

1 **Carbohydrate quality index and mortality risk in the PREDIMED cohort**

2 **Supplemental Methods**

3 **Exposure Assessment**

4 Determination of each of the assessed dimensions of carbohydrate quality (glycemic
5 index, dietary fiber, whole grains to total grains ratio, and solid carbohydrates to total
6 carbohydrates ratio) were defined based on responses to the Spanish validated version of
7 the 137-item, semi-quantitative food frequency questionnaire administered by trained
8 dietitians during face-to-face interviews. Intake of each food item was calculated by
9 multiplying serving size by frequency of consumption, where there were nine options for
10 the average frequency of consumption from never/almost never to at least six times per
11 day, and nutrients (e.g., dietary fiber, carbohydrates) were derived from Spanish food
12 composition tables ^{1,2}. Carbohydrate quality dimensions were defined as follows:

13 Glycemic index was determined using international tables ³.

14 Dietary fiber was determined based on derivations from the Spanish food
15 composition tables in grams per day ^{1,2}.

16 Whole grains intake was calculated based on FFQ responses to “whole grains:
17 muesli, oat flakes, all-bran” [translated from the Spanish: “Cereales integrales: muesli,
18 copos avena, all bran (30 gr.)”], “Whole-wheat and whole grain bread” [“Pan negro o
19 integral (3 rodajas, 75 gr.)”, and [Galletas integrales o de fibra (4-6 unidades, 50 gr.)”] by
20 multiplying serving size by frequency of consumption and summing intakes across the
21 noted relevant variables. Refined grain intake was calculated based on food frequency
22 questionnaire responses to “white bread” [translated from Spanish: “Pan blanco, pan de
23 molde (3 rodajas, 75 gr.)”], “white rice” [“Arroz blanco (60 gr. en crudo)”], “pasta:
24 noodles, macaroni, spaghetti, others” [“Pasta: fideos, macarrones, espaguetis, otras (60

25 gr. en crudo)”, “breakfast cereals” [Cereales desayuno (30 gr.)]”, pizza [“Pizza (1 ración,
26 200 gr.)”], “Maria cookies (4-6 pieces, 50 gr)” [“Galletas tipo Maria (4-6 unidades, 50
27 gr.)”], “chocolate cookies (4 pieces, 50 gr)” [“Galletas con chocolate (4 unidades, 50
28 gr.)”], “homemade cakes and pastries (50 gr)” [“Reposteria y bizcochos hechos en casa
29 (50 gr.)”], “croissants, ensaimada, tea biscuits, or other commercial pastries... (one, 50
30 gr)” [“Croissant, ensaimada, pastas de té u otra bollería industrial comercial (uno, 50
31 gr.)”], “donuts (one)” [“Donuts (uno)”], “muffins (1-2 pieces)” [“Magdalenas (1-2
32 unidades)”], “cakes (one, 50 gr)” [“Pasteles (uno, 50 gr.)”], chocolates and pralines (30
33 gr)” [“Chocolates y bombones (30 gr.)”], “cocoa powder - instant cocoa (1 dessert
34 spoon)” [“Cacao en polvo-cacaos solubles (1 cucharada de postre)”], “turrón (1/8 bar, 40
35 gr)” [“Turrón (1/8 de barra, 40 gr.)”], “shortbread, marzipan (90 gr)” [“Mantecados,
36 mazapan (90 gr.)”] by multiplying serving size by frequency of consumption and
37 summing intakes across the noted relevant variables. Total grains intake was calculated
38 by summing the intakes of whole grains, refined grains and their products.

39 Solid carbohydrates included the carbohydrate content as derived from Spanish
40 food composition tables contained in all solid foods assessed in the food frequency
41 questionnaire. Liquid carbohydrates were calculated as the sum of carbohydrates from
42 sugar-sweetened beverages [translated from the Spanish: Bebidas carbonatadas con
43 azúcar: bebidas con cola, limonadas, tónicas, etc. (1 botellín, 200 cc)], , and fruit juices
44 [Zumo de naranja natural (1 vaso, 200 cc), Zumos naturales de otras frutas (1 vaso, 200
45 cc), Zumos de frutas en botella o enlatados (200 cc), Frutas en almíbar o en su jugo (2
46 unidades)]. Total carbohydrates intake was calculated by summing the intakes of solid
47 carbohydrates and liquid carbohydrates.

48 These four components were selected based on the pre-defined composite
49 carbohydrate quality index (CQI) ⁴, to capture complementary aspects of carbohydrate

50 quality with established but distinct associations with cardiometabolic health, including
51 glycemic control (GI), lipid metabolism and satiety (dietary fiber), nutrient density and
52 food structure (whole grains), and excess energy intake and metabolic dysregulation
53 (solid-to-liquid carbohydrates ratio) ^{5,6}.

54 Each CQI component was assigned equal weight. This aligns with the previously
55 defined composite CQI and given the absence of robust comparative evidence to support
56 differential weighting across components, equal weighting was applied as a conservative
57 and transparent approach that minimizes arbitrary bias and ensures balanced
58 representation of each dimension. This equal weighting of CQI components also reflects
59 a conceptual assumption rather than an empirically derived weighting scheme. The CQI
60 is intended to capture the multidimensional and potentially synergistic nature of
61 carbohydrate quality, rather than to quantify the relative contribution of individual
62 components.

63 **References:**

64 1J. Mataix-Solera, G. L, E. Martinez de Victoria and J. Llopis, *Tabla de Composición*
65 *de Alimentos 4ª Edición Corregida y Aumentada*, 2003.

66 2O. Moreiras Tuny, Á. Carbajal Azcona, L. Cabrera Forneiro and C. Cuadrado Vives,
67 *Tablas de composición de alimentos*, Pirámide, 2016.

68 3F. S. Atkinson, K. Foster-Powell and J. C. Brand-Miller, *Diabetes Care*, 2008, **31**,
69 2281–2283.

70 4I. Zazpe, A. Sánchez-Taínta, S. Santiago, C. de la Fuente-Arrillaga, M. Bes-Rastrollo,
71 J. A. Martínez, M. Á. Martínez-González, and SUN Project Investigators, *Br. J. Nutr.*,
72 2014, **111**, 2000–2009.

73 5L. S. A. Augustin, C. W. C. Kendall, D. J. A. Jenkins, W. C. Willett, A. Astrup, A. W.
74 Barclay, I. Björck, J. C. Brand-Miller, F. Brighenti, A. E. Buyken, A. Ceriello, C. La

- 75 Vecchia, G. Livesey, S. Liu, G. Riccardi, S. W. Rizkalla, J. L. Sievenpiper, A.
76 Trichopoulou, T. M. S. Wolever, S. Baer-Sinnott and A. Poli, *Nutr. Metab.*
77 *Cardiovasc. Dis. NMCD*, 2015, **25**, 795–815.
- 78 6A. Reynolds, J. Mann, J. Cummings, N. Winter, E. Mete and L. Te Morenga, *Lancet*
79 *Lond. Engl.*, 2019, **393**, 434–445.
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Supplemental Tables

Table S1.- Baseline characteristics of the PREDIMED participants by cumulative average carbohydrate quality index

Characteristic ^a	Quintiles					Total	P value ^b
	1	2	3	4	5		
Participants, No. (%)	1,979 (27.4%)	1,456 (20.2%)	1,513 (21.0%)	1,204 (16.7%)	1,058 (14.7%)	7,210 (100.0%)	
Age, mean (SD), years	67.4 (6.4)	67.4 (6.2)	66.9 (6.2)	66.7 (6.1)	66.5 (5.9)	67.0 (6.2)	<0.001
Women, No. (%)	1,050 (53.1%)	790 (54.3%)	856 (56.6%)	754 (62.6%)	693 (65.5%)	4,143 (57.5%)	<0.001
Intervention group, No. (%)							
MedDiet + EVOO	638 (32.2%)	497 (34.1%)	515 (34.0%)	456 (37.9%)	367 (34.7%)	2,473 (34.3%)	<0.001
MedDiet + Nuts	569 (28.8%)	474 (32.6%)	531 (35.1%)	395 (32.8%)	390 (36.9%)	2,359 (32.7%)	
Low fat diet	772 (39.0%)	485 (33.3%)	467 (30.9%)	353 (29.3%)	301 (28.4%)	2,378 (33.0%)	
Educational level, No. (%)							
Primary education or less	1,547 (78.2%)	1,131 (77.7%)	1,160 (76.7%)	943 (78.3%)	820 (77.5%)	5,601 (77.7%)	0.969
Secondary	300 (15.2%)	223 (15.3%)	235 (15.5%)	176 (14.6%)	161 (15.2%)	1,095 (15.2%)	
College	132 (6.7%)	102 (7.0%)	118 (7.8%)	85 (7.1%)	77 (7.3%)	514 (7.1%)	
Smoking status, No. (%)							
Never smoked	146 (7.4%)	87 (6.0%)	108 (7.1%)	94 (7.8%)	87 (8.2%)	522 (7.2%)	0.154
Former smoker	1,487 (75.1%)	1,142 (78.4%)	1,177 (77.8%)	910 (75.6%)	786 (74.3%)	5,502 (76.3%)	
Current smoker	346 (17.5%)	227 (15.6%)	228 (15.1%)	200 (16.6%)	185 (17.5%)	1,186 (16.4%)	
Physical activity, mean (SD), MET/min/d	219.6 (246.9)	218.7 (212.7)	244.0 (257.9)	238.5 (248.6)	262.4 (270.6)	234.0 (247.2)	<0.001
BMI, mean (SD), kg/m ²	30.2 (3.8)	30.0 (3.7)	30.0 (3.8)	29.8 (4.0)	29.7 (4.1)	30.0 (3.9)	0.016
Obesity, No. (%)	958 (48.4%)	701 (48.1%)	736 (48.6%)	528 (43.9%)	460 (43.5%)	3,383 (46.9%)	0.222
Type 2 diabetes, No. (%)	944 (47.7%)	694 (47.7%)	715 (47.3%)	607 (50.4%)	566 (53.5%)	3,526 (48.9%)	0.008
Waist circumference, mean (SD), cm	101.4 (10.1)	101.1 (10.0)	100.8 (10.1) 145.7 (20.3)	99.3 (10.5)	98.8 (11.1)	100.5 (10.3)	<0.001
Systolic blood pressure, mean (SD), mm Hg	145.7 (20.4)	146.4 (19.3)		144.9 (20.1)	144.5 (20.5)	145.5 (20.1)	0.516

Diastolic blood pressure, mean (SD), mm Hg	81.9 (10.2)	82.1 (10.7)	81.6 (10.4)	80.7 (10.6)	82.7 (31.9)	81.8 (15.5)	0.269
Blood parameters, mean (SD), mg/dL							
Glucose	142.3 (445.4)	130.1 (262.0)	137.5 (364.5)	133.9 (289.9)	126.0 (48.7)	135.0 (332.5)	0.710
Total cholesterol	220.4 (313.4)	209.0 (36.3)	225.5 (358.4)	220.4 (286.2)	209.8 (36.3)	217.6 (260.9)	0.369
Low-density lipoprotein	134.8 (19.2)	133.0 (18.9)	133.3 (17.7)	132.6 (18.0)	133.5 (17.5)	133.6 (18.4)	0.005
High-density lipoprotein	52.7 (8.0)	52.9 (8.0)	53.5 (8.1)	53.1 (8.0)	53.1 (8.1)	53.1 (8.1)	0.042
Triglycerides	152.1 (323.0)	140.3 (87.3)	148.4 (365.7)	146.0 (295.6)	131.5 (76.8)	144.9 (271.5)	0.326

Abbreviations: BMI, body mass index (calculated as weight in kilograms divided 2 by height in meters squared); CVD, cardiovascular disease; EVOO, extra virgin olive oil; MedDiet, Mediterranean Diet; METs, metabolic equivalents; PREDIMED = Prevención con Dieta Mediterránea.

^aData are presented as No. (%) or mean (SD) for categorical and continuous variables, respectively.

^b*P* values for intergroup differences by tertiles of carbohydrate quality index in the overall population were calculated with the Pearson χ^2 test or univariate ANOVA, as appropriate. Significance was set at *P* values less than 0.05 and boldfaced.

Table S2.- Dietary intake# by baseline carbohydrate quality index categories

Characteristic ^a	Quintiles					Total	P value ^b
	1	2	3	4	5		
Participants, No. (%)	1,979 (27.4%)	1,456 (20.2%)	1,513 (21.0%)	1,204 (16.7%)	1,058 (14.7%)	7,210 (100.0%)	
Carbohydrate quality index	7.7 (1.3)	10.5 (0.5)	12.5 (0.5)	14.5 (0.5)	17.0 (1.1)	11.7 (3.3)	<0.001
Glycemic index	56.1 (3.5)	54.3 (3.7)	52.8 (4.0)	51.7 (3.7)	50.2 (3.4)	53.5 (4.2)	<0.001
Fiber, g/day	20.9 (4.2)	23.9 (4.7)	26.2 (5.7)	28.6 (5.4)	31.7 (5.9)	25.5 (6.3)	<0.001
Whole-grain-to-total grain ratio	0.0 (0.0)	0.1 (0.1)	0.2 (0.2)	0.4 (0.3)	0.5 (0.2)	0.2 (0.3)	<0.001
Solid to total carbohydrates ratio	1.0 (0.0)	1.0 (0.0)	1.0 (0.0)	1.0 (0.0)	1.0 (0.0)	1.0 (0.0)	<0.001
Energy, kcal/day	2179 (411)	2226 (431)	2185 (440)	2172 (385)	2158 (372)	2186 (412)	<0.001
Carbohydrates, %	41.7 (5.5)	41.3 (5.5)	41.3 (5.6)	41.4 (5.0)	41.0 (5.2)	41.4 (5.4)	0.029
Protein, %	15.8 (2.1)	16.3 (2.1)	16.6 (2.2)	17.0 (2.0)	17.4 (2.2)	16.5 (2.2)	<0.001
Fat, %	39.8 (5.1)	40.0 (5.1)	39.6 (5.2)	40.0 (4.8)	39.8 (5.0)	39.8 (5.1)	0.268
Alcohol, g/day	8.9 (12.9)	8.2 (11.9)	8.2 (11.4)	5.7 (9.0)	5.8 (9.5)	7.6 (11.4)	<0.001
Cereals, g/day	225.3 (72.2)	220.6 (72.0)	210.6 (74.0)	205.9 (62.4)	203.0 (60.4)	214.7 (69.9)	<0.001
Fruits, g/day	318.0 (121.2)	381.3 (138.0)	404.6 (149.9)	424.8 (148.0)	450.3 (148.7)	386.2 (147.3)	<0.001
Vegetables, g/day	279.8 (87.0)	323.4 (95.7)	351.1 (101.6)	365.9 (109.1)	392.4 (113.3)	334.5 (107.3)	<0.001
Legumes, g/day	19.2 (7.0)	21.2 (7.9)	22.6 (9.5)	23.1 (9.8)	24.8 (10.3)	21.8 (9.0)	<0.001
Nuts, g/day	11.0 (11.2)	13.8 (12.8)	15.1 (13.2)	16.3 (14.1)	16.9 (13.3)	14.2 (13.0)	<0.001
Dairy products, g/day	333.3 (168.0)	364.9 (173.1)	377.8 (177.4)	395.8 (171.5)	419.1 (185.9)	372.0 (176.7)	<0.001
Meat, g/day	123.3 (40.0)	124.6 (41.5)	121.3 (39.8)	119.6 (37.9)	117.4 (40.1)	121.7 (40.0)	<0.001
Red meat, g/day	80.6 (35.6)	78.9 (36.4)	74.8 (34.5)	71.3 (32.1)	68.0 (34.5)	75.6 (35.1)	<0.001
Eggs, g/day	20.3 (8.6)	20.6 (7.9)	19.9 (7.7)	20.1 (7.3)	19.8 (7.5)	20.2 (7.9)	0.051
Fish, g/day	95.1 (37.0)	100.8 (35.8)	101.5 (36.3)	105.3 (37.0)	108.7 (46.2)	101.3 (38.4)	<0.001
Olive oil, ml/day	42.9 (13.5)	43.4 (13.3)	41.8 (13.3)	42.3 (12.6)	42.2 (12.7)	42.6 (13.2)	0.016
Sugar and honey, g/day	8.3 (11.0)	7.4 (10.0)	7.0 (9.5)	5.9 (8.5)	4.6 (7.2)	6.9 (9.6)	<0.001
Cookies, g/day	21.8 (21.1)	21.3 (22.8)	18.6 (19.4)	17.5 (19.0)	13.0 (14.6)	19.0 (20.2)	<0.001
Sugar -sweetened beverages, g/day	52.5 (80.5)	30.2 (52.7)	27.8 (49.6)	24.8 (44.6)	13.8 (35.7)	32.5 (59.6)	<0.001
Coffee and tea, mL/day	34.8 (41.9)	32.7 (41.0)	31.9 (40.6)	28.4 (38.5)	32.7 (42.6)	32.4 (41.0)	<0.001

Adherence to MedDiet, points	6.9 (3.0)	7.7 (2.8)	7.8 (2.9)	8.1 (2.8)	8.2 (3.0)	7.6 (2.9)	<0.001
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#Dietary intakes measured at baseline

Abbreviations: MedDiet, Mediterranean Diet.

^aData are presented as No. (%) or mean (SD) for categorical and continuous variables, respectively.

^b*P* values for intergroup differences by tertiles of carbohydrate quality index in the overall population were calculated with the Pearson χ^2 test or univariate ANOVA, as appropriate. Significance was set at *P* values less than 0.05 and boldfaced.

Table S3.- Relationship between quintiles of cumulative average carbohydrate quality index and all-cause and specific-cause mortality in the PREDIMED cohort^a.

	Crude incidence rate (95% CI) ^b	No. Deaths (%)	Models			
			Model 1 ^c	Model 2 ^d	Model 3 ^e	Model 4 ^f
All-cause mortality						
Continuous# (n=7,210)	10.0 (9.1 to 11.0)	425 (5.9%)	0.96 [0.94 to 0.99]	0.97 [0.94 to 1.00]	0.97 [0.94 to 1.00]	0.98 [0.94 to 1.01]
Quintiles						
1 (n=1,979)	13.3 (11.4 to 15.7)	155 (7.8%)	Reference	Reference	Reference	Reference
2 (n=1,456)	9.3 (7.5 to 11.6)	80 (5.5%)	0.71 [0.54 to 0.93]	0.74 [0.56 to 0.97]	0.74 [0.56 to 0.97]	0.76 [0.57 to 1.00]
3 (n=1,513)	8.1 (6.4 to 10.2)	73 (4.8%)	0.65 [0.49 to 0.86]	0.69 [0.52 to 0.92]	0.69 [0.52 to 0.92]	0.71 [0.53 to 0.94]
4 (n=1,204)	9.3 (7.3 to 11.8)	66 (5.5%)	0.79 [0.59 to 1.06]	0.83 [0.61 to 1.12]	0.82 [0.61 to 1.11]	0.88 [0.64 to 1.20]
5 (n=1,058)	8.2 (6.3 to 10.9)	51 (4.8%)	0.76 [0.54 to 1.05]	0.81 [0.58 to 1.13]	0.81 [0.58 to 1.13]	0.88 [0.62 to 1.24]
P-trend*			0.054	0.132	0.130	0.317
CVD mortality						
Continuous# (n=7,210)	2.4 (2.0 to 2.9)	103 (1.4%)	0.95 [0.89 to 1.02]	0.97 [0.91 to 1.04]	0.97 [0.90 to 1.04]	0.97 [0.90 to 1.03]
Quintiles*						
1 (n=1,979)	3.5 (2.6 to 4.8)	41 (2.0%)	Reference	Reference	Reference	Reference
2 (n=1,456)	2.3 (1.5 to 3.6)	20 (1.4%)	0.68 [0.40 to 1.16]	0.73 [0.43 to 1.26]	0.70 [0.41 to 1.20]	0.71 [0.42 to 1.20]
3 (n=1,513)	1.8 (1.1 to 2.9)	16 (1.1%)	0.55 [0.31 to 1.00]	0.60 [0.33 to 1.11]	0.58 [0.32 to 1.05]	0.56 [0.32 to 1.00]
4 (n=1,204)	1.8 (1.1 to 3.1)	13 (1.1%)	0.61 [0.31 to 1.17]	0.67 [0.35 to 1.30]	0.65 [0.33 to 1.25]	0.65 [0.33 to 1.27]
5 (n=1,058)	2.1 (1.2 to 3.6)	13 (1.2%)	0.76 [0.39 to 1.48]	0.85 [0.43 to 1.72]	0.86 [0.43 to 1.73]	0.89 [0.45 to 1.74]
P-trend*			0.210	0.391	0.371	0.388

Cancer mortality						
Continuous# (n=7,210)	4.0 (3.4 to 4.6)	169 (2.3%)	0.93 [0.88 to 0.98]	0.94 [0.89 to 0.99]	0.93 [0.89 to 0.98]	0.94 [0.89 to 0.99]
Quintiles						
1 (n=1,979)	5.8 (4.6 to 7.4)	67 (3.4%)	Reference	Reference	Reference	Reference
2 (n=1,456)	4.1 (2.9 to 5.7)	35 (2.4%)	0.70 [0.46 to 1.07]	0.72 [0.47 to 1.09]	0.73 [0.48 to 1.12]	0.77 [0.50 to 1.18]
3 (n=1,513)	2.9 (2.0 to 4.2)	26 (1.7%)	0.51 [0.32 to 0.82]	0.54 [0.34 to 0.85]	0.54 [0.34 to 0.86]	0.55 [0.34 to 0.90]
4 (n=1,204)	3.6 (2.5 to 5.4)	26 (2.2%)	0.70 [0.44 to 1.13]	0.75 [0.47 to 1.21]	0.77 [0.48 to 1.23]	0.82 [0.50 to 1.34]
5 (n=1,058)	2.4 (1.5 to 4.0)	15 (1.4%)	0.49 [0.28 to 0.87]	0.52 [0.29 to 0.91]	0.52 [0.29 to 0.92]	0.54 [0.30 to 0.98]
P-trend*			0.008	0.016	0.018	0.040
Other-causes of mortality						
Continuous# (n=7,210)	3.6 (3.1 to 4.2)	153 (2.1%)	1.02 [0.96 to 1.07]	1.01 [0.96 to 1.07]	1.01 [0.96 to 1.07]	1.03 [0.97 to 1.09]
Quintiles						
1 (n=1,979)	4.1 (3.1 to 5.4)	47 (2.4%)	Reference	Reference	Reference	Reference
2 (n=1,456)	2.9 (2.0 to 4.3)	25 (1.7%)	0.76 [0.47 to 1.24]	0.78 [0.47 to 1.27]	0.77 [0.47 to 1.27]	0.80 [0.48 to 1.32]
3 (n=1,513)	3.4 (2.4 to 4.9)	31 (2.1%)	0.94 [0.60 to 1.50]	1.01 [0.63 to 1.61]	1.01 [0.63 to 1.60]	1.08 [0.67 to 1.74]
4 (n=1,204)	3.8 (2.6 to 5.5)	27 (2.2%)	1.08 [0.66 to 1.78]	1.07 [0.64 to 1.79]	1.07 [0.64 to 1.78]	1.19 [0.71 to 2.00]
5 (n=1,058)	3.7 (2.5 to 5.6)	23 (2.2%)	1.18 [0.71 to 1.97]	1.22 [0.72 to 2.08]	1.22 [0.72 to 2.08]	1.45 [0.82 to 2.57]
P-trend*			0.481	0.430	0.431	0.187

Abbreviations: CI – confidence interval, CQI – carbohydrate quality index, CVD = Cardiovascular Disease, PREDIMED = Prevención con Dieta Mediterránea.

#Cox Regression calculated per unit increase in CQI score with a possible score range of 4-20

*Tests of linear trend were applied for the evaluation of dose–response relationships across quintiles, assigning to each category of the total intake its quintile-specific median and using the resulting variable as continuous.

^aMultivariable regression Cox models stratified by recruiting centre with robust standard errors to account for small deviations from individual randomization. Results are expressed as Hazard Ratios (HR) and 95% Confidence Intervals (95% CI). Cumulative average carbohydrate quality index was treated as a continuous or categorical variable by quintiles. Extremes of total energy intake (≥ 4000 or < 800 kcal/day in men and ≥ 3500 or in women) were excluded. Significance was set at *P* values less than 0.05 and boldfaced. The exposure as continuous is presented per 1 unit increase.

^bCrude incidence rate is expressed per 1000 person-years.

^cModel 1: adjusted for age (years), sex (male or female), intervention group (control, Mediterranean diet + extra virgin olive oil, Mediterranean diet + nuts).

^dModel 2: additionally adjusted for educational level (primary education or less, secondary education, or college/graduate), baseline physical activity (metabolic equivalent task units in min/day), smoking status (never, current, or former), cumulative average dietary alcohol intake (using the linear term and adding a quadratic term, g/day), baseline body mass index (kg/m²), prevalent type 2 diabetes (yes or no) or medication (yes or no), prevalent hypertension (yes or no) or medication (yes or no), prevalent hypercholesterolemia (yes or no) or medication (yes or no).

^eModel 3: additionally adjusted for cumulative average dietary energy intake (kcal/day).

^fModel 4: additionally adjusted for cumulative average dietary protein intake (g/day), dietary saturated fatty acids intake (g/day), dietary monounsaturated fatty acids intake (g/day), and dietary polyunsaturated fatty acids intake (g/day). In the case of CVD and cancer mortality, family history of such diseases were included as covariates.

Table S4.- Sensitivity analysis for the relationship between cumulative average carbohydrate quality index and all-cause and specific-

cause mortality after excluding early cases of mortality in the PREDIMED cohort^a.

	No. Deaths (%)	Models			
		Model 1 ^b	Model 2 ^c	Model 3 ^d	Model 4 ^e
All-cause mortality					
Continuous# (n=7,179)	394 (5.5%)	0.97 [0.93 to 1.00]	0.97 [0.94 to 1.01]	0.97 [0.94 to 1.01]	0.983 [0.95 to 1.01]
Category*					
High (n=5,217)	256 (4.9%)	Reference	Reference	Reference	Reference
Low (n=1,962)	138 (7.0%)	1.33 [1.07 to 1.65]	1.26 [1.01 to 1.57]	1.26 [1.02 to 1.57]	1.23 [0.98 to 1.54]
P-trend**		0.061	0.110	0.124	0.271
CVD mortality					
Continuous# (n=7,179)	94 (1.3%)	0.93 [0.87 to 1.00]	0.95 [0.88 to 1.02]	0.95 [0.88 to 1.02]	0.94 [0.88 to 1.01]
Category*					
High (n=5,217)	55 (1.1%)	Reference	Reference	Reference	Reference
Low (n=1,962)	39 (2.0%)	1.71 [1.11 to 2.64]	1.56 [1.00 to 2.45]	1.61 [1.03 to 2.51]	1.62 [1.06 to 2.47]
P-trend**		0.070	0.156	0.131	0.141
Cancer mortality					
Continuous# (n=7,179)	157 (2.2%)	0.94 [0.89 to 0.99]	0.95 [0.90 to 1.00]	0.95 [0.91 to 1.00]	0.95 [0.90 to 1.01]
Category*					
High (n=5,217)	99 (1.9%)	Reference	Reference	Reference	Reference
Low (n=1,962)	58 (3.0%)	1.46 [1.04 to 2.06]	1.40 [0.99 to 1.97]	1.38 [0.98 to 1.95]	1.34 [0.92 to 1.93]
P-trend**		0.313	0.382	0.431	0.570
Other causes of mortality					
Continuous# (n=7,179)	143 (2.0%)	1.02 [0.96 to 1.07]	1.02 [0.96 to 1.08]	1.02 [0.96 to 1.08]	1.03 [0.97 to 1.09]
Category*					
High (n=5,217)	102 (2.0%)	Reference	Reference	Reference	Reference
Low (n=1,962)	41 (2.1%)	0.96 [0.66 to 1.40]	0.94 [0.64 to 1.37]	0.94 [0.64 to 1.38]	0.89 [0.60 to 1.32]
P-trend**		0.598	0.751	0.796	0.904

Abbreviations: CI – confidence interval, CQI – carbohydrate quality index, CVD = Cardiovascular Disease, PREDIMED = Prevención con Dieta Mediterránea.

#Regression Cox calculated per unit increase in CQI score with a possible score range of 4-20

*Cumulative average carbohydrate quality index treated as a categorical variable by low intake (lowest quintile) vs. high intake (the four upper quintiles merged, as a reference category).

**Tests of linear trend were applied for the evaluation of dose–response relationships across quintiles, assigning to each category of the total intake its quintile-specific median and using the resulting variable as continuous.

^aMultivariable regression Cox models stratified by recruiting centre with robust standard errors to account for small deviations from individual randomization. Results are expressed as Hazard Ratios (HR) and 95% Confidence Intervals (95% CI).

Extremes of total energy intake (≥ 4000 or < 800 kcal/day in men and ≥ 3500 or in women) were excluded. Significance was set at *P* values less than 0.05 and boldfaced. The exposure as continuous is presented per 1 unit increase.

^bModel 1: adjusted for age (years), sex (male or female), intervention group (control, Mediterranean diet + extra virgin olive oil, Mediterranean diet + nuts).

^cModel 2: additionally adjusted for educational level (primary education or less, secondary education, or college/graduate), baseline physical activity (metabolic equivalent task units in min/day), smoking status (never, current, or former), cumulative average dietary alcohol intake (using the linear term and adding a quadratic term, g/day), baseline body mass index (kg/m²), prevalent type 2 diabetes (yes or no) or medication (yes or no), prevalent hypertension (yes or no) or medication (yes or no), prevalent hypercholesterolemia (yes or no) or medication (yes or no).

^dModel 3: additionally adjusted for cumulative average dietary energy intake (kcal/day).

^eModel 4: additionally adjusted for cumulative average dietary protein intake (g/day), dietary saturated fatty acids intake (g/day), dietary monounsaturated fatty acids intake (g/day), and dietary polyunsaturated fatty acids intake (g/day). In the case of CVD and cancer mortality, family history of such diseases were included as covariates.

Table S5.- Relationship between baseline carbohydrate quality index and all-cause and specific-cause mortality in the PREDIMED cohort^a.

	No. Deaths (%)	Models			
		Model 1 ^b	Model 2 ^c	Model 3 ^d	Model 4 ^e
All-cause mortality					
Continuous# (n=7,210)	425 (5.9%)	0.96 [0.93 to 0.99]	0.97 [0.93 to 1.00]	0.97 [0.93 to 1.00]	0.97 [0.94 to 1.00]
Category*					
High (n=5,026)	265 (5.3%)	Reference	Reference	Reference	Reference
Low (n=2,184)	160 (7.3%)	1.23 [1.00 to 1.52]	1.20 [0.97 to 1.48]	1.20 [0.98 to 1.48]	1.18 [0.95 to 1.45]
P-trend**		0.043	0.070	0.068	0.116
CVD mortality					
Continuous# (n=7,210)	103 (1.4%)	0.95 [0.89 to 1.02]	0.97 [0.90 to 1.03]	0.96 [0.90 to 1.03]	0.96 [0.90 to 1.03]
Category*					
High (n=5,026)	60 (1.2%)	Reference	Reference	Reference	Reference
Low (n=2,184)	43 (2.0%)	1.46 [0.97 to 2.19]	1.36 [0.90 to 2.07]	1.40 [0.92 to 2.13]	1.43 [0.95 to 2.16]
P-trend**		0.152	0.273	0.252	0.231
Cancer mortality					
Continuous# (n=7,210)	169 (2.3%)	0.95 [0.89 to 1.00]	0.95 [0.90 to 1.00]	0.95 [0.90 to 1.00]	0.95 [0.90 to 1.01]
Category*					
High (n=5,026)	100 (2.0%)	Reference	Reference	Reference	Reference
Low (n=2,184)	69 (3.2%)	1.37 [0.99 to 1.91]	1.35 [0.97 to 1.87]	1.33 [0.96 to 1.85]	1.30 [0.93 to 1.83]
P-trend**		0.069	0.086	0.091	0.122
Other causes of mortality					
Continuous# (n=7,210)	153 (2.1%)	0.99 [0.93 to 1.04]	0.99 [0.93 to 1.04]	0.99 [0.93 to 1.04]	0.99 [0.94 to 1.05]
Category*					
High (n=5,026)	105 (2.1%)	Reference	Reference	Reference	Reference
Low (n=2,184)	48 (2.2%)	0.95 [0.65 to 1.37]	0.95 [0.66 to 1.37]	0.95 [0.65 to 1.37]	0.91 [0.63 to 1.32]
P-trend**		0.840	0.787	0.785	0.998

Abbreviations: CI – confidence interval, CQI – carbohydrate quality index, CVD = Cardiovascular Disease, PREDIMED = Prevención con Dieta Mediterránea.

#Regression Cox calculated per unit increase in CQI score with a possible score range of 4-20

*Baseline carbohydrate quality index treated as a categorical variable by low intake (lowest quintile) vs. high intake (the four upper quintiles merged, as a reference category).

**Tests of linear trend were applied for the evaluation of dose–response relationships across quintiles, assigning to each category of the total intake its quintile-specific median and using the resulting variable as continuous.

^aMultivariable regression Cox models stratified by recruiting centre with robust standard errors to account for small deviations from individual randomization. Results are expressed as Hazard Ratios (HR) and 95% Confidence Intervals (95% CI).

Extremes of total energy intake (≥ 4000 or < 800 kcal/day in men and ≥ 3500 or in women) were excluded. Significance was set at *P* values less than 0.05 and boldfaced. The exposure as continuous is presented per 1 unit increase.

^bModel 1: adjusted for age (years), sex (male or female), intervention group (control, Mediterranean diet + extra virgin olive oil, Mediterranean diet + nuts).

^cModel 2: additionally adjusted for educational level (primary education or less, secondary education, or college/graduate), baseline physical activity (metabolic equivalent task units in min/day), smoking status (never, current, or former), baseline dietary alcohol intake (using the linear term and adding a quadratic term, g/day), baseline body mass index (kg/m²), prevalent type 2 diabetes (yes or no) or medication (yes or no), prevalent hypertension (yes or no) or medication (yes or no), prevalent hypercholesterolemia (yes or no) or medication (yes or no).

^dModel 3: additionally adjusted for baseline dietary energy intake (kcal/day).

^eModel 4: additionally adjusted for baseline dietary protein intake (g/day), dietary saturated fatty acids intake (g/day), dietary monounsaturated fatty acids intake (g/day), and dietary polyunsaturated fatty acids intake (g/day). In the case of CVD and cancer mortality, family history of such diseases were included as covariates.

Table S6.- Relationship between cumulative average dietary glycemic index and all-cause and specific-cause mortality in the PREDIMED cohort^a.

No. Deaths (%)

Models

		Model 1 ^b	Model 2 ^c	Model 3 ^d	Model 4 ^e
All-cause mortality					
Continuous# (n=7,210)	425 (5.9%)	0.99 [0.86 to 1.14]	0.99 [0.86 to 1.14]	0.99 [0.86 to 1.14]	0.99 [0.86 to 1.14]
Category*					
High (n=5,768)	348 (6.0%)	Reference	Reference	Reference	Reference
Low (n=1,442)	77 (5.3%)	1.15 [0.89 to 1.49]	1.10 [0.85 to 1.43]	1.11 [0.86 to 1.44]	1.02 [0.79 to 1.33]
P-trend**		0.721	0.814	0.737	0.842
CVD mortality					
Continuous# (n=7,210)	103 (1.4%)	1.07 [0.77 to 1.49]	1.07 [0.77 to 1.49]	1.07 [0.77 to 1.49]	1.07 [0.77 to 1.49]
Category*					
High (n=5,768)	85 (1.5%)	Reference	Reference	Reference	Reference
Low (n=1,442)	18 (1.2%)	1.06 [0.63 to 1.78]	0.97 [0.57 to 1.66]	1.09 [0.63 to 1.88]	0.89 [0.53 to 1.50]
P-trend**		0.273	0.270	0.579	0.148
Cancer mortality					
Continuous# (n=7,210)	169 (2.3%)	0.98 [0.78 to 1.22]	0.98 [0.78 to 1.22]	0.98 [0.78 to 1.22]	0.98 [0.78 to 1.22]
Category*					
High (n=5,768)	141 (2.4%)	Reference	Reference	Reference	Reference
Low (n=1,442)	28 (1.9%)	1.08 [0.71 to 1.66]	1.05 [0.68 to 1.61]	0.98 [0.64 to 1.49]	0.97 [0.62 to 1.50]
P-trend**		0.837	0.921	0.728	0.944
Other causes of mortality					
Continuous# (n=7,210)	153 (2.1%)	0.95 [0.77 to 1.18]	0.95 [0.77 to 1.18]	0.95 [0.77 to 1.18]	0.95 [0.77 to 1.18]
Category*					
High (n=5,768)	122 (2.1%)	Reference	Reference	Reference	Reference
Low (n=1,442)	31 (2.1%)	1.29 [0.85 to 1.94]	1.23 [0.82 to 1.86]	1.24 [0.82 to 1.88]	1.21 [0.80 to 1.83]
P-trend**		0.190	0.206	0.185	0.158

Abbreviations: CI – confidence interval, CQI – carbohydrate quality index, CVD = Cardiovascular Disease, PREDIMED = Prevención con Dieta Mediterránea.

#Regression Cox calculated per 5 units increase in glycemic index

*Cumulative average dietary glycemic index treated as a categorical variable by low intake (lowest quintile) vs. high intake (the four upper quintiles merged, as a reference category).

**Tests of linear trend were applied for the evaluation of dose–response relationships across quintiles, assigning to each category of the total intake its quintile-specific median and using the resulting variable as continuous.

^aMultivariable regression Cox models stratified by recruiting centre with robust standard errors to account for small deviations from individual

randomization. Results are expressed as Hazard Ratios (HR) and 95% Confidence Intervals (95% CI). Cumulative average dietary glyce- mic index was treated as a continuous or categorical variable by tertiles. Extremes of total energy intake (≥ 4000 or < 800 kcal/day in men and ≥ 3500 or in women) were excluded. Significance was set at P values less than 0.05 and boldfaced.

^bModel 1: adjusted for age (years), sex (male or female), intervention group (control, Mediterranean diet + extra virgin olive oil, Mediterranean diet + nuts).

^cModel 2: additionally adjusted for educational level (primary education or less, secondary education, or college/graduate), baseline physical activity (metabolic equivalent task units in min/day), smoking status (never, current, or former), cumulative average dietary alcohol intake (using the linear term and adding a quadratic term, g/day), baseline body mass index (kg/m^2), prevalent type 2 diabetes (yes or no) or medication (yes or no), prevalent hypertension (yes or no) or medication (yes or no), prevalent hypercholesterolemia (yes or no) or medication (yes or no).

^dModel 3: additionally adjusted for cumulative average dietary energy intake (kcal/day).

^eModel 4: additionally adjusted for cumulative average dietary protein intake (g/day), dietary saturated fatty acids intake (g/day), dietary monounsaturated fatty acids intake (g/day), and dietary polyunsaturated fatty acids intake (g/day). In the case of CVD and cancer mortality, family history of such diseases were included as covariates.

Table S7.- Relationship between cumulative average dietary fibre and all-cause and specific-cause mortality in the PREDIMED cohort^a.

No. Deaths (%)	Models			
	Model 1 ^b	Model 2 ^c	Model 3 ^d	Model 4 ^e

All-cause mortality					
Continuous # (n=7,210)	425 (5.9%)	0.97 [0.95 to 0.99]	0.98 [0.96 to 1.01]	0.97 [0.95 to 1.00]	0.98 [0.96 to 1.01]
Category*					
High (n=5,768)	309 (5.4%)	Reference	Reference	Reference	Reference
Low (n=1,442)	116 (8.0%)	1.68 [1.32 to 2.12]	1.52 [1.19 to 1.93]	1.67 [1.30 to 2.16]	1.51 [1.16 to 1.97]
P-trend**		0.001	0.001	0.001	0.002
CVD mortality					
Continuous # (n=7,210)	103 (1.4%)	0.99 [0.95 to 1.03]	1.00 [0.96 to 1.05]	0.97 [0.92 to 1.02]	0.99 [0.94 to 1.03]
Category*					
High (n=5,768)	78 (1.4%)	Reference	Reference	Reference	Reference
Low (n=1,442)	25 (1.7%)	1.23 [0.74 to 2.05]	1.06 [0.63 to 1.78]	1.48 [0.86 to 2.56]	1.23 [0.71 to 2.12]
P-trend**		0.424	0.817	0.161	0.465
Cancer mortality					
Continuous # (n=7,210)	169 (2.3%)	0.95 [0.91 to 0.98]	0.96 [0.92 to 0.99]	0.96 [0.92 to 1.01]	0.96 [0.91 to 1.00]
Category*					
High (n=5,768)	121 (2.1%)	Reference	Reference	Reference	Reference
Low (n=1,442)	48 (3.3%)	1.98 [1.37 to 2.85]	1.85 [1.28 to 2.67]	1.76 [1.17 to 2.64]	1.72 [1.13 to 2.62]
P-trend**		0.001	0.001	0.006	0.012
Other causes of mortality					
Continuous # (n=7,210)	153 (2.1%)	0.99 [0.96 to 1.02]	1.00 [0.96 to 1.03]	1.00 [0.96 to 1.03]	1.01 [0.98 to 1.06]
Category*					
High (n=5,768)	110 (1.9%)	Reference	Reference	Reference	Reference
Low (n=1,442)	43 (3.0%)	1.65 [1.12 to 2.43]	1.50 [1.02 to 2.21]	1.62 [1.08 to 2.44]	1.43 [0.93 to 2.20]
P-trend**		0.011	0.042	0.020	0.100

Abbreviations: CI – confidence interval, CQI – carbohydrate quality index, CVD = Cardiovascular Disease, PREDIMED = Prevención con Dieta Mediterránea.

#Regression Cox calculated per 1 unit increase in dietary fibre

*Cumulative average dietary fibre treated as a categorical variable by low intake (lowest quintile) vs. high intake (the four upper quintiles merged, as a reference category).

**Tests of linear trend were applied for the evaluation of dose–response relationships across quintiles, assigning to each category of the total intake its quintile-specific median and using the resulting variable as continuous.

^aMultivariable regression Cox models stratified by recruiting centre with robust standard errors to account for small deviations from individual randomization. Results are expressed as Hazard Ratios (HR) and 95% Confidence Intervals (95% CI). Cumulative average dietary fibre was

treated as a continuous or categorical variable by tertiles. Extremes of total energy intake (≥ 4000 or < 800 kcal/day in men and ≥ 3500 or in women) were excluded. Significance was set at *P* values less than 0.05 and boldfaced. The exposure as continuous is presented per 1 unit increase.

^bModel 1: adjusted for age (years), sex (male or female), intervention group (control, Mediterranean diet + extra virgin olive oil, Mediterranean diet + nuts).

^cModel 2: additionally adjusted for educational level (primary education or less, secondary education, or college/graduate), baseline physical activity (metabolic equivalent task units in min/day), smoking status (never, current, or former), cumulative average dietary alcohol intake (using the linear term and adding a quadratic term, g/day), baseline body mass index (kg/m²), prevalent type 2 diabetes (yes or no) or medication (yes or no), prevalent hypertension (yes or no) or medication (yes or no), prevalent hypercholesterolemia (yes or no) or medication (yes or no).

^dModel 3: additionally adjusted for cumulative average dietary energy intake (kcal/day).

^eModel 4: additionally adjusted for cumulative average dietary protein intake (g/day), dietary saturated fatty acids intake (g/day), dietary monounsaturated fatty acids intake (g/day), and dietary polyunsaturated fatty acids intake (g/day). In the case of CVD and cancer mortality, family history of such diseases were included as covariates.

Table S8.- Relationship between cumulative average whole-grain-to-total grain ratio and all-cause and specific-cause mortality in the PREDIMED cohort^a.

No. Deaths (%)	Models			
	Model 1 ^b	Model 2 ^c	Model 3 ^d	Model 4 ^e
All-cause mortality				

Category*						
High (n=4,138)	179 (4.3%)	Reference	Reference	Reference	Reference	
Low (n=3,072)	246 (8.0%)	1.68 [1.36 to 2.06]	1.60 [1.29 to 1.97]	1.59 [1.29 to 1.97]	1.53 [1.24 to 1.90]	
P-trend**		0.236	0.387	0.406	0.604	
CVD mortality						
Category*						
High (n=4,138)	43 (1.0%)	Reference	Reference	Reference	Reference	
Low (n=3,072)	60 (2.0%)	1.75 [1.14 to 2.68]	1.64 [1.06 to 2.54]	1.59 [1.02 to 2.47]	1.51 [0.97 to 2.36]	
P-trend**		0.858	0.915	0.710	0.666	
Cancer mortality						
Category*						
High (n=4,138)	68 (1.6%)	Reference	Reference	Reference	Reference	
Low (n=3,072)	101 (3.3%)	1.83 [1.31 to 2.55]	1.74 [1.25 to 2.43]	1.76 [1.26 to 2.45]	1.71 [1.22 to 2.40]	
P-trend**		0.242	0.352	0.283	0.377	
Other causes of mortality						
Category*						
High (n=4,138)	68 (1.6%)	Reference	Reference	Reference	Reference	
Low (n=3,072)	85 (2.8%)	1.47 [1.04 to 2.06]	1.42 [1.00 to 2.01]	1.42 [1.00 to 2.01]	1.34 [0.93 to 1.93]	
P-trend**		0.551	0.598	0.603	0.808	

Abbreviations: CI – confidence interval, CQI – carbohydrate quality index, CVD = Cardiovascular Disease, PREDIMED = Prevención con Dieta Mediterránea.

*Cumulative average whole-grain-to-total grain ratio treated as a categorical variable by low intake (lowest quintile) vs. high intake (the four upper quintiles merged, as a reference category).

**Tests of linear trend were applied for the evaluation of dose–response relationships across quintiles, assigning to each category of the total intake its quintile-specific median and using the resulting variable as continuous.

^aMultivariable regression Cox models stratified by recruiting centre with robust standard errors to account for small deviations from individual randomization. Results are expressed as Hazard Ratios (HR) and 95% Confidence Intervals (95% CI). Cumulative average whole-grain-to-total grain ratio was treated as a categorical variable by tertiles. Extremes of total energy intake (≥ 4000 or < 800 kcal/day in men and ≥ 3500 or in women) were excluded. Significance was set at *P* values less than 0.05 and boldfaced.

^bModel 1: adjusted for age (years), sex (male or female), intervention group (control, Mediterranean diet + extra virgin olive oil, Mediterranean diet + nuts).

^cModel 2: additionally adjusted for educational level (primary education or less, secondary education, or college/graduate), baseline physical activity (metabolic equivalent task units in min/day), smoking status (never, current, or former), cumulative average dietary alcohol intake (using

the linear term and adding a quadratic term, g/day), baseline body mass index (kg/m²), prevalent type 2 diabetes (yes or no) or medication (yes or no), prevalent hypertension (yes or no) or medication (yes or no), prevalent hypercholesterolemia (yes or no) or medication (yes or no).

^dModel 3: additionally adjusted for cumulative average dietary energy intake (kcal/day).

^eModel 4: additionally adjusted for cumulative average dietary protein intake (g/day), dietary saturated fatty acids intake (g/day), dietary monounsaturated fatty acids intake (g/day), and dietary polyunsaturated fatty acids intake (g/day). In the case of CVD and cancer mortality, family history of such diseases were included as covariates.

Table S9.- Relationship between cumulative average solid- to-all carbohydrates ratio and all-cause and specific-cause mortality in the PREDIMED cohort^a.

No. Deaths (%)	Models			
	Model 1 ^b	Model 2 ^c	Model 3 ^d	Model 4 ^e
All-cause mortality				
Category*				

High (n=5,745)	320 (5.6%)	Reference	Reference	Reference	Reference
Low (n=1,434)	74 (5.2%)	1.09 [0.85 to 1.40]	1.12 [0.87 to 1.44]	1.12 [0.87 to 1.44]	1.07 [0.83 to 1.38]
P-trend**		0.936	0.894	0.914	0.914
CVD mortality					
Category*					
High (n=5,745)	78 (1.4%)	Reference	Reference	Reference	Reference
Low (n=1,434)	16 (1.1%)	0.86 [0.51 to 1.44]	0.84 [0.50 to 1.40]	0.81 [0.49 to 1.34]	0.75 [0.44 to 1.28]
P-trend**		0.344	0.303	0.220	0.179
Cancer mortality					
Category*					
High (n=5,745)	120 (2.1%)	Reference	Reference	Reference	Reference
Low (n=1,434)	37 (2.6%)	1.56 [1.09 to 2.24]	1.58 [1.09 to 2.29]	1.60 [1.10 to 2.32]	1.51 [1.03 to 2.21]
P-trend**		0.055	0.051	0.039	0.065
Other causes of mortality					
Category*					
High (n=5,745)	122 (2.1%)	Reference	Reference	Reference	Reference
Low (n=1,434)	21 (1.5%)	0.79 [0.49 to 1.26]	0.87 [0.54 to 1.40]	0.87 [0.54 to 1.40]	0.84 [0.52 to 1.35]
P-trend**		0.142	0.305	0.293	0.246

Abbreviations: CI – confidence interval, CQI – carbohydrate quality index, CVD = Cardiovascular Disease, PREDIMED = Prevención con Dieta Mediterránea.

*Cumulative average solid- to all- carbohydrates ratio treated as a categorical variable by low intake (lowest quintile) vs. high intake (the four upper quintiles merged, as a reference category).

**Tests of linear trend were applied for the evaluation of dose–response relationships across quintiles, assigning to each category of the total intake its quintile-specific median and using the resulting variable as continuous.

^aMultivariable regression Cox models stratified by recruiting centre with robust standard errors to account for small deviations from individual randomization. Results are expressed as Hazard Ratios (HR) and 95% Confidence Intervals (95% CI). Cumulative average solid to all carbohydrates ratio was treated as a categorical variable by tertiles. Extremes of total energy intake (>4000 or < 800 kcal/day in men and >3500 or in women) were excluded. Significance was set at *P* values less than 0.05 and boldfaced.

^bModel 1: adjusted for age (years), sex (male or female), intervention group (control, Mediterranean diet + extra virgin olive oil, Mediterranean diet + nuts).

^cModel 2: additionally adjusted for educational level (primary education or less, secondary education, or college/graduate), baseline physical activity (metabolic equivalent task units in min/day), smoking status (never, current, or former), cumulative average dietary alcohol intake (using the linear term and adding a quadratic term, g/day), baseline body mass index (kg/m²), prevalent type 2 diabetes (yes or no) or medication (yes

or no), prevalent hypertension (yes or no) or medication (yes or no), prevalent hypercholesterolemia (yes or no) or medication (yes or no).

^dModel 3: additionally adjusted for cumulative average dietary energy intake (kcal/day).

^eModel 4: additionally adjusted for cumulative average dietary protein intake (g/day), dietary saturated fatty acids intake (g/day), dietary monounsaturated fatty acids intake (g/day), and dietary polyunsaturated fatty acids intake (g/day). In the case of CVD and cancer mortality, family history of such diseases were included as covariates.

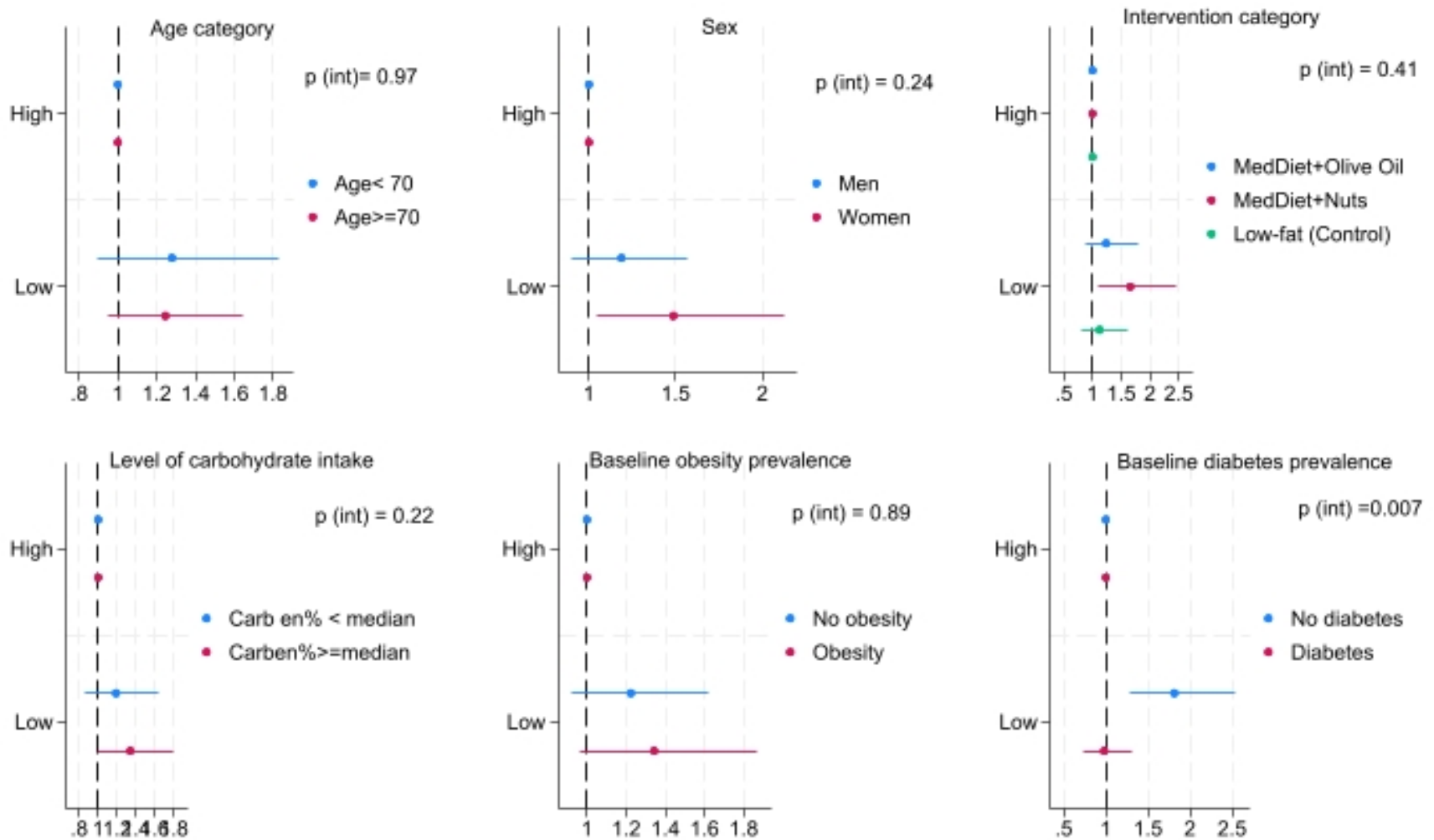


Figure S1.- Interaction for carbohydrate quality index and baseline the variables of the study.