

Clustering of adherence to personalised dietary recommendations and changes in healthy eating index within the Food4Me study

Article

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**Title** 

Clustering of adherence to personalised dietary recommendations and changes in healthy eating index within the Food4Me Study

#### **Author names**

Katherine M. Livingstone\*<sup>1</sup>, Carlos Celis-Morales\*<sup>1</sup>, Jose Lara<sup>1</sup>, Clara Woolhead<sup>2</sup>, Clare B. O'Donovan<sup>2</sup>, Hannah Forster<sup>2</sup>, Cyril F.M. Marsaux<sup>3</sup>, Anna L. Macready<sup>4</sup>, Rosalind Fallaize<sup>4</sup>, Santiago Navas-Carretero<sup>5</sup>, Rodrigo San-Cristobal<sup>5</sup>, Silvia Kolossa<sup>6</sup>, Lydia Tsirigoti<sup>7</sup>, Christina P. Lambrinou<sup>7</sup>, George Moschonis<sup>7</sup>, Agnieszka Surwiłło<sup>8</sup>, Christian A. Drevon<sup>9</sup>, Yannis Manios<sup>7</sup>, Iwona Traczyk<sup>8</sup>, Eileen R. Gibney<sup>2</sup>, Lorraine Brennan<sup>2</sup>, Marianne C. Walsh<sup>2</sup>, Julie A. Lovegrove<sup>4</sup>, J. Alfredo Martinez<sup>5</sup>, Wim H. Saris<sup>3</sup>, Hannelore Daniel<sup>6</sup>, Mike Gibney<sup>2</sup>, John C. Mathers<sup>1</sup>, on behalf of the Food4Me Study.

#### **Author affiliations**

- 1, Human Nutrition Research Centre, Institute of Cellular Medicine, Newcastle University, Newcastle Upon Tyne, UK (KML, katherine.livingstone@newcastle.ac.uk, JL, jose.lara@newcastle.ac.uk, CCM, carlos.celis@newcastle.ac.uk; JCM, john.mathers@newcastle.ac.uk)
- 2, UCD Institute of Food and Health, University College Dublin, Belfield, Dublin 4, Republic of Ireland (CBD, clare.odonovan@ucdconnect.ie; HF, hannah.forster@ucdconnect.ie; CW, clara.woolhead@ucdconnect.ie; EG, eileen.gibney@ucd.ie; LB, lorraine.brennan@ucd.ie; MCW, marianne.walsh@ucd.ie; MG, mike.gibney@ucd.ie)
- 3, Department of Human Biology, NUTRIM School of Nutrition and Translational Research in Metabolism, Maastricht University Medical Centre, Maastricht, The Netherlands (CFMM, c.marsaux@maastrichtuniversity.nl; WHMS, w.saris@maastrichtuniversity.nl)
- 4, Hugh Sinclair Unit of Human Nutrition and Institute for Cardiovascular and Metabolic Research, University of Reading, Reading, UK (ALM, a.l.macready@reading.ac.uk; RF, r.fallaize@.reading.ac.uk; JAL, j.a.lovegrove@reading.ac.uk)
- 5, Center for Nutrition Research, University of Navarra, Pamplona, Spain; CIBER Fisiopatología Obesidad y Nutrición (CIBERobn), Instituto de Salud Carlos III, Madrid, Spain (SNC, snavas@unav.es; RSC, rsan.1@alumni.unav.es; JAM, jalfmtz@unav.es) 6, ZIEL Research Center of Nutrition and Food Sciences, Biochemistry Unit, Technical University of Munich, Germany (SK, silvia.kolossa@tum.de; HD, hannelore.daniel@tum.de)

- 7, Department of Nutrition and Dietetics, Harokopio University, Athens, Greece (LT, tsirigoti.lydia@gmail.com; CPL, cplambrinos@gmail.com; GM, gmoschi@hua.gr; YM, manios@hua.gr)
- 8, National Food & Nutrition Institute (IZZ), Poland (AS, asurwillo@izz.waw.pl; IT, itraczyk@izz.waw.pl)
- 9, Department of Nutrition, Institute of Basic Medical Sciences, Faculty of Medicine, University of Oslo, Oslo, Norway (CAD, c.a.drevon@medisin.uio.no)
- \*KML and CCM are joint first-authors

#### **Corresponding author; request for reprints**

Professor John C. Mathers

**Human Nutrition Research Centre** 

Institute of Cellular Medicine

Newcastle University

Biomedical Research Building

Campus for Ageing and Vitality

Newcastle upon Tyne

NE4 5PL, UK

john.mathers@newcastle.ac.uk

Tel: +44 (0) 1912081133 Fax: +44 (0) 1912081101

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**Abbreviations:** Body mass index (BMI); Cardiovascular disease (CVD); Food frequency questionnaire (FFQ); Healthy eating index (HEI); Physical activity level (PAL); Personalised Nutrition (PN); Randomized controlled trial (RCT); Sedentary behaviour (SB); Waist

circumference (WC)

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Ethical standards disclosure: This study was conducted according to the guidelines laid down in the Declaration of Helsinki and all procedures involving human subjects/patients were approved by the Research Ethics Committees at each University or Research Centre delivering the intervention. The Food4Me trial was registered as a RCT (NCT01530139) at Clinicaltrials.gov. All participants expressing an interest in the study were asked to sign online consent forms at two stages in the screening process. These consent forms were automatically directed to the local study investigators to be counter-signed and archived.

#### 1 Abstract (words count=250)

#### 2 Objective

- 3 To characterize clusters of individuals based on adherence to dietary recommendations and to
- 4 determine whether changes in Healthy Eating Index (HEI) scores in response to a
- 5 personalised nutrition (PN) intervention varied between clusters.

#### 6 Design

- 7 Food4Me study participants were clustered according to whether their baseline dietary
- 8 intakes met European dietary recommendations. Changes in HEI scores between baseline and
- 9 month 6 were compared between clusters and stratified by whether individuals received
- 10 generalized or PN advice.

#### 11 Setting

12 Pan-European, internet-based, 6-month randomized controlled trial.

#### 13 Subjects

14 Adults aged 18-79 years (*n* 1480).

#### 15 Results

- 16 Individuals in cluster 1 (C1) met all recommended intakes except for red meat, those in
- cluster 2 (C2) met two recommendations and those in cluster 3 (C3) and cluster 4 (C4) met
- one recommendation each. C1 had higher intakes of white fish, beans and lentils and low fat
- dairy products and lower percentage energy intakes from saturated fatty acids (P<0.05). C2
- 20 consumed less chips and pizza and fried foods than C3 and C4 (P<0.05). C1 were lighter, had
- 21 lower BMI and WC than C3 and were more physical active than C4 (P<0.05). More
- individuals in C4 were smokers and wanted to lose weight than C1 (P<0.05). Individuals who
- 23 received PN advice in C4 reported greater improvements in HEI compared with C3 and C1
- 24 (*P*<0.05).

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#### Conclusions

- 26 The cluster where the fewest recommendations were met (C4), reported greater
- 27 improvements in HEI following a 6-month trial of PN whereas there was no difference
- between clusters for those randomised to the Control, non-personalised dietary intervention.

- **Trial registration** Clinicaltrials.gov NCT01530139
- **Key Words** Clustering; personalised nutrition; dietary recommendations; healthy eating
- 31 index

# INTRODUCTION

33	Global obesity prevalence has reached epidemic proportions with 37% of men and 38% of
34	women now either overweight or obese (1). Poor dietary choices and inadequate physical
35	activity are the primary causes of obesity (2). Current strategies for improving diet and other
36	lifestyle behaviours, such as consuming 5 portions of fruit and vegetables per day (3), are
37	based on "one size fits all" generalised dietary guidelines. Given that the burden of obesity is
38	increasing (1), alternative strategies for improving dietary behaviours are being developed,
39	including predictive, personalised, preventative and participatory interventions (4). Recent
40	evidence suggests that genetic-based personalised nutrition (PN) improves dietary intakes
41	more than non-personalised advice <sup>(5)</sup> . However, since dietary intakes tend to cluster <sup>(6; 7)</sup> , it
42	may be possible to enhance the efficacy of interventions by further characterization of
43	participants according to their dietary and lifestyle behaviours and, subsequently, use this
44	information to strengthen the basis for personalization of the intervention. For example, lower
45	intakes of fruit, vegetables and wholegrains are often associated with higher intakes of red or
46	processed meat (8). In addition, less healthy dietary clusters are associated with increased
47	disease risk (9), and unhealthy dietary and lifestyle behaviours is associated with higher levels
48	of sedentary behaviour (7) and mortality (10; 11). Clustering individuals based on whether they
49	meet dietary recommendations may be a useful predictive tool for estimating response to an
50	intervention (12; 13; 14) and may help to stratify or personalise interventions.
51	The Food4Me proof-of-principle (PoP) study was the first internet-based study to
52	demonstrate that PN advice was more effective in improving dietary intakes, including
53	lowering intakes of red meat when compared with conventional "one size fits all" population-
54	based advice. However, the characteristics of individuals clustered on the basis of adherence
55	to current recommended dietary intake of fruit and vegetables, wholegrains, oily fish, dairy
56	products and red and processed meat, are unknown. Thus, the aims of this analysis were to i)
57	characterise European adults participating in the Food4Me study (15) according to clustering
58	based on European recommendations for healthy eating and ii) determine whether cluster
59	membership predicted dietary changes following a PN intervention.

# **METHODS**

# Study design and population

63	The Food4Me study was a 6-month, 4-arm, internet-based, RCT in 1607 individuals						
64	conducted across 7 European countries (15). Participants were recruited via the Food4Me						
65	website (16) to emulate a web-based PN service. This was aided by local and national						
66	advertising of the study via the Internet, radio, newspapers, posters, e-flyers, social media and						
67	word of mouth. Recruitment took place between August 2012 and August 2013 in the						
68	following sites: University College Dublin (Ireland), Maastricht University (The						
69	Netherlands), University of Navarra (Spain), Harokopio University (Greece), University of						
70	Reading (United Kingdom, UK), National Food and Nutrition Institute (Poland), Technical						
71	University of Munich (Germany). The Research Ethics Committees at each University or						
72	Research Centre delivering the intervention granted ethical approval for the study. The						
73	Food4Me trial was registered as a RCT (NCT01530139) at Clinicaltrials.gov. All participants						
74	expressing an interest in the study were asked to sign online consent forms at two stages in						
75	the screening process.						
76							
77	Intervention arms						
78	Participants were randomized to receive non-personalised, generalised dietary advice						
79	(Control), or one of three levels of PN (Level 1, Level 2 or Level 3). Briefly, non-						

Participants were randomized to receive non-personalised, generalised dietary advice (Control), or one of three levels of PN (Level 1, Level 2 or Level 3). Briefly, non-personalised dietary advice was based on national dietary recommendations in each of the 7 European countries. These "standardised" recommendations included advice on energy intake and on the consumption of fruits and vegetables, wholegrains, fish, dairy products, meat, type of fat and salt. Participants randomised to Level 1 received personalised dietary advice on how their intakes of these food groups compared with guideline amounts. Participants randomised to Level 2 received advice based on their dietary intake (as for Level 1) and also on their baseline phenotypic data. The phenotypic feedback was based on anthropometric measurements and nutrient- and metabolic-related biomarkers. Participants randomised to Level 3 received advice based on their dietary intake, phenotypic and genotypic data collected at baseline. The genotypic feedback was based on specific variants in five nutrient-responsive genes selected specifically for the study. Further details are provided elsewhere (15).

#### Screening questionnaires and dietary intakes

Participants eligible for inclusion in the RCT completed an online questionnaire to collect detailed information on socio-demographic, health and anthropometric characteristics and dietary habits. Following completion of this questionnaire, participants were asked to complete an online food frequency questionnaire (FFQ) to estimate usual dietary intake. This FFQ, which was developed and validated for this study (17; 18), included 157 food items consumed frequently in each of the 7 recruitment countries. Intakes of foods and nutrients were computed in real time using a food composition database based on McCance & Widdowson's "The composition of foods" (19). Intakes of nutrients were assessed based on standardised recommendations (Supplementary Table 1) for dietary intakes of foods and food groups (20), which were integrated and harmonised across 8 European countries (UK, Ireland, Germany, The Netherlands, Spain, Greece, Poland and Norway) (21; 22; 23; 24). The following 4 food group recommendations were used in the present analysis: eat at least 5 portions of fruit and vegetables every day (operationalised as >400g); eat at least 3 portions of wholegrain products daily (>50g); eat at least 1 portion of oily fish per week (>150g) and eat less than 3 portions of red or processed meat per week (<450g) (20). The Healthy Eating Index 2010 (HEI) was derived based on intakes of the following components: ratio of monoand polyunsaturated fatty acids to saturated fatty acids, protein, salt, "empty calories", refined grains, seafood and plant protein, fruit, whole fruit, vegetables, greens and beans, wholegrains, dairy products (25).

#### Personalised feedback report

Participants randomized to PN received personalised reports via email at baseline, month 3 and month 6 of the intervention based on diet, anthropometric measurements and physical activity. Using information on the individual's intakes of nutrients, algorithms were used to rank information on need for dietary change and to provide participants with 3 specific dietary, food-based goals. For participants randomized to Level 2 and Level 3, the dietary advice was also based on phenotypic data (Level 2) and phenotypic plus genotypic data (Level 3). Reported intakes were compared with recommended intakes and determined to be adequate, high or low. If intakes were too high or too low, contributing foods were identified and specific messages developed to advise change in intake of those foods. Dietary intakes relative to recommendations were illustrated using a three-colour sliding scale: green representing "Good, no change recommended," amber representing "Improvement recommended". For the

genotype-based information, risk was indicated using "Yes" or "No" according to whether the participant did, or did not, carry the higher risk variant for each of the 5 nutrient-related genes included in the study. Additionally, each report contained a personalized message from the dietitian/ nutritionist to the participant. Further details of the protocol are provided elsewhere <sup>(15)</sup>.

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#### Anthropometric, socio-demographic and physical activity measures

Detailed standardised online instructions were given for participants to self-measure and selfreport their body weight, height and waist circumference (WC) via the Food4Me website (www.Food4me.org). Body mass index (BMI) was estimated from body weight and height. Self-reported measurements were validated in a sub-sample of the participants (n=140) and showed a high degree of reliability (26). Physical activity levels (PALs) and time spent in sedentary behaviours (SB) were estimated from triaxial accelerometers (TracmorD, Philips Consumer Lifestyle, the Netherlands). Participants self-reported smoking habits and occupation. Occupations were grouped according to the European classifications of occupations and their salaries (the European wide average salary for each occupation was compared to the mean overall salary. If the standard deviation of the salary was >0.5 they were placed in group 1, between 0.5 to -0.5 were placed into group 2 and <-0.5 were placed into group 3): Group 1: Professional and managerial (professionals; managers); Group 2: Intermediate (Armed forces occupations; technicians and associate professionals; clerical support workers); Group 3: Routine and manual (craft and related trades workers; plant and machine operators and assemblers; service and sales workers; elementary occupations; skilled agricultural, forestry and fishery workers) (27; 28). Categories for "Students" and "Retired and unemployed" were added.

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#### Statistical analysis

Data were analysed using Stata (version 13; StataCorp, College Station, TX, USA) and IBM SPSS (V.22, IBM Corporation, Armonk, NY, USA). Clusters of dietary recommendations were generated based on whether participants met the following 4 food group recommendations at baseline and were coded as 0 or 1 accordingly: eat at least 5 portions of fruit and vegetables every day (operationalised as >400g); eat at least 3 portions of wholegrain products daily (>50g); eat at least 1 portion of oily fish per week (>150g) and eat

159	less than 3 portions of red or processed meat per week (<450g). Clusters were derived using
160	the SPSS Two Step cluster analysis procedure (29). Small pre-clusters were generated based
161	on log-likelihood distance criterion (Step 1), and were merged into distinct groups using
162	agglomerative hierarchical clustering (Step 2). Automatic selection and the Bayesian
163	Information Criterion (BIC) were used to determine the optimal number of clusters.
164	Robustness and stability of the final clusters were re-evaluated by random ordering of cases
165	(four times). This clustering methodology identified the percentage of participants within
166	each cluster who met recommended intakes of each of the 4 food groups of public health
167	importance. Logistic regression was used to test for significant differences across categorical
168	variables and ANOVA was used for continuous variables. Tukey's pairwise comparisons
169	were used to test for significant differences between clusters. Analyses were adjusted for age
170	sex, country, BMI, PAL and smoking, except when those (or related) variables were being
171	assessed i.e. analyses were not adjusted for BMI when assessing BMI, body weight or WC.
172	Results were deemed significant at $P$ <0.05. To exclude extreme intakes of the food groups
173	used for clustering, the top and bottom 3SD of these intakes were excluded prior to
174	clustering.

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#### RESULTS

Of the 5562 individuals who registered on the Food4Me website, 1607 were randomised into the study and a total of 1480 provided baseline data on dietary intakes <sup>(15)</sup>.

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#### Dietary adequacies across Food4Me cohort

- Recommended intakes for nutrients are summarised in Supplementary Table 1. On average, 181 50% of individuals met the recommendations for total fat (Supplementary Table 1). The 182 percentage of individuals who met the recommendations for saturated (SFA), mono- (MUFA) 183 and polyunsaturated fatty acids (PUFA) intake was 54, 24 and 36%, respectively 184 (Supplementary Table 1). Only 56% of individuals met the recommendation for carbohydrate 185 intake, whereas 91% of individuals had adequate protein intakes. Only 7 and 46% of 186 individuals met the recommendations for salt and dietary fibre intakes, respectively. Meeting 187 recommended micronutrient intakes ranged from 61% (folate) to 99% (vitamin B12; 188
- Supplementary Table 1).

As summarised in **Supplementary Table 2**, approximately half (52%) of participants reported consuming at least 5 portions of fruit and vegetables per day and 32% consumed at least 1 portion of oily fish per week. Nearly three quarters (74%) of participants consumed more than 3 servings of wholegrains per day and approximately half of participants (51%) consumed less than 3 servings of red meat per week (>450g/week). 14% of individuals met the recommendation for dairy product intake (>600g/day).

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#### **Cluster characterization**

Clustering of individuals according to whether they met the recommendations for dairy products, fruit and vegetable, oily fish, red meat and wholegrain intake at baseline did not create clear clustering due to the low percentage of individuals who met the recommendation for dairy products (2 clusters). Exclusion of dairy products as a clustering variable provided improved clustering, as estimated by silhouette measure of cohesion and separation (average silhouette: 0.3 vs 0.5; 4 clusters, **Supplementary Table 3**). Cluster one (C1) was the largest (n=475) and was particularly characterised by individuals meeting the recommended intake for oily fish (100% of individuals); 74 and 69% of C1 members met the recommendations for wholegrains and fruit and vegetables, respectively, whereas only 46% met the recommendation for red meat. Cluster 2 (C2; n=398) was the second largest and was particularly characterised by all members meeting recommendations for wholegrains (100%) and red meat (100%), only 50% met the recommendation for fruit and vegetables and no one meeting the recommendation for oily fish. All individuals in cluster 3 (C3; n=348) met the recommendation for wholegrains, but no one met the recommendation for oily fish, or red meat, whereas only 48% met the recommended intake for fruit and vegetables. None of the participants in Cluster 4 (C4; n=259) met the recommended intakes for either oily fish or wholegrains; only 50 and 71% of C4 members achieved the recommended intakes for red meat and fruit and vegetables, respectively (Supplementary Table 3).

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#### **Dietary intakes by clusters**

- Intakes of oily fish and fruit and vegetables were higher in C1 than in C2, C3 and C4
- 219 (P<0.05), and wholegrain intakes were higher in C1, C2 and C3 than in C4 (**Table 1**;
- 220 P<0.05). Red meat intake was lower in C1, C2 and C3 than in C4 (P<0.05). Intakes of fruit

221	juice, eggs, chicken, white fish, fish products, beans and lentils and low fat dairy products
222	were higher in C1 than C4, whereas intakes of non-wholegrain products were lower
223	(P<0.05). Participants in C2 consumed lower intakes of chips and pizza and fried foods than
224	C3 and C4 (P<0.05; Table 1). Total energy intake and energy intake to basal metabolic rate
225	ratio (EI: BMR) were higher in C1 than in C2 and C4 and higher in C3 than in C2 (P<0.05;
226	Table 1). Individuals in C1 derived higher percentages of energy intake from PUFA and
227	protein than those in C2 and C4 (P<0.05) and individuals in C2 higher percentage energy
228	from carbohydrates than participants in C3 and C4 (P<0.05). In contrast, individuals in C1
229	had lower percentage energy intakes from total fat and SFA than those in C4 (P<0.05) and
230	higher percentage energy intake from monounsaturated fatty acids (MUFA) than participants
231	in C2 and C3 (P<0.05). Subjects in C1 had lower percentage energy intake from sugar than
232	C2 (P<0.05). Participants in C1 consumed more dietary fibre and salt than those in C2 and
233	C4 ( <i>P</i> <0.05).
234	More individuals in C1 met the recommendation for total fat intake (51%), SFA (62%),
235	PUFA (42%) and dietary fibre (56%) than C4 cluster members (Supplementary Table 4).
236	Fewer individuals in C1 met the recommendations for protein intake (86%) than those in C2
237	(97%) and C3 (93%). Furthermore, fewer individuals in C1 met the recommendation for salt
238	intake (5%) than C2 (11%) and C4 (17%; Table 4).
239	
240	Socio-demographic, anthropometric and health characteristic by clusters
241	Individuals in C1 were on average 4.5 years older than C4 (P<0.05; <b>Table 2</b> ). Body weight
242	was significantly lower in C1 than in C3, and lower in C2 compared with C3 and C4
243	(P<0.05). Individuals in C1 had 1.4kg/m <sup>2</sup> lower BMI and 5cm lower WC than participants in
244	C3 (P<0.05) and PAL was higher in C1 than C2 and C4 (P<0.05). 11% more individuals in
245	C4 wanted to lose weight than those in C1 (P<0.05; Table 2) and C4 was characterised by
246	more current smokers than C1 (P<0.05). 12% more individuals in C1 had a professional or
247	managerial occupation than C4, and similarly 7% more individuals had a manual occupation
248	in C4 compared with C1 (P<0.05; Table 2). No other significant differences were observed
249	(Table 2).
250	

Changes in Healthy Eating Index (HEI) by cluster after 6 months intervention

252	Baseline and follow up HEI scores and their components are presented in <b>Table 3</b> . There
253	were no significant differences in changes in HEI between clusters for those randomised to
254	non-personalised dietary advice. In contrast, for individuals who received PN advice (based
255	on information of current diet alone or combined with information on phenotype and
256	genotype), changes in HEI differed between clusters (P<0.001). There were bigger
257	improvements in HEI for participants in C4 compared with C1 and C2 (P<0.05) and in C2
258	compared with C4 (P<0.05; Figure 1). There were no significant differences in changes in
259	HEI between clusters when PN was stratified by L1, L2 or L3 (data not shown).
260	
261	Sensitivity analyses
262	Exclusion of participants with reported intakes more than 3 SD above or below the mean
263	dietary intakes of wholegrain, oily fish, red meat and fruit and vegetables revealed similar
264	clusters (Supplementary Table 5). The pattern of the main results remained the same, with
265	individuals in C3 and C4 making greater changes in HEI at month 6 than those in C1, and
266	participants in C4 compared with those in C2 ( <i>P</i> <0.05).
267	
268	DISCUSSION
268 269	DISCUSSION Main findings
269	Main findings
269 270	Main findings  Based on our secondary analysis in the Food4Me PoP study, we identified four distinct
<ul><li>269</li><li>270</li><li>271</li></ul>	Main findings  Based on our secondary analysis in the Food4Me PoP study, we identified four distinct clusters of individuals according to their adherence to current European dietary
<ul><li>269</li><li>270</li><li>271</li><li>272</li></ul>	Main findings  Based on our secondary analysis in the Food4Me PoP study, we identified four distinct clusters of individuals according to their adherence to current European dietary recommendations. Individuals in C1 and C2 met more dietary recommendations than those in
<ul><li>269</li><li>270</li><li>271</li><li>272</li><li>273</li></ul>	Main findings  Based on our secondary analysis in the Food4Me PoP study, we identified four distinct clusters of individuals according to their adherence to current European dietary recommendations. Individuals in C1 and C2 met more dietary recommendations than those in C3 and C4. Moreover, on average individuals in C1 and C2 had a healthier diet, lower BMI
<ul><li>269</li><li>270</li><li>271</li><li>272</li><li>273</li><li>274</li></ul>	Main findings  Based on our secondary analysis in the Food4Me PoP study, we identified four distinct clusters of individuals according to their adherence to current European dietary recommendations. Individuals in C1 and C2 met more dietary recommendations than those in C3 and C4. Moreover, on average individuals in C1 and C2 had a healthier diet, lower BMI and WC and smoked less compared with those in C3 and C4. When randomised to a 6-month
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dietary guidelines at baseline for participants in the Food4Me intervention study. This 284 approach identified groups of individuals who differed in the number, and groupings, of 285 dietary recommendations they met. Clusters where more individuals met the 286 recommendations were characterised by being slightly older and in more highly educated 287 occupations, which is a well-established characteristic of healthy dietary clusters (31). 288 Clustering of dietary intakes and adequacies have been investigated in relation to several 289 health outcomes (7; 8; 32) and can be strong predictors of these outcomes (33). A recent review of 290 dietary clusters and health outcomes by the USDA (34) concluded that the strongest evidence 291 for an association between unhealthy dietary patterns and increased disease risk, is for 292 cardiovascular disease (CVD), followed by obesity and then type 2 diabetes. This USDA 293 review concluded that there was a lack of studies assessing dietary intakes at follow-up and 294 295 using a universal and quantitative indicator of dietary intake. Our study is in line with these recommendations as we utilised the HEI, which is a validated estimate of dietary adequacy, 296 297 and we assessed dietary change using the same instrument at both baseline and follow-up. Although more limited, some prospective and RCT studies have investigated the effect of 298 clustering on changes in health outcomes (12; 35; 36), and some studies have used adherence to 299 dietary recommendations to derive clusters (12; 13; 14; 37; 38). Dietary recommendations used in 300 studies included in the systematic review by the USDA (34) varied according to the study, but 301 all included a measure of fruit and vegetable, wholegrains and meat intake. 302 To our knowledge, no previous research has evaluated the impact of clustering of dietary 303 recommendations on the response to a PN intervention. We observed that individuals in the 304 cluster where the fewest recommendations were met (C4) reported the biggest improvement 305 in HEI following PN intervention but there were no differences between clusters in response 306 to conventional, non-personalised dietary advice. Given that adverse lifestyle behaviours and 307 the prevalence and risk of death from obesity-related diseases are strongly socioeconomically 308 patterned <sup>(39)</sup>, it is important that appropriate interventions are targeted to those most in need 309 of improved lifestyle. Whilst research on the development and implementation of PN 310 interventions and their effects on changing diets is in its infancy (40), the findings from the 311 present study provide encouragement that PN interventions can be more effective than 312 current "one size fits all" interventions and that they may be particularly effective amongst 313 individuals with the poorest diets. There have been concerns that PN may be taken up only by 314 the 'worried well' (41), who already have adequate dietary intakes. However, our findings 315 316 suggest that PN is most effective in people who have the least adequate diets, and therefore

the greatest need for improvement in dietary intakes with the potential for significant reductions in disease risk.

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#### **Strengths and limitations**

The present study had a number of strengths. Our findings are derived from a relatively large number of participants who were broadly representative of European adults from 7 different European countries. The Food4Me RCT collected extensive information on anthropometrics, physical activity and socio-demographic and health-related data, which contributed to detailed characterization of participants in the clusters. Our study design allowed us to estimate changes in dietary intakes using the same validated instrument at baseline and at month 6. Furthermore, we quantified responses using the HEI, which has been shown to be an effective indicator of overall diet quality (25) and, therefore, a better measure of overall dietary change than outcomes based on single foods or nutrients. A limitation of the study is that our data were self-reported via the internet, which may have introduced measurement error. However, the validity of internet-based, self-reported anthropometric data is high <sup>(42)</sup> and has been confirmed in the present study <sup>(26)</sup>. We were not able to include dairy products as a dietary recommendation in the present analyses due to so few individuals meeting the recommendation. However, dairy products do not have a recommended intake in the UK and so habitual diets would not necessarily be expected to comply with this recommendation, even if they were very health conscious. Dietary intakes were estimated by a FFQ, which is known to be subject to misreporting error (43) but this was minimised by validating our FFQ against a 4-day weighed food record (18). Moreover, our estimation of dietary change was based on the HEI, which is a validated indicator of overall diet (25), and which may be less susceptible to reporting errors than approaches measuring change in specific nutrients or individual foods. Our study participants were almost solely Caucasian – thus, further research in wider ethnicity groups is required to generalise our findings to other populations. One of the primary aims of the Food4Me PoP study was to evaluate change in intakes of food groups across 4 treatment arms. Thus, although the present study is a secondary analysis of these data, clustering was based on how individuals adhered to food group recommendations and included 4 clusters. As a result, our analyses are likely to be powered to detect differences between clusters.

349	Implications of findings
350	Our findings suggest that the efficacy of PN in modifying dietary intakes depends on the
351	clustering of adherence to dietary recommendations, with those with the poorest diets
352	benefiting most from the PN intervention. As a result, the implementation of PN-based
353	interventions in individuals with the least healthy diets may help to address health
354	inequalities. Understanding the characteristics of individuals within coherent clusters which
355	are linked with their responsiveness to interventions may help in the design and
356	implementation of more effective health promotion actions. Future PN interventions may
357	benefit from tailoring PN advice based on clustering of overall dietary behaviours rather than
358	on single nutrients or foods.
359	
360	Conclusions
361	We identified four distinct clusters of individuals based on adherence to current food-based
362	dietary recommendations. The cluster where the fewest recommendations were met (C4)
363	reported significantly greater improvements in their diets (as estimated by the HEI) following
364	a 6-month trial of PN, whereas there was no difference between clusters for those randomized

to the Control, non-personalised dietary intervention.

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#### FIGURE LEGENDS

**Figure 1** Changes from baseline to month 6 in Healthy Eating Index by clusters of adherence to recommendations at baseline

Values represent predicted means and SE. Models were adjusted for age, sex, body mass index, physical activity level, smoking habits and country and Posthoc Tukey's tests was used to test for significant differences between clusters (C); C4>C1 (P<0.001), C3>C1 (P=0.005)

Table 1 Food and nutrient and intakes by participants by clusters of adherence to recommendations at baseline

				Clu	sters				P*
	1 (r	n=475)	2 (n=	2 (n=398) 3 (n=348)			4 (n=	259)	_
	Mean	SD	Mean	SD	Mean	SD	Mean	SD	-
Dietary recommendations, g/d									
Oily fish	48	$32^{2,3,4}$	8	7	10	7	8	7	<0.001
Wholegrains	183	$182^{2,4}$	216	$184^{3,4}$	205	$165^{4}$	22	16	<0.001
Red meat	85	80.9 <sup>2,3,4</sup>	30	$20^{3,4}$	119	$53^{4}$	84	96	<0.001
Fruit and vegetables	610	371 <sup>2,3,4</sup>	470	3033,4	456	288	339	218	<0.001
Other food intakes, g/d									
Fruit Juice	117	1813,4	114	165	94	144	76	108	0.008
Non-wholemeal	116	$140^{2,4}$	78	$76^{4}$	114	$103^{4}$	149	189	<0.001
Eggs	41	$41^{2,3}$	22	24	31	47	30	51	<0.001
Chicken, grilled or roast	36	37 <sup>2,3,4</sup>	17	$21^{3}$	28	25	25	27	<0.001
White fish	26	$26^{2,3,4}$	10	14	13	14	11	14	<0.001
Fish products	19	$30^{2,4}$	10	$11^{3}$	14	16	13	15	<0.001
Beans and lentils	30	$40^{2,3}$	15	24	16	27	22	28	<0.001
Butter	4	93	6	$11^{3}$	9	$18^{4}$	5	12	0.005
Low fat dairy	293	$296^{2,3,4}$	217	203	221	212	173	219	<0.001
High fat dairy	64	120	60	119	83	113	83	204	0.44
Sugar sweetened beverages	36	176	18	55	40	139	41	84	0.39
Low calorie soft drinks	66	194	46	154	80	239	72	190	0.53
Added sugar	4	9	4	11	5	13	7	13	0.11
Chocolate and sweets	21	37	19	23	26	61	17	26	0.10
Cakes	22	31	18	25	20	25	22	39	0.08
Biscuits	30	55	21	37	35	88	27	55	0.38
Ice-cream	7	19	6	11	7	12	7	13	0.62
Pastries	8	34	4	6	6	10	10	39	0.49
Crisps	4	10	3	$5^3$	5	10	4	8	0.06
Chips and pizza	30	41	24	$22^{3,4}$	35	30	34	35	0.001
Fried foods	33	$52^{2}$	21	$28^{3,4}$	34	35	33	30	0.047
Nutrient intake									
Total energy, kcal/d	2870	1219 <sup>2,4</sup>	2218	$745^{3}$	2855	$1065^{4}$	2106	978	<0.001
EI:BMR ratio	1.9	$0.7^{2,4}$	1.5	$0.5^{3}$	1.8	$0.6^{4}$	1.4	0.6	< 0.001
Total fat, % energy	36.0	$5.7^{2,4}$	34.1	$5.6^{3,4}$	36.4	5.5	37.9	6.6	< 0.001
SFA, % energy	13.4	$2.8^{3,4}$	13.6	$3.3^{3,4}$	14.9	3.0	15.3	3.3	< 0.001
MUFA, % energy	14.2	$3.2^{2,3}$	12.6	$2.8^{3,4}$	13.6	$2.6^{4}$	14.8	3.5	< 0.001
PUFA, % energy	6.0	$1.4^{2,4}$	5.7	1.4	5.6	1.3	5.5	1.7	0.003
Protein, % energy	18.3	$4.1^{2,3,4}$	15.5	$3.2^{3,4}$	17.0	2.9	17.3	3.7	< 0.001

Carbohydrate, % energy	44.5	$7.5^{2,3}$	49.6	$7.0^{3,4}$	45.6	6.4	43.7	8.3	< 0.001
Sugars, % energy	21.0	$5.9^{2}$	22.5	$6.1^{3,4}$	19.8	5.6	20.8	5.9	< 0.001
Dietary fibre, g/d†	34.0	$15.8^{2,4}$	30.2	$14.4^{4}$	31.7	$12.8^{4}$	18.7	8.2	< 0.001
Salt, g/d†	8.3	$4.0^{2,4}$	6.1	$2.7^{3}$	8.7	$3.6^{4}$	5.9	3.6	< 0.001

Values represent means and SD

<sup>\*,</sup> ANOVA were adjusted for age, sex, BMI, PAL, smoking habits and country; Posthoc Tukey tests were performed to test for significant differences between clusters Superscript numbers denote where the differences lie across the clusters. For example, 1 means significantly different from cluster 1.

<sup>†,</sup> P values are also adjusted for total energy intake.

**Table 2** Socio-demographic characteristics of participants by clusters of adherence to recommendations at baseline

				Clu	sters				$\mathbf{P}^*$
	1 (n=475)		2 (n	=398)	3 (n:	=348)	4 (n=	=259)	_
	Mean	SD	Mean	SD	Mean	SD	Mean	SD	
Age, years	41.2	12.7 <sup>2,3</sup>	39.2	14.24	41.2	12.74	36.7	11.5	<0.001
Female, %	5	6.0	6	7.3	4	7.4	64	1.1	0.79
Ethnicity, %									
Caucasian	9:	5.6	9	6.5	9′	7.7	98	3.1	0.16
Occupation, %									
Professional and managerial	44	$1.2^{4}$	3	7.8	39	9.4	32	2.2	0.014
Intermediate occupations	2:	5.9	2	2.4	23	8.5	28	3.7	0.16
Routine and manual	7	.44	ć	5.8	12	2.9	14	1.3	0.006
Student	1:	3.5	2	1.7	9	.5	14	1.7	0.18
Not currently working	9	0.1	1	1.3	9	.8	10	).1	0.38
Anthropometrics									
Body weight, kg	74.6	$15.1^{3}$	70.5	$15.0^{3,4}$	80.3	$16.0^{4}$	74.1	16.3	< 0.001
BMI, kg/m <sup>2</sup>	25.4	$4.4^{2,3}$	24.1	$4.4^{3,4}$	26.8	4.9	26.0	5.7	< 0.001
Waist circumference, cm	85.4	$13.0^{3}$	81.8	13.2	90.4	14.1	85.9	14.1	< 0.001
Physical activity									
PAL	1.8	$0.2^{2,4}$	1.7	$0.2^{3}$	1.8	$0.2^{4}$	1.7	0.2	< 0.001
SB, min/d	746	73	742	77	750	76	744	7	0.96
Dietary conditions, %									
Want to lose weight	46	$5.1^{4}$	4	1.2	48	8.6	57	7.5	0.013
Restricted diet	$\epsilon$	5.1	1	1.6	3	.7	5	.8	0.47
Medication use, %									
Prescribed medication	2	6.1	3	5.7	29	9.9	27	7.0	0.79
Non-prescribed medication	8	3.6	1	0.6	9	.2	11	1.2	0.18
Health and disease									
Current smoker, %	9	$.8^{4}$	Ģ	0.0	10	0.3	22	2.0	0.005
Total cholesterol, mmol/L	4.6	0.9	4.5	1.0	4.7	1.0	4.6	0.9	0.09
High blood pressure, %	8	3.2	7	7.0	9	.8	5	.8	0.89
Heart disease, %	2	2.1	1	1.8	0	.6	1	.2	0.17

Values represent means and SD or percentages; PAL, physical activity level; SB, sedentary behaviour

<sup>\*,</sup> ANOVA and logistic regression were used to test for significant differences across clusters in continuous and categorical variables, respectively. Analyses were adjusted for age, sex, BMI, PAL, smoking habits and country. Post hoc Tukey tests (continuous data) and logistic regression (categorical) were used to test for significant differences between clusters. Superscripts denote where the differences lie across the clusters. For example, 2 means significantly different from cluster 2.

**Table 3** Healthy Eating Index (HEI) score and its constituents at baseline and month 6 by clusters of adherence to recommendations

				Clu	ster				P†
	1 (n=475)		2 (n=	=398)	3 (n=	348)	4 (n=259)		
	Mean	SD	Mean	SD	Mean	SD	Mean	SD	
Baseline score									
Total HEI	53.3	$8.9^{2,3,4}$	50.5	$8.9^{3,4}$	47.5	$8.9^{4}$	41.8	10.1	< 0.001
Fatty acid ratio*	3.2	$2.4^{2,3,4}$	2.2	$2.4^{3,4}$	1.7	1.7	2.0	2.0	< 0.001
Protein	3.7	$0.7^{2,3,4}$	3.2	$0.6^{3,4}$	3.5	0.6	3.5	0.7	< 0.001
Salt	0.1	0.5	0.1	0.7	0.1	0.6	0.1	0.6	0.002
Empty calories	8.8	$4.0^{2}$	7.7	4.3	8.5	3.8	7.5	4.1	0.012
Refined grains	6.1	$3.7^{2,3,4}$	4.8	3.7	4.4	3.7	4.7	4.0	< 0.001
Seafood and plant protein	5.0	$0.2^{2,3,4}$	4.5	1.0	4.3	1.1	4.4	1.1	< 0.001
Fruit	3.8	$1.3^{3}$	3.8	$1.4^{3}$	3.3	1.5	3.3	1.5	< 0.001
Whole fruit	4.2	$1.3^{3,4}$	4.1	$1.3^{3,4}$	3.6	1.5	3.6	1.6	< 0.001
Vegetables	2.5	$1.1^{3,4}$	2.3	$1.1^{3,4}$	2.0	0.9	2.1	1.1	< 0.001
Greens and beans	4.2	$1.1^{2,3,4}$	3.8	$1.3^{3}$	3.5	1.3	3.7	1.4	< 0.001
Wholegrains	7.3	3.5	9.5	1.2	8.8	1.9	2.9	2.2	< 0.001
Dairy products	4.7	$2.6^{2,3,4}$	4.7	$2.7^{4}$	4.3	$2.2^{4}$	4.4	2.7	0.27
Follow up score									
Total HEI	55.7	$9.1^{1,3,4}$	53.3	$9.6^{4}$	51.4	8.7	48.0	10.3	< 0.001
Fatty acid ratio <sup>1</sup>	3.8	$2.6^{2,3,4}$	3.1	$2.7^{3}$	2.5	2.1	2.6	2.2	< 0.001
Protein	3.8	$0.7^{2,3,4}$	3.3	$0.6^{3,4}$	3.6	0.6	3.6	0.6	< 0.001
Salt	0.1	0.6	0.2	$0.9^{3}$	0.1	0.6	0.1	0.6	0.002
Empty calories	8.7	$4.0^{2}$	7.4	4.1	8.8	4.0	8.1	4.1	0.002
Refined grains	6.2	$3.8^{4}$	5.4	3.8	5.1	3.8	4.9	3.8	0.004
Seafood and plant protein	5.0	$0.2^{2,3}$	4.7	0.8	4.6	1.0	4.7	±0.9	< 0.001
Fruit	4.1	1.3	4.2	$1.2^{3}$	3.7	1.4	3.7	±1.5	0.009
Whole fruit	4.4	1.2	4.4	1.1	4.1	1.4	4.0	±1.5	0.023
Vegetables	2.8	$1.2^{3,4}$	2.7	$1.3^{3,4}$	2.3	1.0	2.4	1.0	< 0.001
Greens and beans	4.3	$1.0^{2,3}$	4.0	1.2	3.9	1.2	4.1	1.2	0.001
Wholegrains	7.9	$3.1^{2,3,4}$	9.2	$1.9^{4}$	8.5	$2.7^{4}$	5.5	3.7	< 0.001
Dairy products	4.8	2.7	4.7	2.8	4.4	2.3	4.5	2.6	0.52

Values represent means and SD.

<sup>\*,</sup> Fatty acid ratio is the ratio of unsaturated fatty acids (mono- and polyunsaturated fatty acids) to saturated fatty acids

<sup>†</sup> ANOVA were used to test for significant differences across clusters. Models were adjusted for age, sex, body mass index, physical activity level, smoking habits and country. Posthoc Tukey's tests used to test for significant differences between clusters. Superscript numbers denote where the differences lie across the clusters relative to the reference category (1). For example, 2 means significantly different from cluster 2.

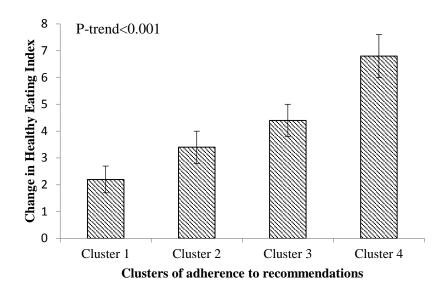


Figure 1

 $\underline{\text{Supplementary Table 1. Summary of criteria for assessing dietary intakes}^*}$ 

			Deficient	Adequate	In excess
		Food g	roups		
Fruit and vegetable	es, g/d		<400	<u>≥</u> 400	NA
Wholegrains, g/d			<50	<u>&gt;</u> 50	NA
Dairy products, g/c	i		<600	<u>&gt;</u> 600	NA
Oily fish, g/wk			<150	<u>≥</u> 150	NA
Red meat, g/wk			NA	<u>&lt;</u> 450	>450
		Nutri	ients		
Protein, g/kg body	weight		<0.66	<u>&gt;</u> 0.66 & <u>&lt;</u> 2.4	>2.4
Carbohydrate, % of	f total energy		<45	45-65	>65
Total fat, % of total	l energy		<20	20-35	>35
Monounsaturated,	% of total ener	gy	<15	15-20	>20
Polyunsaturated, %	6 of total energy	<i>'</i>	<6	6-11	>11
Saturated fat, % of	total energy		<10	<u>&gt;</u> 10 & <u>&lt;</u> 15	>15
Salt, g/d		18-50yrs	<u>&lt;</u> 3.75	>3.75 & <u>&lt;</u> 5.75	>5.75
		51-70yrs	<u>&lt;</u> 3.25	>3.25 & <u>&lt;</u> 5.75	>5.75
		>70yrs	<3	<u>&gt;</u> 3 & <5.75	>5.75
Omega-3, % of tota	al energy		<0.2	<u>&gt;</u> 0.2 & <0.6	<u>&gt;</u> 0.6
Fibre, g/d	Males	18-50yrs	<28	<u>&gt;</u> 28 & <38	<u>&gt;</u> 38
		>50yrs	<20	<u>&gt;</u> 20 & <30	<u>&gt;</u> 30
	Females	18-50yrs	<15	<u>&gt;</u> 15 & <25	<u>≥</u> 25
		>50yrs	<14	<u>&gt;</u> 14 & <21	<u>≥</u> 21
Calcium, mg/d	Males	18-70yrs	<800	<u>&gt;</u> 800 & <u>&lt;</u> 2500	>2500
		>70yrs	<1000	<u>&gt;</u> 1000 & <u>&lt;</u> 2500	>2500
	Females	18-50yrs	<800	<u>&gt;</u> 800 & <u>&lt;</u> 2500	>2500
		>50yrs	<1000	<u>&gt;</u> 1000 & <u>&lt;</u> 2500	>2500
Iron, mg/d	Males	>18yrs	<u>&gt;</u> 4 & <6	<u>&gt;</u> 6.0 & <u>&lt;</u> 45	>45
	Females	18-50yrs	<8.1	<u>&gt;</u> 8.1 & <u>&lt;</u> 45	>45
		>50yrs	<5	<u>≥</u> 5 & <u>&lt;</u> 45	>45
Vitamin A, μg/d	Males		<625	<u>&gt;</u> 625 & <u>&lt;</u> 3000	>3000
	Females		<500	<u>&gt;</u> 500 & <u>&lt;</u> 3000	>3000
Folate, μg/d			<320	<u>&gt;</u> 320 & <u>&lt;</u> 1000	>1000
Thiamin, mg/d	Males		<0.8	<u>&gt;</u> 0.8 & ≤1.0	>1.0
	Females		<0.7	<u>&gt;</u> 0.7 & <u>&lt;</u> 0.9	>0.9
Riboflavin, mg/d	Males		<0.9	<u>&gt;</u> 0.9 & <u>&lt;</u> 1.1	>1.1
	Females		<0.7	<u>&gt;</u> 0.7 & <u>&lt;</u> 0.9	>0.9
Vitamin B12, μg/d			<1.6	≥1.6 & <u>&lt;</u> 2.0	>2.0
Vitamin C, mg/d	Males		<75	<u>&gt;</u> 75 & <u>&lt;</u> 2000	>2000
	Females		<60	≥60 & <u>&lt;</u> 2000	>2000

<sup>\*,</sup> Cut-offs were used to deliver personalized dietary advice during the intervention (20-23)

Supplementary Table 2. Percentage of individuals meeting current European dietary recommendations at baseline

	Meet recommendation				
	Percentage	95% CI			
Food group intake, %					
Fruit and vegetables	52.0	45.7-58.1			
Oily fish	32.1	18.7-49.3			
Red meat	50.5	39.8-61.3			
Wholegrains	74.2	51.9-88.5			
Dairy products	13.7	9.2-19.9			
Nutrient intake, %					
Total fat	50.4	43.5-57.3			
Saturated fat	54.3	45.2-63.0			
Mono-unsaturated fat	24.3	16.0-35.0			
Poly-unsaturated fat	36.2	28.2-45.1			
Protein	91.1	87.7-93.6			
Carbohydrate	55.6	47.4-63.6			
Salt	7.4	3.6-14.8			
Dietary fibre	45.5	35.9-55.6			
Calcium	73.8	65.8-80.5			
Folate	61.4	48.5-72.8			
Iron	95.1	91.8-97.1			
Riboflavin	95.5	89.9-98.0			
Thiamine	97.1	92.6-98.9			
Vitamin A	83.7	77.8-88.3			
Vitamin B12	98.6	96.9-99.4			
Vitamin C	90.1	84.7-93.8			

Values represent percentages (95% CI) of individuals meeting current European dietary recommendations (20-23)

Supplementary Table 3. Description of dietary clusters and the percentage of individuals within each cluster who met the dietary recommendations at baseline (met recommended intake:  $\checkmark$ ; did not meet recommended intake:  $\ast$ )

	Clusters				
	1	2	3	4	
	(n=475)	(n=398)	(n=348)	(n=259)	
Total, n	475	398	348	259	
Food group					
Oily fish	√ (100%)	<b>×</b> (100%)	<b>×</b> (100%)	× (100%)	
Wholegrains	<b>√</b> (74.1%)	√ (100%)	√ (100%)	× (100%)	
Red meat	<b>×</b> (53.7%)	√ (100%)	<b>×</b> (100%)	√ (50.2%)	
Fruit and vegetables	√ (69.3%)	<b>×</b> (50.3%)	<b>×</b> (52.3%)	<b>×</b> (70.7%)	

Values represent the percentage of individuals meeting the following recommendations: Fruit and vegetables >5 servings/day; Oily fish >1 serving/week; Wholegrains >3 servings/day; Red meat <3 servings/week (20-23)

Supplementary Table 4 Percentage of individuals meeting nutrient-based guidelines by clusters of adherence to recommendations at baseline\*

	Clusters				P†
	1 (n=475)	2 (n=398)	3 (n=348)	4 (n=259)	_
Total fat, % energy	50.5 <sup>2,4</sup>	58.5	50.0	38.2	0.046
SFA, % energy	$62.1^{3,4}$	53.5	50.6	46.0	< 0.001
MUFA, % energy	29.1	12.6	22.1	36.3	0.68
PUFA, % energy	$42.1^{3,4}$	36.2	32.2	30.9	0.005
Protein, g/kg/d	85.9	96.5	93.4	89.2	0.99
Carbohydrate, % energy	46.5	75.6	54.0	43.6	0.93
Dietary fibre, g/d	56.24	50.8	50.3	11.6	< 0.001
Salt, g/d	$4.6^{2,4}$	11.3	0.0	16.6	0.034

Values represent percentages of individuals that meet the dietary guidelines:

<sup>\*,</sup> Dietary recommendations: Total fat: 20-35 % energy; SFA: 10-15% energy; MUFA: 15-20% energy; PUFA: 6-11% energy; protein: 0.66-2.4g/kg/day; carbohydrate: 45-65% energy; dietary fibre: males (18-50yrs  $\geq 38g/day$ ;  $>50yrs \geq 30g/day$ ) and females (18-50yrs  $\geq 25g/day$ ;  $>50yrs \geq 21g/day$ ); salt:  $18-50yrs \leq 3.75g/day$ ;  $51-70yrs \leq 3.25g/day$ ;  $>70yrs \leq 3g/day$ 

<sup>†,</sup> Logistic regression was used to test for significant differences across and between clusters (cluster 1 was used as the base category) (20; 21; 22; 23).

Supplementary Table 5 Percentage of individuals meeting dietary recommendations by clusters of adherence to recommendations after exclusion of 3SD of each of the four dietary components at baseline (met recommended intake: ✓; did not meet recommended intake: ×)

	Clusters				
	1	2	3	4	
	(n=475)	(n=398)	(n=348)	(n=259)	
Total, n	439	341	328	275	
Food group					
Oily fish	<b>√</b> (93.6%)	<b>×</b> (100%)	× (100%)	<b>×</b> (100%)	
Fruit and vegetables	√ (68.8%)	√ (100%)	<b>×</b> (86.3%)	× (100%)	
Red meat	<b>×</b> (55.6%)	<b>√</b> (53.7%)	<b>√</b> (100%)	<b>×</b> (100%)	
Wholegrains	<b>√</b> (68.8%)	√ (100%)	√ (86.3%)	√ (100%)	

Values represent the percentage of individuals meeting the following recommendations: Fruit and vegetables >5 servings/day; Oily fish >1 serving/week; Wholegrains >3 servings/day; Red meat <3 servings/week (20-23)