

Fruits	<b>-0.055</b>	0.000	-0.021	<b>-0.061</b>
Vegetables	-0.045	<b>0.059</b>	-0.002	-0.038
Fish	-0.037	0.023	-0.001	-0.016
Haricots verts	-0.022	0.010	0.004	0.006
Chou-fleur	-0.029	0.036	0.000	-0.019
Tomatoes	0.017	0.005	0.006	0.015
Carrots	<b>-0.064</b>	0.024	-0.001	<b>-0.098</b>
Green salad	-0.040	0.014	-0.020	<b>-0.061</b>
Thick vegetables soup	-0.033	0.040	-0.005	0.007
Tomato sauce	-0.028	0.006	-0.032	0.019
Tofu	<b>-0.091</b>	0.013	-0.026	<b>-0.086</b>
Bananas, apples	-0.035	-0.004	0.007	<b>-0.059</b>
Oranges, tangerines	-0.023	-0.005	-0.027	-0.021
Peaches, nectarines	0.030	-0.036	0.044	0.027
Strawberries, blackcurrants	0.034	-0.041	-0.002	-0.011
Kiwis	-0.034	0.012	0.028	<b>-0.077</b>
Fresh fruit juice	-0.043	-0.031	-0.031	-0.002
Patterns				
Meat & Chips	<b>0.072</b>	0.030	0.029	<b>0.094</b>
Fruits & Vegetables	<b>-0.090</b>	0.030	-0.022	<b>-0.131</b>
Pastries & Fat	-0.048	-0.007	-0.017	<b>-0.095</b>
Dietary scores				
Mediterranean (Trichopoulou)	<b>-0.076</b>	0.046	-0.022	-0.032
Mediterranean (Vormund)	<b>-0.054</b>	0.027	-0.018	-0.038
AHEI <sup>1</sup>	<b>-0.111</b>	0.033	-0.016	<b>-0.072</b>

AHEI, alternate healthy eating index. Data from 4027 participants. Results are expressed as Spearman correlations. Significant ( $p < 0.01$ ) correlations are indicated in bold.



**Table 2:** multivariate associations between selected inflammatory and dietary markers CoLaus study, Lausanne, Switzerland, 2009-2012.

	CRP (log)	IL-6 (log)	Leucocytes
<b>Macronutrients</b>			
Total fat	-	0.001	-
Monounsaturated	-	0.006	-
Polyunsaturated	-	0.005	-
Fiber	-	-	-0.028
<b>Micronutrients</b>			
Carotene	-0.037	-	-0.047
<b>Food Items</b>			
Fruits	<b>-0.043</b>	-	-0.026
Vegetables	-	0.025	-
Carrots	-0.037	-	-0.041
Green salad	-	-	-0.043
Tofu	-0.024	-	-0.013
Bananas, apples	-	-	-0.010
Kiwis	-	-	-0.016
<b>Dietary patterns</b>			
Meat & Chips	0.011	-	0.045
Fruits & Vegetables	<b>-0.054</b>	-	<b>-0.057</b>
Pastries & Fat	-	-	-0.021
<b>Dietary scores</b>			
Mediterranean (Trichopoulou)	<b>-0.043</b>	-	-
Mediterranean (Vormund)	-0.039	-	-
AHEI <sup>1</sup>	<b>-0.067</b>	-	-0.041

AHEI, alternate healthy eating index; -, not assessed. Data from 4027 participants. Results are expressed as standardized regression coefficients adjusted for age (continuous), body mass index (continuous), gender, smoking (never, former, current), educational level (high, middle, low), physical activity, diabetes (yes/no) and total caloric intake (continuous). Significant ( $p < 0.01$ ) associations are indicated in bold.

## Supplementary files

**Supplementary table 1:** food groups and subgroups used to derive dietary patterns.

Main group	Subgroup	Examples
Dairy products	Plain	Milk, yogurt
	Low fat	Cottage cheese, 0% fat; yogurt, light
	Cheese	Emmental, Gruyère, fondue, feta, tomme, parmesan
Bread & cereals	White bread	
	Wholemeal bread	
	Breakfast cereals	Puffed rice, corn flakes, muesli, porridge
	Toasts, crackers	
Meat	Red meat	Beef, horsemeat, veal, spareribs, lamb chops
	White meat	Poultry
	Processed meat	Sausage, salami, cured ham, cervelas, pâté, terrine
	Liver	Liver (veal, pork, poultry)
Fish	Oily fish	Salmon (fresh or smoked)
	Canned or fried fish	
	Lean fish & seafood	Cod, pollock, trout, shrimp
Vegetables	Vegetables	Spinach, green beans, tomatoes, broccoli, carrots, vegetable soup
Potatoes	Boiled	
	French fries	
Sauces	Sauces	Mayonnaise, vinaigrette, tomato
Other	Cafeteria	Ravioli, tortellini, cannelloni, pizza, quiche
	Starch	Rice, pasta, semolina
Eggs	Eggs	
Tofu	Tofu	
Fruit	Fresh fruit	Banana, pear, apple, plum, grapes, berries, orange, tangerine, melon, apricot, peach, nectarine, kiwi
	Canned fruit	
	Fresh fruit juice	
	Bottled fruit juice	
	Low-fat products	
Fats & oils	Hard fats	Butter, margarine
	Olive oil	
	Other vegetable oils	Sunflower seed, peanut
Sugar & sweets	Bakery	Croissant, chocolate bread, cream cake, English cake, biscuits, cookies, fruit pie
	Chocolate	Chocolate, jam, honey, ice-cream
	Sweeteners	Aspartame
	Sugar	
	Supplements	Vitamins
	Other supplements	Fibre, bran, garlic pills

Drinks

Sodas

Tea & coffee

Water

Alcoholic

Beer, wine, champagne, spirits, aperitifs

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**Supplementary table 2:** Comparison between excluded and included participants, CoLaus study, Lausanne, Switzerland, 2009-2012.

	<b>Included</b>	<b>Excluded</b>	<b>P-value</b>
Sample size	4027	1037	
Gender (male)	46.51%	46.67%	0.926
Age (years)	57.2 ± 10.2	60.0 ± 11.3	<0.001
Education			<0.001
High	23.09%	14.44%	
Middle	26.89%	21.61%	
Low	50.01%	63.95%	
BMI (kg/m <sup>2</sup> )	26.0 ± 4.5	27.0 ± 5.1	<0.001
BMI categories			<0.001
Normal	45.10%	37.45%	
Overweight	38.99%	40.12%	
Obese	15.92%	22.43%	
Smoking status			<0.001
Current	20.34%	27.55%	
Former	37.94%	36.22%	
Never	41.72%	36.22%	
Sedentary	56.41%	65.79%	<0.001
Diabetes	9.56%	15.13%	<0.001

BMI, body mass index. Results are expressed as percentage or as average standard deviation. Between group comparisons using chi-square for categorical variables or student's t-test for continuous variables.

**Supplementary table 3:** factor loadings, percentage of total variance explained, and nutritional intake of the highest quintile for the three dietary patterns identified, CoLaus study, Lausanne, Switzerland, 2009-2012.

		<b>Meat &amp; chips</b>	<b>Fruits &amp; vegetables</b>	<b>Pastries &amp; fat</b>
Bread & cereals	White bread			0.307
Meat	Red meat	0.419		
	White meat	0.403		
	Processed meat	0.417		
	Liver, kidneys	0.421		
Vegetables	Vegetables		0.342	
Potatoes	French fries	0.334		
Fruit	Fresh fruit		0.346	
Fats & oils	Hard fats			0.373
Sugar & sweets	Bakery			0.346
	Chocolate			0.327
	Sugar			0.314
% of variance explained §		8%	7%	5%
Total energy intake (kcal)		2267 [1849 - 2755]	2160 [1788 - 2655]	2466 [2121 - 2948]
Macronutrients (% TEI)				
	Protein	16.3 [14.6 - 18.3]	14.9 [13.2 - 17.1]	13.9 [12.4 - 15.6]
	Carbohydrates	42.9 [37.4 - 48.3]	48.1 [42.7 - 54.2]	47.4 [41.4 - 53.2]
	Fat	35.6 [31.4 - 40.2]	34.1 [29.3 - 38.9]	35.0 [30.5 - 39.5]
	Alcohol	2.6 [0.6 - 6.7]	1.1 [0.2 - 2.9]	1.7 [0.4 - 4.6]
Fatty acids (% fat)				
	Saturated	36.6 [32.7 - 40.0]	33.8 [29.8 - 37.8]	39.5 [35.5 - 43.4]
	Monounsaturated	40.1 [37.5 - 42.7]	40.4 [37.4 - 44.2]	38.2 [35.7 - 41.1]
	Polyunsaturated	14.3 [12.3 - 17.0]	14.9 [12.7 - 17.5]	13.2 [10.9 - 16.0]
Fibre (g/day)		15.6 [11.4 - 21.3]	24.0 [17.6 - 32.2]	17.4 [13.1 - 23.4]

Only items with absolute factor loading >0.300 are shown. Results for energy and nutrient intake are expressed as median and [interquartile range]. TEI, total energy intake. §, by the rotated components using varimax procedure.

**Supplementary table 4:** multivariate associations between selected dietary patterns or scores with C-reactive protein levels, further adjusted for fruit intake, CoLaus study, Lausanne, Switzerland, 2009-2012.

	<b>Fruits &amp; Vegetables</b>	<b>Mediterranean diet</b>	<b>AHEI</b>
Fruit intake	-0.017 (0.399)	-0.032 (0.063)	-0.015 (0.399)
Selected score/pattern	-0.045 (0.021)	-0.035 (0.027)	-0.061 (<0.001)

AHEI, alternate healthy eating index. Data from 4027 participants. Results are expressed as standardized regression coefficients adjusted for age (continuous), body mass index (continuous), gender, smoking (never, former, current), educational level (high, middle, low), physical activity, diabetes (yes/no) and total caloric intake (continuous) and (p-value).