

Supplementary material

Evaluating agreement between evidence from randomised controlled trials and cohort studies in nutrition: a meta-research replication study

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Table of contents

Supplementary Appendix S1 Changes made to the original study	4
Supplementary Appendix S2 Search strategy for systematic reviews.....	5
Supplementary Appendix S3 Criteria for Rating Population (P), Intervention/Exposure (I/E), Comparator (C), and Outcome (O) similarities	7
Supplementary Appendix S4 Reasons for exclusion of systematic reviews	9
Supplementary Table S1 Exclusion reasons for highly correlated outcomes.....	10
Supplementary Table S2 Exclusion reasons for BoE-pairs due to overlap	13
Supplementary Table S3 Minimal important differences (MID) for the included continuous outcomes.....	14
Supplementary Table S4 Description of BoE-pairs	15
Supplementary Table S5 Overview of transformations made to the original data extraction.....	23
Supplementary Table S6 Characteristics of included BoE from randomised controlled trials	36
Supplementary Table S7 Characteristics of included BoE from cohort studies.....	48
Supplementary Table S8 Methodological quality assessment of the included systematic reviews (AMSTAR 2).....	57
Supplementary Table S9 Ratings of PI/ECO similarity degree for included BoE-pairs	60
Supplementary Table S10 Analysis of concordance of the included BoE-pairs	63
Supplementary Table S11 Subgroup analysis by PI/ECO similarity degree for each domain.....	67
Supplementary Table S12 Overlap of primary studies in BoE-pairs with highly similar PI/ECO questions – comparison between the present sample and the sample in Schwingshackl 2021	68
Supplementary Fig. S1 Forest plot, analysis of BoE-pairs with continuous outcomes and standardised mean difference	71
Supplementary Fig. S2 Forest plot, subgroup analysis of BoE-pairs with binary outcomes by type of intervention/exposure	72
Supplementary Fig. S3 Forest plot, subgroup analysis of BoE-pairs with binary outcomes by type of intake/exposure.....	73
Supplementary Fig. S4 Forest plot, subgroup analysis of BoE-pairs with binary outcomes by type of outcome	74
Supplementary Fig. S5 Forest plot, subgroup analysis of BoE-pairs with binary outcomes by overall PI/ECO similarity degree	75
Supplementary Fig. S6 Forest plot, subgroup analysis of BoE-pairs with binary outcomes by population similarity degree	76
Supplementary Fig. S7 Forest plot, subgroup analysis of BoE-pairs with binary outcomes by intervention/exposure similarity degree	77
Supplementary Fig. S8 Forest plot, subgroup analysis of BoE-pairs with binary outcomes by comparator similarity degree	78
Supplementary Fig. S9 Forest plot, subgroup analysis of BoE-pairs with binary outcomes by outcome similarity degree	79

Supplementary Fig. S10 Forest plot, subgroup analysis of BoE-pairs with binary outcomes by AMSTAR 2 rating	80
Supplementary Fig. S11 Forest plot, sensitivity analysis including one BoE-pair per systematic review	81
Supplementary Fig. S12 Forest plot, sensitivity analysis by direction of cohort study summary effect estimate (cohort studies with risk ratio [RR] <1)	82
Supplementary Fig. S13 Forest plot, sensitivity analysis by direction of cohort study summary effect estimate (cohort studies with risk ratio [RR] ≥1)	83
Supplementary Fig. S14 Forest plot, sensitivity analysis for interventions with micronutrients	84
Supplementary Fig. S15 Forest plot, sensitivity analysis for cancer outcomes.....	85
Supplementary Fig. S16 Forest plot, sensitivity analysis excluding BoE-pairs with highly similar PI/ECO questions and overlapping primary studies.....	86

Supplementary Appendix S1 Changes made to the original study

	Schwingshackl 2021 [1]	present replication study
Sample	Systematic reviews of RCTs (Cochrane reviews) and systematic reviews of cohort studies of the same patient-relevant outcome or intermediate disease marker* → matching approach Exclusion of highly correlated outcomes in post hoc sensitivity analyses	Systematic reviews including both RCTs and cohort studies of the same patient-relevant outcome or intermediate disease marker* Exclusion of highly correlated outcomes Exclusion of systematic review with overlapping PI/ECO with another included systematic review
Search	database: Cochrane Database of Systematic Reviews and Medline (via OVID) filter: published between 1 January 2010 and 31 December 2019	database: Cochrane Database of Systematic Reviews, Medline (via OVID), and Epistemonikos filter: published between 1 January 2011 and 6 September 2021
Screening	Title and abstracts were screened by one reviewer, and full texts by two reviewers independently.	Title and abstracts, and full texts were screened by two reviewers independently.
Selection of outcomes	Based on the ranking in the summary of findings table in the identified Cochrane reviews (from top to bottom)	Based on the list of primary outcomes, and thereafter selection with an top down approach (highest number of studies included in BoE from RCTs; highest number of study participants; highest number of cases)
Additional analyses	N/A	AMSTAR 2 assessment Analysis of concordance

* There is an overlap of nine systematic reviews between both samples: The reviews of Aburto 2013 [2], Chowdhury 2014 [3], Feng 2015 [4], Jiang 2019 [5], Kastorini 2011 [6], Vinceti 2018 [7], Wolf 2017 [8] and Zhang 2016 [9] were also included in the review by Schwingshackl 2021 [1].

AMSTAR 2=a measurement tool to assess systematic reviews, version 2; BoE=body of evidence; PI/ECO=population, intervention/exposure, comparator, outcome; RCT=randomised controlled trial

Supplementary Appendix S2 Search strategy for systematic reviews

Medline (via Ovid), date: 06.09.2021	
ID	Query
#1	Systematic Review/ or (systematic-review or systematic literature review or systematic search).ti,ab,kf or systematic review.pt.
#2	Meta-analysis/ or (meta-analys* or meta analys* or metaanalys* or metasynthe* or meta-synthe* or meta-review or methodological review).ti,ab,kf or meta-analysis.pt.
#3	(randomi* or randomly or RCT or RCTs or (intervention* adj2 (stud* or trial*)) or (controlled adj1 (stud* or trial*))).ti,ab,kf.
#4	Cohort Studies/ or (cohort* or observational or longitudinal or prospective or case-cohort or nested case-control or (epidemiologic* adj2 stud*)).ti,ab,kf.
#5	(#1 or #2) and #3 and #4
#6	Diet/ or (mediterranean or vegetarian or low-carb* or low-fat or high-protein or gluten* or FODMAP*).ti,ab,kf.
#7	Edible Grain/ or Fruit/ or Vegetables/ or Milk/ or Dairy Products/ or Meat/ or Eggs/ or Nuts/ or Chocolate/ or Oil/ or beverages/ or (grain* or fruit* or vegetabl* or legume* or milk or dairy or soy or meat or fish or eggs or nuts or chocolate or oil* or beverage*).ti,ab,kf.
#8	Dietary Carbohydrates/ or Dietary Fats/ or Dietary Proteins/ or (carbohydrat* or fructose or glucose or starch or sucrose or sweetener* or aspartame or saccharin* or protein* or amino acid* or fatty acid* or omega or unsaturated or monounsaturated or polyunsaturated or eicosapentaenoic or docosahexaenoic or linoleic acid*).ti,ab,kf.
#9	Micronutrients/ or Trace Elements/ or Vitamins/ or (Vitamin* or multivitamin* or retinol or caroten* or thiamine or riboflavin or niacin or pantothenic or pyridoxine or biotin or folic or folate or folacin or cobalamin or ascorbic or ascorbate or calciferol* or cholecalciferol* or ergocalciferol* or tocopherole or tocotrienol or phyloquinone or menaquinone or calcium or chromium or copper or iodine or iron or fluorine or fluoride or magnesium or manganese or phosphor* or potassium or selenium or sodium or zinc).ti,ab,kf.
#10	Dietary Fiber/ or Dietary Supplements/ or (fibre* or fiber* or psyllium or inulin or cellulose or prebiotic* or probiotic* or symbiotic*).ti,ab,kf.
#11	(diet* or nutrition or eat* or feed* or food* or consum* or intake* or supplement*).ti,ab,kf.
#12	(#6 or #7 or #8 or #9 or #10) and #11
#13	#5 and #12
	limit 13 to last 10 years
	results: 1,351

Cochrane Library (via Website, advanced search), date: 06.09.2021	
ID	Query
#1	(randomi* or randomly or RCT or RCTs):ti,ab,kw
#2	(controlled NEAR/1 (stud* or trial)):ti,ab,kw
#3	(clinical NEAR/2 (stud* or trial)):ti,ab,kw
#4	#1 or #2 or #3
#5	(cohort* or observational or longitudinal or prospective or case-cohort or “nested case-control” or epidemiologic*):ti,ab,kw
#6	MeSH descriptor: [Cohort Studies] explode all trees
#7	#5 or #6
#8	#4 and #7
#9	(diet* or mediterranean or vegetarian or low-carb* or low-fat or high-protein or gluten* or FODMAP*):ti,ab,kw
#10	MeSH descriptor: [Diet] explode all trees
#11	(grain* or fruit* or vegetabl* or legume* or milk or dairy or soy or meat or fish or eggs or nuts or chocolate or oil* or beverage*):ti,ab,kw
#12	MeSH descriptor: [Food] explode all trees
#13	MeSH descriptor: [Beverages] explode all trees

#14	(carbohydrat* or fructose or glucose or starch or sucrose or sweetener* or aspartame or saccharin* or omega or unsaturated or monounsaturated or polyunsaturated or eicosapentaenoic or docosahexaenoic or "linoleic acid*" or "fatty acid*" or protein* or "amino acid*"):ti,ab,kw
#15	MeSH descriptor: [Dietary Carbohydrates] explode all trees
#16	MeSH descriptor: [Dietary Fats] explode all trees
#17	MeSH descriptor: [Fatty acids] explode all trees
#18	MeSH descriptor: [Dietary Proteins] explode all trees
#19	(Vitamin* or multivitamin* or retinol or caroten* or thiamine or riboflavin or niacin or pantothenic or pyridoxine or biotin or folic or folate or folacin or cobalamin or ascorbic or ascorbate or calciferol* or cholecalciferol* or ergocalciferol* or tocopherole or tocotrienol or phylloquinone or menaquinone or calcium or chromium or copper or iodine or iron or fluorine or fluoride or magnesium or manganese or phosphor* or potassium or selenium or sodium or zinc):ti,ab,kw
#20	MeSH descriptor: [Micronutrients] explode all trees
#21	(fibre* or fiber* or psyllium or inulin or cellulose or prebiotic* or probiotic* or symbiotic*):ti,ab,kw
#22	#9 or #10 or #11 or #12 or #13 or #14 or #15 or #16 or #17 or #18 or #19 or #20 or #21
#23	(diet* or nutrition or eat* or feed* or food* or consum* or intake* or supplement*):ti,ab,kw
#24	#22 and #23
#25	#8 and #24
	#25 in Cochrane Reviews
	with Cochrane Library publication date from Jan 2011 to Sep 2021
	results: 126

Epistemonikos (via Website, advanced search, title/abstract), 06.09.2021	
ID	Query
#1	"systematic review" OR systematic-review OR "systematic literature review" OR "systematic search"
#2	meta-analys* OR "meta analys*" OR metaanalys* OR metasynthe* OR meta-synthe* OR meta-review OR "methodological review"
#3	#1 or #2
#4	randomi* OR randomly OR RCT OR RCTs OR "intervention* studies" OR "intervention* trials" OR "controlled studies" OR "controlled trials"
#5	cohort* OR observational OR longitudinal OR prospective OR case-cohort OR "nested case-control" OR epidemiologic*
#6	#3 and #4 and #7
#7	diet* OR mediterranean OR vegetarian OR low-carb* OR low-fat or high-protein OR gluten* OR FODMAP*
#8	grain* OR fruit* OR vegetabl* OR legume* OR milk OR dairy OR soy OR meat OR fish OR eggs OR nuts OR chocolate OR oil* OR beverage*
#9	carbohydrat* OR fructose OR glucose OR starch OR sucrose OR sweetener* OR aspartame OR saccharin* OR omega OR unsaturated OR monounsaturated OR polyunsaturated OR eicosapentaenoic OR docosahexaenoic OR "linoleic acid*" OR "fatty acid*" OR protein* OR "amino acid"
#10	Vitamin* OR multivitamin* OR retinol OR caroten* OR thiamine OR riboflavin OR niacin OR pantothenic OR pyridoxine OR biotin OR folic OR folate or folacin OR cobalamin OR ascorbic OR ascorbate OR calciferol* OR cholecalciferol* OR ergocalciferol* or tocopherole OR tocotrienol OR phylloquinone OR menaquinone OR calcium OR chromium OR copper OR iodine OR iron OR fluorine OR fluoride OR magnesium OR manganese OR phosphor* OR potassium OR selenium OR sodium OR zinc
#11	fibre* OR fiber* OR psyllium OR inulin OR cellulose OR prebiotic* OR probiotic* OR symbiotic*
#12	#7 OR #8 OR #9 OR #10 OR #11
#13	diet* OR nutrition OR eat* OR feed* or food* OR consum* OR intake* OR supplement*
#14	#12 AND #13
#15	#6 AND #14
	publication year: "Last 10 years"
	results: 1,408

Supplementary Appendix S3 Criteria for Rating Population (P), Intervention/Exposure (I/E), Comparator (C), and Outcome (O) similarities
(modified from Schwingshackl 2021 [1])

Rating	Population	Intervention/ Exposure	Comparator	Outcome
1 = more or less identical	Both BoE include primary <u>or</u> secondary prevention Example: - both BoE with healthy population, general population - both BoE with diseased population (e.g. people with cardiovascular disease) Both BoE include same type of population regarding basic characteristics (age, gender) Example: - both BoE with adults - both BoE with postmenopausal women	Both BoE use intake, supplementation, <u>or</u> status Example: - folate supplementation in BoE from RCTs and BoE from cohort studies - apple intake in BoE from RCTs and BoE from cohort studies	Both BoE use nil/ low intake, supplementation/ placebo, <u>or</u> status. Example: - no/ low intake from BoE from RCTs and BoE from cohort studies	Both BoE examine the same outcome. Example: - all-cause mortality in BoE from RCTs and BoE from cohort studies
2 = similar but not identical	Both BoE include primary and secondary prevention <u>or</u> mixed population vs. primary or secondary prevention <u>or</u> high-risk population vs. primary or secondary prevention Example: - mixed population (healthy people and partly people with cardiovascular disease) in BoE from RCTs vs. general healthy population in BoE from cohort studies - population at high-risk in BoE from RCTs vs. general healthy population in BoE from cohort studies	Both BoE use similar intervention/ exposure: - similar products with different routes of administration - different products of the same class - similar products with different co-interventions - similar time frame of intervention Example: - intake vs. supplementation; intake vs. urinary excretion; intake + supplementation vs. intake - supplementation vs. supplementation of similar products (multi-micronutrient supplementation vs. multivitamin supplementation) - intake of product + medicament (e.g. aspirin) vs. intake of product	BoE use similar but not identical comparator. Example: see Intervention/ Exposure	Both BoE examine the same cluster of outcomes. Example: - cardiovascular disease in BoE from RCTs and coronary heart disease in BoE from cohort studies - cardiovascular mortality in BoE from RCTs vs. cardiovascular incidence + mortality in BoE from cohort studies

3 = broadly similar	One BoE include primary prevention and the other BoE (mostly) secondary prevention Example: - people with cancer in BoE from RCTs vs. general healthy population in BoE from cohort studies - mixed population (most $\geq 66\%$) of the people with cardiovascular disease) in BoE from RCTs vs. general population in BoE from cohort studies	BoE use broadly similar intervention/ exposure: - intake or supplementation vs. status - broadly similar products - different time frame/ early treatment vs. any treatment Example: - selenium supplementation vs. high selenium status - Vitamin A supplementation vs. β-carotene intake	BoE use broadly similar comparator. Example: see Intervention/ Exposure	BoE examine similar outcomes. Example: - colorectal adenoma in BoE from RCTs vs. colorectal cancer in BoE from cohort studies
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BoE=bodies of evidence; RCT=randomised controlled trial

Supplementary Appendix S4 Reasons for exclusion of systematic reviews

References	Reason for exclusion
[10-98]	Ineligible study design
[99-130]	Ineligible PI/ECO
[131-151]	No RCTs or cohort studies included or RCTs analysed as cohort studies
[152-187]	Intervention/exposure or outcome differ
[188-203]	Ineligible effect size
[204-206]	Updated version included
[207-217]	Overlapping PI/ECO with another included systematic review

PI/ECO=population, intervention/exposure, comparator, outcome; RCT=randomised controlled trial

Supplementary Table S1 Exclusion reasons for highly correlated outcomes

Reference: Author, Year	Intervention	Outcome	Reason for exclusion
Aburto 2013 [2]	Sodium	Stroke	Highly likely correlated with outcome “Cardiovascular disease”
Aburto 2013 [2]	Sodium	Coronary heart disease	Highly likely correlated with outcome “Cardiovascular disease”
Aguilar-Cordero 2020 [218]	Vitamin D (maternal)	Prematurity	Highly likely correlated with outcome “Pre-eclampsia”
Alexander 2017 [219]	Omega-3 fatty acids	Any fatal coronary heart disease event	Highly likely correlated with outcome “Coronary heart disease”
Alexander 2017 [219]	Omega-3 fatty acids	Any nonfatal coronary heart disease event	Highly likely correlated with outcome “Coronary heart disease”
Alexander 2017 [219]	Omega-3 fatty acids	Coronary death	Highly likely correlated with outcome “Coronary heart disease”
Azad 2017 [220]	Nonnutritive sweeteners	Weight (change)	Highly likely correlated with outcome “Body mass index”
Azad 2017 [220]	Nonnutritive sweeteners	Waist circumference	Highly likely correlated with outcome “Body mass index”
Bolland 2015 [221]	Calcium	Hip fractures	Highly likely correlated with outcome “All fractures”
Bolland 2015 [221]	Calcium	Vertebral fractures	Highly likely correlated with outcome “All fractures”
Bolland 2015 [221]	Calcium	Forearm fractures	Highly likely correlated with outcome “All fractures”
Grosso 2015 [222]	Mediterranean diet	Myocardial infarction	Highly likely correlated with outcome “Cardiovascular disease”
Grosso 2015 [222]	Mediterranean diet	Stroke	Highly likely correlated with outcome “Cardiovascular disease”
Grosso 2015 [222]	Mediterranean diet	Cardiovascular disease mortality	Highly likely correlated with outcome “Cardiovascular disease”
Kim 2018 [223]	Multivitamins/ Minerals	Coronary heart disease mortality	Highly likely correlated with outcome “Cardiovascular disease mortality”

Kim 2018 [223]	Multivitamins/ Minerals	Stroke mortality	Highly likely correlated with outcome “Cardiovascular disease mortality”
Stratton 2011 [224]	Vitamin E	Metastatic prostate cancer/ death from prostate cancer	Highly likely correlated with outcome “Prostate cancer”
Stratton 2011 [224]	β -carotene	Metastatic prostate cancer/ death from prostate cancer	Highly likely correlated with outcome “Prostate cancer”
Thorne-Lyman 2012 [225]	Vitamin D	Mean gestational age	Highly likely correlated with outcome “Mean birth weight”
Vinceti 2018a [7]	Selenium	Colorectal cancer	Highly likely correlated with outcome “Any cancer”
Vinceti 2018a [7]	Selenium	Lung cancer	Highly likely correlated with outcome “Any cancer”
Vinceti 2018a [7]	Selenium	Breast cancer	Highly likely correlated with outcome “Any cancer”
Vinceti 2018a [7]	Selenium	Bladder cancer	Highly likely correlated with outcome “Any cancer”
Vinceti 2018a [7]	Selenium	Prostate cancer	Highly likely correlated with outcome “Any cancer”
Wien 2012 [226]	Folic acid	Pancreatic cancer	Highly likely correlated with outcome “Any cancer”
Wien 2012 [226]	Folic acid	Breast cancer	Highly likely correlated with outcome “Any cancer”
Wien 2012 [226]	Folic acid	Lung cancer	Highly likely correlated with outcome “All cancer”
Wolf 2017 [8]	Multivitamins	Low birthweight	Highly likely correlated with outcome “Preterm birth”
Wolf 2017 [8]	Multivitamins	Small for gestational age	Highly likely correlated with outcome “Preterm birth”
Wolf 2017 [8]	Multivitamins	Neural tube defect	Highly likely correlated with outcome “Stillbirth”
Yang 2020 [227]	Calcium	Coronary heart disease	Highly likely correlated with outcome “Cardiovascular disease”
Yang 2020 [227]	Calcium	Stroke	Highly likely correlated with outcome “Cardiovascular disease”

Yao 2019 [228]	Vitamin D	Hip fracture	Highly likely correlated with outcome “Any fracture”
Yu 2021 [229]	Folic acid	Gestational hypertension	Highly likely correlated with outcome “Pre-eclampsia”
Zhang 2016 [9]	Selenium	Coronary heart disease	Highly likely correlated with outcome “Cardiovascular disease”
Zhang 2016 [9]	Selenium	Myocardial infarction	Highly likely correlated with outcome “Cardiovascular disease”
Zhang 2018 [230]	Vitamin D	Fasting insulin	Highly likely correlated with outcome “Fasting plasma glucose”
Zhang 2018 [230]	Vitamin D	HOMA-IR	Highly likely correlated with outcome “Fasting plasma glucose”
Zhang 2018 [230]	Vitamin D	HbA1c	Highly likely correlated with outcome “Fasting plasma glucose”

HbA1c=Haemoglobin A1c; HOMA-IR=Homeostasis model assessment-estimated insulin resistance

Supplementary Table S2 Exclusion reasons for BoE-pairs due to overlap

Reference: Author, Year	Intervention/Exposure	Outcome	Reason
Amegah 2017 [231]	Vitamin D	Preterm birth	Most of the included studies (5/7) are also identified by Zhou 2017 [232].
Becerra-Tomas 2020 [207]	Mediterranean diet	Cardiovascular disease mortality	Some of the included studies (4/18) are identified by Grosso 2015 [222], which also used a larger sample of studies for BoE from RCTs (n=4) and BoE from cohort studies (n=20).
Bloomfield 2016 [208]	Mediterranean diet	Breast cancer	Some of the included studies (4/14) are identified by Morze 2021 [233], which also used the corrected publication of the PREDIMED for BoE from RCTs.
Chowdhury 2014a [234]	Omega-3 fatty acids	Coronary disease	Many of the included studies (14/25) are also identified by Alexander 2017 [219], which also used a larger sample of studies for BoE from RCTs (n=18) and BoE from cohort studies (n=17).
Chung 2016 [209]	Calcium	Cardiovascular disease mortality	Some of the included studies (2/6) are also identified by Yang 2020 [227], which also used a larger sample of studies for BoE from RCTs (n=4) and BoE from cohort studies (n=13).
Hua 2016 [210]	Folic acid	Gestational hypertension and pre-eclampsia	All of the included studies (6/6) are identified by Yu 2021 [229] or Yang 2016 [235].
Hyppönen 2013 [211]	Vitamin D	Pre-eclampsia	All of the included studies (6/6) are identified by Aguilar-Cordero 2020 [218].
Kohler 2018 [212]	Selenium	Type 2 diabetes	All of the included studies (4/4) are identified by Vinceti 2018b [236].
Koloverou 2014 [213]	Mediterranean diet	Type 2 diabetes	Most of the included studies (9/10) are identified by Schwingshackl 2015 [237], which also used a larger sample of the PREDIMED study for BoE from RCTs [238].
Kuria 2020 [214]	Selenium	Cardiovascular disease	All of the included studies (7/7) are also identified by Zhang 2016 [9].
Liu 2018 [215]	Folic acid	Pre-eclampsia	All of the included studies (10/10) are identified by Yu 2021 [229].
Martinez-Gonzalez 2014a [216]	Mediterranean diet	Cardiovascular disease	All of the included studies (14/14) are also identified by Grosso 2015 [222].
Shim 2016 [217]	Folic acid	Gestational hypertension	All of the included studies (2/2) are also identified by Yang 2016 [235].
Trikalinos 2012 [239]	Omega-3 fatty acids	Cardiovascular mortality	Most of the included studies (11/16) are identified by Alexander 2017 [219], which also used a larger sample of studies for BoE from RCTs (n=18) and BoE from cohort studies (n=17).

BoE=bodies of evidence; PREDIMED=Prevención con Dieta Mediterránea; RCT=randomised controlled trial

Supplementary Table S3 Minimal important differences (MID) for the included continuous outcomes

Outcome	Normal value	MID	Lower 95% CI	Upper 95% CI
Body weight	75 kg	2.5%	-1.9	1.9
Body mass index	24.4	2.5%	-1.2	1.2
Birth weight	3,400 g	2.5%	-85	85
Gestational length	40 weeks	2.5%	-1	1
HDL-cholesterol	1.036 mmol/l	5%	-0.05	0.05
Mini Mental State Examination Score	>26	5%	-1.3	1.3
Systolic blood pressure	120 mmHg	2.5%	-3	3
Triglyceride	1.7 mmol/l	2.5%	-0.04	0.04

CI=confidence interval; HDL=high density lipoprotein; MID=minimal important difference

Supplementary Table S4 Description of BoE-pairs

	BoE from randomised controlled trials					BoE from cohort studies					
Reference	Intervention and type of intake	Comparison	Outcome	Studies (n)	Summary measure, and effect estimate (95% CI)	Exposure and type of intake	Comparison	Outcome	Studies (n)	Summary measure, and effect estimate (95% CI)	Included in the meta-analysis (Yes/No)
Aburto 2013 [2]	Sodium/ Intake	Low vs. High	All-cause mortality	4	RR 0.70 (0.44 to 1.14)	Sodium/ Intake	High vs. Low	All-cause mortality	2	RR 1.06 (0.94 to 1.20)	Yes
Aburto 2013 [2]	Sodium/ Intake	Low vs. High	Cardiovascular disease	2	RR 0.84 (0.57 to 1.23)	Sodium/ Intake	High vs. Low	Cardiovascular disease	3	RR 1.12 (0.93 to 1.34)	Yes
Afshin 2014 [240]	Nuts/ Intake	Dose-response; per 4 weekly servings (28.4g) of nuts	Coronary heart disease	1	RR 0.79 (0.55 to 1.14)	Nuts/ Intake	Dose-response; per 4 weekly servings (28.4g) of nuts	Coronary heart disease	6	RR 0.76 (0.69 to 0.83)	Yes
Afshin 2014 [240]	Nuts/ Intake	Dose-response; per 4 weekly servings (28.4g) of nuts	Stroke	1	RR 0.62 (0.44 to 0.87)	Nuts/ Intake	Dose-response; per 4 weekly servings (28.4g) of nuts	Stroke	2	RR 0.95 (0.83 to 1.09)	Yes
Afshin 2014 [240]	Nuts/ Intake	Dose-response; per 4 weekly servings (28.4g) of nuts	Type 2 diabetes	1	RR 0.86 (0.68 to 1.08)	Nuts/ Intake	Dose-response; per 4 weekly servings (28.4g) of nuts	Type 2 diabetes	4	RR 0.87 (0.80 to 0.95)	Yes
Aguilar-Cordero 2020 [218]	Vitamin D/ Supplementation	High vs. Low	Pre-eclampsia	7	OR 0.61 (0.36 to 1.04)	Vitamin D/ Status	Low vs. High	Pre-eclampsia	7	OR 1.25 (0.89 to 1.75)	Yes
Alexander 2017 [219]	Omega-3 fatty acids/ Intake + Supplementation	High vs. Low	Coronary heart disease	18	RR 0.94 (0.85 to 1.05)	Omega-3 fatty acids/ Intake	High vs. Low	Coronary heart disease	17	RR 0.82 (0.74 to 0.92)	Yes

Amegah 2017 [231]	Vitamin D/ Supplementation	High vs. Low	Gestational length	2	MD 0.16 (0.28 to 0.59)	Vitamin D/ Status	Low vs. High	Gestational length	1	MD 0.80 (0.20 to 1.40)	Yes
Azad 2017 [220]	Nonnutritive sweeteners/ Intake + Supplementation	High vs. Low	Body mass index	3	MD -0.37 (-1.10 to 0.36)	Nonnutritive sweeteners/ Intake	High vs. Low	Body mass index	1	MD 0.77 (0.47 to 1.07)	Yes
Bolland 2015 [221]	Calcium/ Supplementation	High vs. Low	All fractures	20	RR 0.89 (0.81 to 0.96)	Calcium/ Supplementation	High vs. Low	All fractures	5	RR 1.02 (0.93 to 1.11)	Yes
Chowdhury 2012 [241]	Omega-3 fatty acids/ Supplementation	High vs. Low	Cerebrovascular disease	2	RR 0.98 (0.89 to 1.08)	Omega-3 fatty acids/ Intake	High vs. Low	Cerebrovascular disease	10	RR 0.90 (0.80 to 1.01)	Yes
Chowdhury 2014a [234]	α -linolenic acid/ Supplementation	High vs. Low	Coronary disease	4	RR 0.97 (0.69 to 1.36)	α -linolenic acid/ Intake	High vs. Low	Coronary disease	7	RR 0.99 (0.86 to 1.14)	Yes
Chowdhury 2014a [234]	Omega-6 fatty acids/ Intake + Supplementation	High vs. Low	Coronary disease	8	RR 0.86 (0.69 to 1.07)	Omega-6 fatty acids/ Intake	High vs. Low	Coronary disease	8	RR 0.98 (0.90 to 1.06)	Yes
Chowdhury 2014b [3]	Vitamin D/ Supplementation	High vs. Low	All-cause mortality	22	RR 0.98 (0.94 to 1.03)	Vitamin D/ Status	Low vs. High	All-cause mortality	68	RR 1.44 (1.34 to 1.55)	Yes
Chung 2011 [242]	Vitamin D/ Supplementation	High vs. Low	Colorectal cancer	1	RR 1.02 (0.60 to 1.74)	Vitamin D/ Status	Dose-response; per 10nmol/l increase	Colorectal cancer	9	OR 0.94 (0.91 to 0.97)	Yes
Chung 2011 [242]	Vitamin D/ Supplementation	High vs. Low	Breast cancer	1	RR 0.99 (0.25 to 4.00)	Vitamin D/ Status	Dose-response; per 10nmol/l increase	Breast cancer	4	OR 0.99 (0.97 to 1.01)	Yes
Ding 2017 [243]	Dairy/ Intake	High vs. Low	Systolic blood pressure	8	MD -0.21 (-0.98 to 0.57)	Dairy/ Intake	Dose-response; per serving increase	Systolic blood pressure	27	MD -0.11 (-0.20 to -0.02)	Yes
Feng 2015 [4]	Folic acid/ Supplementation	High vs. Low	Congenital heart defects	1	OR 0.42 (0.19 to 0.98)	Folic acid/ Supplementation	High vs. Low	Congenital heart defects	1	OR 0.60 (0.38 to 0.96)	Yes

Filippini 2020 [244]	Green Tea/ Supplementation	High vs. Low	Prostate cancer	3	RR 0.50 (0.18 to 1.36)	Green Tea/ Intake	High vs. Low	Prostate cancer	5	RR 1.09 (0.89 to 1.32)	Yes
Filippini 2020 [244]	Green Tea/ Supplementation	High vs. Low	Endometrial cancer	1	RR 0.33 (0.01 to 8.15)	Green Tea/ Intake	High vs. Low	Endometrial cancer	1	RR 0.75 (0.43 to 1.30)	Yes
Fu 2021 [245]	Vitamin C/ Supplementation	High vs. Low	Lung cancer	2	RR 1.30 (0.68 to 2.48)	Vitamin C/ Supplementation	High vs. Low	Lung cancer	3	RR 1.02 (0.85 to 1.23)	Yes
Fu 2021 [245]	Vitamin C/ Supplementation	High vs. Low	Breast cancer	1	RR 1.11 (0.87 to 1.41)	Vitamin C/ Supplementation	High vs. Low	Breast cancer	6	RR 1.02 (0.88 to 1.18)	Yes
Fu 2021 [245]	Vitamin C/ Supplementation	High vs. Low	Colorectal cancer	2	RR 0.84 (0.64 to 1.10)	Vitamin C/ Supplementation	High vs. Low	Colorectal cancer	2	RR 0.80 (0.42 to 1.52)	Yes
Gayer 2019 [246]	Apples/ Intake	High vs. Low	Body weight	5	MD 1.14 (-0.45 to 0.73)	Apples/ Intake	Dose-response; per serving/day	Body weight	1	MD 0.56 (0.39 to 0.73)	Yes
Grosso 2015 [222]	Mediterranean diet/ Intake	High vs. Low adherence	Cardiovascular disease	4	RR 0.55 (0.39 to 0.76)	Mediterranean diet/ Intake	High vs. Low adherence	Cardiovascular disease	20	RR 0.71 (0.65 to 0.78)	Yes
Jiang 2019 [5]	Vitamin E/ Supplementation	High vs. Low	Age-related cataract	6	RR 0.97 (0.91 to 1.03)	Vitamin E/ Intake	High vs. Low	Age-related cataract	6	RR 0.90 (0.80 to 1.00)	Yes
Jiang 2019 [5]	β-carotene/ Supplementation	High vs. Low	Age-related cataract	4	RR 0.99 (0.92 to 1.07)	β-carotene/ Intake	High vs. Low	Age-related cataract	7	RR 0.90 (0.83 to 0.99)	Yes
Johnston 2019 [247]	Red meat/ Intake	Low vs. High	Cardiovascular disease	1	HR 0.98 (0.91 to 1.06)	Red meat/ Intake	Low vs. High	Cardiovascular disease	25	RR 0.86 (0.79 to 0.94)	Yes
Johnston 2019 [247]	Red meat/ Intake	Low vs. High	Type 2 diabetes	1	HR 0.96 (0.90 to 1.03)	Red meat/ Intake	Low vs. High	Type 2 diabetes	14	RR 0.76 (0.68 to 0.86)	Yes
Johnston 2019 [247]	Red meat/ Intake	Low vs. High	Colorectal cancer	1	HR 1.04 (0.90 to 1.20)	Red meat/ Intake	Low vs. High	Colorectal cancer	16	RR 0.94 (0.85 to 1.05)	Yes

Jonker 2020 [248]	Folic acid/Supplementation	High vs. Low	Mean birthweight	3	SMD 0.56 (0.15 to 0.97)	Folic acid/Supplementation	High vs. Low	Mean birthweight	2	SMD 0.33 (0.19 to 0.47)	Yes
Jonker 2020 [248]	Folic acid/Supplementation	High vs. Low	Low birthweight	1	OR 0.93 (0.81 to 1.08)	Folic acid/Supplementation	High vs. Low	Low birthweight	5	OR 0.58 (0.46 to 0.73)	Yes
Jonker 2020 [248]	Folic acid/Supplementation	High vs. Low	Small for gestational age	1	OR 0.98 (0.92 to 1.06)	Folic acid/Supplementation + Intake	High vs. Low	Small for gestational age	4	OR 0.62 (0.29 to 1.31)	Yes
Kastorini 2011 [6]	Mediterranean diet/Intake	High vs. Low adherence	Metabolic syndrome	2	logHR -0.80 (-1.42 to -0.18)	Mediterranean diet/Intake	High vs. Low adherence	Metabolic syndrome	2	logHR -0.76 (-2.16 to 0.63)	Yes
Kastorini 2011 [6]	Mediterranean diet/Intake	High vs. Low adherence	HDL-cholesterol	27	MD 1.25 (0.52 to 1.99)	Mediterranean diet/Intake	High vs. Low adherence	HDL-cholesterol	1	MD 0.30 (-1.77 to 2.37)	Yes
Kastorini 2011 [6]	Mediterranean diet/Intake	High vs. Low adherence	Triglycerides	27	MD -6.45 (-10.91 to -1.99)	Mediterranean diet/Intake	High vs. Low adherence	Triglycerides	1	MD -2.00 (-6.74 to 2.74)	Yes
Kastorini 2011 [6]	Mediterranean diet/Intake	High vs. Low adherence	Systolic blood pressure	14	MD -2.35 (-3.51 to -1.18)	Mediterranean diet/Intake	High vs. Low adherence	Systolic blood pressure	1	MD 0.80 (-0.84 to 2.44)	Yes
Kim 2018 [223]	Multivitamins/Minerals/Supplementation	High vs. Low	Cardiovascular disease	1	RR 0.95 (0.83 to 1.09)	Multivitamins/Minerals/Supplementation	High vs. Low	Cardiovascular disease	9	RR 1.00 (0.96 to 1.05)	Yes
Kim 2018 [223]	Multivitamins/Minerals/Supplementation	High vs. Low	Coronary heart disease	2	RR 0.97 (0.80 to 1.19)	Multivitamins/Minerals/Supplementation	High vs. Low	Coronary heart disease	6	RR 0.90 (0.85 to 0.96)	Yes
Kim 2018 [223]	Multivitamins/Minerals/Supplementation	High vs. Low	Stroke	1	RR 1.05 (0.70 to 1.56)	Multivitamins/Minerals/Supplementation	High vs. Low	Stroke	3	RR 0.98 (0.91 to 1.04)	Yes
Kong 2014 [249]	Vitamins/Supplementation	High vs. Low	Gastric cancer	11	RR 1.11 (0.94 to 1.31)	Vitamins/Intake	High vs. Low	Gastric cancer	7	RR 0.85 (0.66 to 1.08)	Yes
Lin 2019 [250]	Vitamin E/Supplementation	High vs. Low	Bladder cancer	3	RR 1.10 (0.86 to 1.35)	Vitamin E/Intake	High vs. Low	Bladder cancer	8	RR 0.84 (0.72 to 0.96)	Yes

Lin 2020 [251]	Vitamin D/ Supplementation	High vs. Low	Nephrolithiasis	7	RR 0.84 (0.65 to 1.10)	Vitamin D/ Supplementation	High vs. Low	Nephrolithiasis	3	RR 1.24 (1.01 to 1.51)	Yes
Lin 2020 [251]	Calcium/ Supplementation	High vs. Low	Nephrolithiasis	4	RR 1.20 (0.65 to 2.21)	Calcium/ Supplementation	High vs. Low	Nephrolithiasis	2	RR 1.16 (1.00 to 1.35)	Yes
Martinez-Gonzalez 2014b [252]	Olive oil/ Supplementation + Intake	Dose-response; per 25g/day increase	Coronary heart disease	1	RR 0.81 (0.53 to 1.24)	Olive oil/ Intake	Dose-response; per 25g/day increase	Coronary heart disease	3	RR 0.96 (0.78 to 1.18)	Yes
Martinez-Gonzalez 2014b [252]	Olive oil/ Supplementation + Intake	Dose-response; per 25g/day increase	Stroke	1	RR 0.69 (0.48 to 0.98)	Olive oil/ Intake	Dose-response; per 25g/day increase	Stroke	2	RR 0.74 (0.60 to 0.92)	Yes
Miller 2014 [253]	Low-calorie sweeteners/ Intake	High vs. Low	Body weight	13	WGMD -0.80 (-1.17 to -0.43)	Low-calorie sweeteners/ Intake	Mean correlation	Body weight	4	WGMC 0.02 (-0.01 to 0.06)	No
Moazzen 2018 [254]	Folic acid/ Supplementation	High vs. Low	Colorectal cancer	4	RR 1.07 (0.86 to 1.43)	Folic acid/ Supplementation	High vs. Low	Colorectal cancer	3	RR 0.96 (0.76 to 1.21)	Yes
Mocellin 2017 [255]	Vitamin B6/ Supplementation	High vs. Low	Cancer	9	RR 1.03 (0.94 to 1.13)	Vitamin B6/ Intake	High vs. Low	Cancer	29	RR 0.95 (0.90 to 1.01)	Yes
Morze 2021 [233]	Mediterranean diet/ Intake	High vs. Low adherence	Cancer mortality	1	RR 0.75 (0.17 to 3.33)	Mediterranean diet/ Intake	High vs. Low adherence	Cancer mortality	18	RR 0.87 (0.82 to 0.92)	Yes
Morze 2021 [233]	Mediterranean diet/ Intake	High vs. Low adherence	Breast cancer	1	RR 0.41 (0.19 to 0.87)	Mediterranean diet/ Intake	High vs. Low adherence	Breast cancer	12	RR 0.97 (0.94 to 1.00)	Yes
Picasso 2019 [256]	Vegetarian diet/ Intake	Vegetarian vs. Omnivore diet	Triglycerides	5	MD 0.11 (-0.10 to 0.31)	Vegetarian diet/ Intake	Vegetarian vs. Omnivore diet	Triglycerides	1	MD 0.07 (-0.39 to 0.53)	Yes
Picasso 2019 [256]	Vegetarian diet/ Intake	Vegetarian vs. Omnivore diet	HDL-cholesterol	4	MD -0.06 (-0.17 to 0.04)	Vegetarian diet/ Intake	Vegetarian vs. Omnivore diet	HDL-cholesterol	1	MD -0.03 (-0.10 to 0.04)	Yes

Sayehmiri 2018 [257]	Selenium/Supplementation	High vs. Low	Prostate cancer	9	RR 0.90 (0.74 to 1.09)	Selenium/Supplementation + Intake	High vs. Low	Prostate cancer	2	RR 0.89 (0.64 to 1.23)	Yes
Schwingshackl 2015 [237]	Mediterranean diet/Intake	High vs. Low adherence	Type 2 diabetes	1	RR 0.70 (0.54 to 0.91)	Mediterranean diet/Intake	High vs. Low adherence	Type 2 diabetes	8	RR 0.83 (0.74 to 0.92)	Yes
Schwingshackl 2017 [258]	Olive oil/Intake	High vs. Low	Type 2 diabetes	1	RR 0.60 (0.43 to 0.84)	Olive oil/Intake	High vs. Low	Type 2 diabetes	4	RR 0.87 (0.80 to 0.94)	Yes
Setien-Suero 2016 [259]	Homocysteine/Supplementation	Low vs. High	Mini Mental State Examination Score	2	SMD 0.05 (-0.07 to 0.17)	Homocysteine/Status	Correlation	Mini Mental State Examination Score	3	β -0.11 (-0.18 to -0.05)	No
Stratton 2011 [224]	Multivitamins/Supplementation	High vs. Low	Prostate cancer	1	OR 0.90 (0.61 to 1.33)	Multivitamins/Supplementation	High vs. Low	Prostate cancer	1	OR 1.05 (0.94 to 1.16)	Yes
Stratton 2011 [224]	Vitamin E/Supplementation	High vs. Low	Prostate cancer	3	NR	Vitamin E/Supplementation	High vs. Low	Prostate cancer	5	NR	Yes
Stratton 2011 [224]	Vitamin C/Supplementation	High vs. Low	Prostate cancer	1	HR 1.01 (0.92 to 1.10)	Vitamin C/Supplementation	High vs. Low	Prostate cancer	1	NR	Yes
Stratton 2011 [224]	β -carotene/Supplementation	High vs. Low	Prostate cancer	1	OR 1.20 (0.86 to 1.66)	β -carotene/Supplementation	High vs. Low	Prostate cancer	1	OR 0.69 (0.62 to 0.77)	Yes
Te Morenga 2013 [260]	Dietary sugars/Intake	High vs. Low	Weight gain	10	MD 0.75 (0.30 to 1.19)	Dietary sugars/Intake	High vs. Low	Weight gain	4	MD 0.31 (-0.07 to 0.68)	Yes
Te Morenga 2013 [260]	Dietary sugars/Intake	Low vs. High	Body mass index (change)	3	SMD 0.09 (-0.14 to 0.32)	Dietary sugars/Intake	Dose-response; per serving/day or increase of 100g/day SSB intake	Body mass index (change)	4	MD 0.02 (0.00 to 0.05)	Yes
Thorne-Lyman 2012 [225]	Vitamin D/Supplementation	High vs. Low	Mean birthweight	5	MD 58.07 (-63.87 to 180.00)	Vitamin D/Supplementation + Intake	High vs. Low	Mean birthweight	2	MD 52.66 (3.54 to 101.77)	Yes

Trikalinos 2012 [239]	Vitamin E/ Supplementation	High vs. Low	Cardiovascular disease	14	RR 0.97 (0.91 to 1.03)	Vitamin E/ Supplementation + Intake	High vs. Low	Cardiovascular disease	8	HR 0.85 (0.78 to 0.93)	Yes
Vinceti 2018a [7]	Selenium/ Supplementation	High vs. Low	Any cancer	5	RR 0.99 (0.86 to 1.14)	Selenium/ Status	High vs. Low	Any cancer	7	OR 0.72 (0.55 to 0.93)	Yes
Vinceti 2018a [7]	Selenium/ Supplementation	High vs. Low	Cancer mortality	2	RR 0.81 (0.49 to 1.32)	Selenium/ Intake	High vs. Low	Cancer mortality	1	OR 0.93 (0.83 to 1.04)	Yes
Vinceti 2018b [236]	Selenium/ Supplementation	High vs. Low	Type 2 diabetes	5	RR 1.11 (1.01 to 1.22)	Selenium/ Intake	High vs. Low	Type 2 diabetes	1	OR 2.39 (1.32 to 4.32)	Yes
Wien 2012 [226]	Folic acid/ Supplementation	High vs. Low	Cancer	10	RR 1.07 (1.00 to 1.14)	Folic acid/ Supplementation	High vs. Low	Cancer	5	RR 1.03 (0.90 to 1.18)	Yes
Wolf 2017 [8]	Multivitamins/ Supplementation	High vs. Low	Preterm birth	1	RR 1.09 (0.43 to 2.77)	Multivitamins/ Supplementation	High vs. Low	Preterm birth	4	RR 0.84 (0.69 to 1.03)	Yes
Wolf 2017 [8]	Multivitamins/ Supplementation	High vs. Low	Stillbirth	1	RR 2.43 (0.12 to 50.05)	Multivitamins/ Supplementation	High vs. Low	Stillbirth	2	RR 0.78 (0.59 to 1.03)	Yes
Yang 2016 [235]	Folic acid/ Supplementation	High vs. Low	Gestational hypertension	1	RR 0.62 (0.40 to 0.94)	Folic acid/ Supplementation + Intake	High vs. Low	Gestational hypertension	3	RR 1.02 (0.95 to 1.09)	Yes
Yang 2020 [227]	Calcium/ Supplementation	High vs. Low	Cardiovascular disease	4	RR 0.91 (0.56 to 1.46)	Calcium/ Intake	High vs. Low	Cardiovascular disease	13	RR 0.98 (0.90 to 1.06)	Yes
Yao 2019 [228]	Vitamin D/ Supplementation	High vs. Low	Any fracture	11	Rate Ratio 1.06 (0.98 to 1.14)	Vitamin D/ Status	Dose-response; per 25nmol/l increase in 25(OH)D level in blood	Any fracture	10	Rate Ratio 0.93 (0.89 to 0.96)	Yes
Yu 2021 [229]	Folic acid/ Supplementation	High vs. Low	Pre-eclampsia	3	RR 1.11 (0.75 to 1.62)	Folic acid/ Supplementation + Intake	High vs. Low	Pre-eclampsia	13	RR 0.85 (0.76 to 0.94)	Yes

Zhang 2016 [9]	Selenium/Supplementation	High vs. Low	Cardiovascular disease	9	RR 0.91 (0.74 to 1.10)	Selenium/Status	High vs. Low	Cardiovascular disease	14	RR 0.87 (0.76 to 0.99)	Yes
Zhang 2018 [230]	Vitamin D/Supplementation	High vs. Low	Gestational diabetes	4	RR 0.72 (0.39 to 1.31)	Vitamin D/Status	Low vs. High	Gestational diabetes	24	OR 1.82 (1.36 to 2.44)	Yes
Zhang 2018 [230]	Vitamin D/Supplementation	High vs. Low	Fasting plasma glucose	16	SMD -0.43 (-0.67 to -0.18)	Vitamin D/Status	Correlation	Fasting plasma glucose	13	CC -0.09 (-0.20 to 0.01)	No
Zhao 2014 [261]	Multivitamins/Minerals/Supplementation	High vs. Low	Nuclear cataract	2	RR 0.73 (0.59 to 0.90)	Multivitamins/Minerals/Supplementation	High vs. Low	Nuclear cataract	5	RR 0.73 (0.64 to 0.82)	Yes
Zhao 2014 [261]	Multivitamins/Minerals/Supplementation	High vs. Low	Cortical cataract	2	RR 0.92 (0.69 to 1.23)	Multivitamins/Minerals/Supplementation	High vs. Low	Cortical cataract	4	RR 0.81 (0.68 to 0.94)	Yes
Zhao 2014 [261]	Multivitamins/Minerals/Supplementation	High vs. Low	Posterior subcapsular cataract	2	RR 1.81 (1.30 to 2.54)	Multivitamins/Minerals/Supplementation	High vs. Low	Posterior subcapsular cataract	3	RR 0.96 (0.72 to 1.20)	Yes
Zhou 2017 [232]	Vitamin D/Supplementation	High vs. Low	Preterm birth	5	OR 0.57 (0.36 to 0.91)	Vitamin D/Status	Low vs. High	Preterm birth	12	OR 1.14 (0.95 to 1.38)	Yes

BoE=bodies of evidence; CC=correlation coefficient; CI=confidence interval; HR=hazard ratio; MD=mean difference; NR=not reported; OR=odds ratio; RR=risk ratio; SMD=standardised mean difference; SSB=sugar sweetened beverages; WGMC=weighted group mean correlation; WGMD=weighted group mean difference

Supplementary Table S5 Overview of transformations made to the original data extraction

Systematic review	Intervention/ Exposure	Outcome	Type of BoE (RCTs/ CSs)	n (studies)	Original		What we used		Rationale
					HR	95% CI	RR	95% CI	
Johnston 2019 [247]	Low red meat	Cardiovascular disease	RCT	1	0.98	0.91 to 1.06	1.00	0.84 to 1.19	Number of patients and events in intervention and control group for coronary heart disease death and fatal stroke, mean 8.1 years follow-up; Table 4 in Howard 2006 [262]
Johnston 2019 [247]	Low red meat	Colorectal cancer	RCT	1	1.04	0.90 to 1.20	1.02	0.89 to 1.18	Number of patients and events in intervention and control group for colorectal cancer, mean 5 years follow-up; Supplementary Table 2 in Thomson 2014 [263]
Johnston 2019 [247]	Low red meat	Type 2 diabetes	RCT	1	0.96	0.90 to 1.03	0.95	0.89 to 1.02	Number of patients and events in intervention and control group for type 2 diabetes, mean 8.1 years follow-up; Figure 2 in Tinker 2008 [264]
Stratton 2011 [224]	Vitamin C	Prostate cancer	RCT	1	1.01	0.92 to 1.10	1.10	0.93 to 1.29	Number of patients and events in intervention and control group for prostate cancer, mean 8 years follow-up; Figure 3 in Gaziano 2009 [265]
Trikalinos 2012 [239]	Vitamin E	Cardiovascular disease	CS	8	0.85	0.78 to 0.93	0.85	0.78 to 0.93	Data not available to calculate a RR for the HR; assumption HR=RR
					logHR	95% CI	RR	95% CI	
Kastorini 2011 [6]	Mediterranean diet	Metabolic syndrome	RCT	2	-0.80	-1.42 to -0.18	0.45	0.24 to 0.84	Meta-analysis of RCTs (clinical trials) shown in Figure 2; HRs pooled with inverse-variance method; data not available to calculate a RR for the HR; assumption HR=RR

			CS	2	-0.76	-2.16 to 0.63	0.46	0.11 to 1.88	Meta-analysis of CSs (prospective studies) shown in Figure 2; HRs pooled using inverse-variance models with random effects; data not available to calculate a RR for the HR; assumption HR=RR
					OR	95% CI	RR	95% CI	
Aguilar-Cordero 2020 [218]	Vitamin D	Pre-eclampsia	RCT	7	0.61	0.36 to 1.04	0.66	0.42 to 1.05	Meta-analysis of RCTs (n=7) shown in Figure 27; RRs calculated using number of patients and events in intervention and control groups; pooled using inverse-variance models with random effects
			CS	7	1.47	1.29 to 1.67	0.82	0.60 to 1.11	Meta-analysis of observational studies (n=11) shown in Figure 4; prospective studies were separated (Achkar 2015 [266], Arisoy 2015 [267], Bener 2013 [268], Burris 2014 [269], Gidlöf 2015 [270], Schneuer 2014 [271], van Weert 2016 [272]); ORs inverted (from low vs. high to high vs. low); ORs converted to RR with ACR= 0.10; risk observed in the control group of the corresponding BoE of RCTs (Figure 24); pooled using inverse-variance models with random effects
Chung 2011 [242]	Vitamin D	Colorectal cancer	CS	9	0.94	0.91 to 0.97	0.94	0.91 to 0.97	ACR= 0.0201; risk observed in the control group of the corresponding BoE of RCTs (Trivedi 2003 [273], colorectal cancer)
Chung 2011 [242]	Vitamin D	Breast cancer	CS	4	0.99	0.97 to 1.01	0.99	0.97 to 1.01	ACR= 0.0124; risk observed in the control group of the corresponding BoE of RCTs (Trivedi 2003 [273], breast cancer)

Feng 2015 [4]	Folic acid	Congenital heart defects	RCT	1	0.42	0.19 to 0.98	0.58	0.33 to 1.02	RR calculated using number of patients and events in intervention and control group for cardiovascular defects; Table 4 in Czeizel 1998 [274]; converted to a standardized food portion of the corresponding CS (from 0.8mg/day to 0.6mg/day)
			CS	1	0.60	0.38 to 0.96	0.60	0.38 to 0.96	ACR= 0.0084; risk observed in the control group of the corresponding BoE of RCTs (Czeizel 1998 [274], cardiovascular defects)
Jonker 2020 [248]	Folic acid	Low birthweight	RCT	1	0.93	0.81 to 1.08	0.96	0.88 to 1.04	Meta-analysis shown in Figure 4; RR calculated using number of patients and events in intervention and control group of Christian 2003 [275]
			CS	5	0.55	0.44 to 0.70	0.71	0.60 to 0.83	Meta-analysis for non-randomised studies (n=7) in Figure 4; prospective cohorts were separated (Balarajan 2013 [276], Chaudhary 2012 [277], Joseph 2011 [278], Nisar 2014 [279], Wang 2016 [280]); OR converted to RR with ACR= 0.4341; risk observed in the control group of the corresponding BoE of RCTs (Christian 2003 [275])
Jonker 2020 [248]	Folic acid	Small for gestational age	RCT	1	0.96	0.81 to 1.14	0.98	0.92 to 1.06	Meta-analysis shown in Figure 6; RR calculated using number of patients and events in intervention and control group of Christian 2003 [275]
			CS	4	0.66	0.41 to 1.07	0.80	0.50 to 1.11	Meta-analysis for non-randomised studies (n=4) in Figure 4; we combined post-conceptional and pre-conceptional folic acid use in Zheng 2016 [281]; OR converted to RR with ACR= 0.5870; risk observed in the control group of the corresponding BoE of RCTs Christian 2003 [275]

Stratton 2011 [224]	Multivitamins	Prostate cancer	RCT	1	0.90	0.61 to 1.33	0.90	0.62 to 1.33	Meta-analysis shown in Figure 2a; RR calculated using number of patients and events in intervention and control group of Meyer 2005 [282]
			CS	1	1.05	0.94 to 1.16	1.05	0.94 to 1.16	ACR= 0.0215; risk observed in the control group of the corresponding BoE of RCTs (Meyer 2005 [282], Occurrence of prostate cancer)
Stratton 2011 [224]	β -carotene	Prostate cancer	RCT	1	1.20	0.86 to 1.66	1.19	0.87 to 1.65	Meta-analysis shown in Figure 2E; RR calculated using number of patients and events in intervention and control group of Heinonen 1998 [283]
			CS	1	0.69	0.62 to 0.77	0.69	0.63 to 0.77	ACR= 0.0092; risk observed in the control group of the corresponding BoE of RCTs (Heinonen 1998 [283]); Figure 2E
Stratton 2011 [224]	Vitamin E	Prostate cancer	RCT	3	1.02	0.89 to 1.16	0.98	0.81 to 1.20	Meta-analysis for all study types (Figure 2B) was separated; effect estimates for RCTs and CS were converted from OR to RR; pooled using inverse-variance models with random effects
			CS	5			1.09	0.94 to 1.27	
Vinceti 2018a [7]	Selenium	Any cancer	CS	7	0.72	0.55 to 0.93	0.75	0.59 to 0.94	ACR= 0.1509; risk observed in the control group of the corresponding BoE of RCTs; Analysis 1.1., Outcome 1 Any cancer risk
Vinceti 2018a [7]	Selenium	Cancer mortality	CS	1	0.93	0.82 to 1.04	0.93	0.83 to 1.04	ACR= 0.0597; risk observed in the control group of the corresponding BoE of RCTs; Analysis 1.2., Outcome 2 Cancer mortality
Vinceti 2018b [236]	Selenium	Type 2 diabetes	CS	1	2.39	1.32 to 4.32	2.28	1.31 to 3.89	ACR= 0.0335; risk observed in the control group of the corresponding BoE of RCTs; Table 2

Zhang 2018 [230]	Vitamin D	Gestational diabetes	CS	24	1.85	1.47 to 2.32	0.58	0.44 to 0.76	Meta-analysis for observational studies (n=38) in Figure 1; we included only prospective cohort (Alvarez-Silvares 2016 [284], Burris 2011 [285], Burris 2012 [286], Davies-Tuck 2013 [287], Davies-Tuck 2015 [288], Fernandez-Alonso 2010 [289], Hu 2015 [290], Inancli 2016 [291], Kramer 2014 [292], Loy 2015 [293], Park 2014 [294], Song 2015 [295], Tomedi 2013 [296], Zhou 2014 [297], Zhu 2015 [298]) and nested-case-control studies (Arnold 2015 [299], Baker 2012 [300], Dodds 2013 [301], Dodds 2016 [302], Giguere 2014 [303], Li 2016 [304] Makgoba 2011 [305], Schneuer 2014 [271], Zhang 2008 [306]); ORs were inverted (from low vs. high to high vs. low); ORs were pooled using inverse-variance models with random effects; ACR= 0,1060; risk observed in the control group of the corresponding BoE of RCTs (Hossain 2014 [307], Mojibian 2015 [308], Sablok 2015 [309], Yap 2014 [310]; Figure S19
Zhou 2017 [232]	Vitamin D	Preterm birth	RCT	5	0.57	0.36 to 0.91	0.58	0.30 to 1.11	Meta-analysis for RCTs (n=6) in Figure 3; we excluded Asemi 2013 [311] due to retraction; RRs calculated using number of patients and events in intervention and control groups; pooled using inverse-variance models with random effects

			CS	12	1.25	1.13 to 1.38	0.88	0.73 to 1.05	Meta-analysis of observational studies (n=14) in Supplementary Figure 2; we separated the studies of the prospective, case-cohort and nested case-cohort subgroups; ORs were inverted (from low vs. high to high vs. low); ORs were pooled using inverse-variance models with random effects; ACR= 0.0460; risk observed in the control group of the corresponding BoE of RCTs, Figure 3
					Rate Ratio	95% CI	RR	95% CI	
Yao 2019 [228]	Vitamin D	Any fracture	RCT	11	1.06	0.98 to 1.14	1.06	0.97 to 1.15	Meta-analysis for RCTs (n=11) in Figure 2, Any fracture; RRs calculated using number of patients and events in intervention and control groups; pooled using inverse-variance models with random effects
			CS	10	0.93	0.89 to 0.96	0.89	0.83 to 0.96	Meta-analysis for CSs (n=10) in Figure 1, Any fracture; RRs calculated using number of patients and events in intervention and control groups; pooled using inverse-variance models with random effects
					RR	95% CI	RR	95% CI	
Aburto 2013 [2]	Low sodium	All-cause mortality	CS	2	1.06	0.94 to 1.20	0.95	0.71 to 1.27	Meta-analysis of CSs (n=7) shown in Figure 3.20 [312]; we included only CSs reporting on effect of dietary intake (Cohen 2008 [313], He 1999 [314]); RRs were inverted (from high vs. low to low vs. high); pooled using inverse-variance models with random effects

Aburto 2013 [2]	Low sodium	Cardiovascular disease	CS	3	1.12	0.93 to 1.34	0.87	0.64 to 1.18	Meta-analysis of CSs (n=9) shown in Figure 3.4 [312]; we included only CSs reporting on effect of dietary intake (Cohen 2008 [313], He 1999 [314], Umesawa 2008 [315]); RRs were inverted (from high vs. low to low vs. high); pooled using inverse-variance models with random effects
Afshin 2014 [240]	Nuts	Coronary heart disease	RCT	1	Fatal: 0.76 Nonfatal: 0.78	0.69 to 0.84	0.79	0.55 to 1.14	Meta-analysis for RCTs and CSs (Figure 2A and 2B; Fatal and nonfatal CHD) was separated; RRs for CSs with nonfatal and fatal outcomes were pooled using inverse-variance models with random effects
			CS	6		0.67 to 0.92	0.76	0.69 to 0.83	
Afshin 2014 [240]	Nuts	Stroke	RCT	1	0.89	0.74 to 1.05	0.62	0.44 to 0.87	Meta-analysis for RCTs and CSs (Figure 2C; Total stroke) was separated; RRs for CS were pooled using inverse-variance models with random effects
			CS	2			0.95	0.83 to 1.09	
Afshin 2014 [240]	Nuts	Type 2 diabetes	RCT	1	0.87	0.81 to 0.94	0.86	0.68 to 1.08	Meta-analysis for RCTs and CSs (Figure 2D; diabetes) was separated; RRs for CS were pooled using inverse-variance models with random effects
			CS	4			0.87	0.80 to 0.95	
Chowdhury 2014b	Vitamin D	All-cause mortality	CS	68	1.44	1.34 to 1.55	0.69	0.65 to 0.75	Value of RR and 95% CI displayed in Figure 1, All-cause mortality was inverted (from low vs. high to high vs. low).
Kim 2018 [223]	Multivitamins/Minerals	Cardiovascular disease	RCT	1	1.00	0.97 to 1.04	0.95	0.83 to 1.09	Meta-analysis for RCTs and CSs (Figure 2; CVD mortality) was separated; RRs for CS were pooled using inverse-variance models with random effects
			CS	9			1.00	0.96 to 1.05	
Kim 2018 [223]	Multivitamins/Minerals	Stroke	RCT	1	0.98	0.91 to 1.05	1.05	0.70 to 1.56	Meta-analysis for RCTs and CSs (Figure 2; Stroke Incidence) was separated; RRs for CS were pooled using inverse-variance models with random effects
			CS	3			0.98	0.91 to 1.04	

Lin 2020 [251]	Vitamin D	Nephrolithiasis	CS	3	1.22	1.01 to 1.49	1.24	1.01 to 1.51	Meta-analysis for observational studies (Figure 5, Total Vitamin D, n=4); prospective studies were separated (NHS I [316], NHS II [317], HPFS [317]; RRs pooled using inverse-variance models with random effects
Sayehmiri 2018 [257]	Selenium	Prostate cancer	CS	2	0.77	0.52 to 1.14	0.89	0.64 to 1.23	Meta-analysis of CSs (n=9) shown in Table 2, Type of study; we included only CSs reporting on effect of dietary intake (Hartman 1998 [318], Peters 2008 [319]; RRs pooled using inverse-variance models with random effects
Stratton 2011 [224]	Vitamin C	Prostate cancer	CS	1	NR	NR	0.96	0.86 to 1.06	Number of patients and events in intervention and control group for prostate cancer reported in Table 2
Wien 2012 [226]	Folic acid	Cancer	CS	5	1.03	0.92 to 1.16	1.03	0.90 to 1.18	Meta-analysis of CSs (n=6); we included only prospective CSs (Maruti 2009 [320], Oaks 2010 [321], Skinner 2004 [322], Slatore 2008 [323], Stolzenberg-Solomon 2006 [324]); RRs pooled using inverse-variance models with random effects
Wolf 2017 [8]	Multivitamins	Preterm birth	RCT	1	NR	NR	1.09	0.43 to 2.77	Number of patients and events in intervention and control group for preterm birth; Table 4 in Brough 2010 [325]
Wolf 2017 [8]	Multivitamins	Stillbirth	RCT	1	NR	NR	2.43	0.12 to 50.05	Number of patients and events in intervention and control group for preterm birth; Table 2 in Kirke 1992 [326]
Yang 2016 [235]	Folic acid	Gestational hypertension	RCT	1	1.02	1.00 to 1.04	0.62	0.40 to 0.94	Meta-analysis for RCTs and CSs (Figure 2) was separated; we included only prospective CSs reporting on postconceptional folic acid use (Li 2013 [327], Oken 2007 [328], Timmermans (Postcon) 2011 [329]; RRs of CSs were pooled using inverse-variance models with random effects
			CS	3			1.05	1.00 to 1.11	

Yu 2021 [229]	Folic acid	Pre-eclampsia	CSs	13	0.83	0.74 to 0.93	0.85	0.76 to 0.94	Meta-analysis for all study types (Figure 3); we included only prospective CSs (Bodnar 2006 [330], Bukowski 2009 [331], Catov 2009 [332], Catov 2011 [333], Li 2020 [334], Liu 2015 [335], Martinussen 2015 [336], Oken 2007 [328], Timmermanns 2011 [329], Vanderlelie 2014 [337], Wang 2015 [338], Wen 2008 [339], Wen 2016 [340]; RRs pooled using inverse-variance models with random effects
Zhao 2014 [261]	Multivitamins/Minerals	Nuclear cataract	RCT	2	NR	NR	0.73	0.59 to 0.90	RRs of reported RCTs (Maraini 2008 [341], Sperduto 1993 [342]) were pooled using inverse-variance models with random effects
Zhao 2014 [261]	Multivitamins/Minerals	Cortical cataract	RCT	2	NR	NR	0.92	0.69 to 1.23	RRs of reported RCTs (Maraini 2008 [341], Sperduto 1993 [342]) were pooled using inverse-variance models with random effects
Zhao 2014 [261]	Multivitamins/Minerals	Posterior subcapsular cataract	RCT	2	NR	NR	1.81	1.30 to 2.54	RRs of reported RCTs (Maraini 2008 [341], Sperduto 1993 [342]) were pooled using inverse-variance models with random effects
					MD	95% CI	MD	95% CI	
Amegah 2017 [231]	Vitamin D	Gestational length	RCT	2	-0.24	-0.69 to 0.22	0.16	-0.28 to 0.59	Meta-analysis for RCTs and CS (Figure 3C) was separated; MDs were inverted (from low vs. high to high vs. low)
			CS	1			0.08	0.20 to 1.40	
Kastorini 2011 [6]	Mediterranean diet	HDL-cholesterol	RCT	27	1.17	0.38 to 1.96	0.03	0.01 to 0.05	Meta-analyses for clinical trials shown in Table 4; we excluded non-randomised studies (Papadaki 2008 [343], Gomez 2005 [344]); MD was converted from mg/dl to mmol/l (factor: 0.026)
			CS	1	0.30	-1.77 to 2.37	0.01	-0.05 to 0.06	MD was converted from mg/dl to mmol/l (factor: 0.026)

Kastorini 2011 [6]	Mediterranean diet	Triglycerides	RCT	27	-6.14	-10.35 to 1.93	-0.07	-0.12 to -0.02	Meta-analyses for clinical trials shown in Table 4; we excluded non-randomised studies (Papadaki 2008 [343], Gomez 2005 [344]); MD was converted from mg/dl to mmol/l (factor: 0.0114)
			CS	1	-2.00	-6.74 to 2.74	-0.02	-0.08 to 0.03	MD was converted from mg/dl to mmol/l (factor: 0.0114)
Te Morenga 2013 [260]	Low dietary sugars	Weight gain	RCT	10	0.75	0.30 to 1.19	-0.75	-1.19 to 0.30	Meta-analysis of RCTs shown in Figure 4; MD was inverted (from high vs. low to low vs. high)
			CS	4	0.31	-0.07 to 0.68	-0.31	-0.68 to 0.07	Meta-analysis of CS (n=4) shown in Web figure 3 (Weight difference between highest and lowest intakes at baseline); MD inverted (from high vs. low to low vs. high)
Te Morenga 2013 [260]	Low dietary sugars	Body mass index	CS	4	0.02	-0.00 to 0.05	-0.02	-0.05 to -0.00	Meta-analysis of CS (n=4) shown in Web figure 6; MD inverted (from high vs. low to low vs. high)
					MD	95% CI	SMD	95% CI	
Azad 2017 [220]	Nonnutritive sweeteners	Body mass index	RCT	3	-0.37	-1.10 to 0.36	-0.14	-0.43 to 0.16	Meta-analysis for RCTs shown in Figure 2A using MD; we converted MD to SMD using mean, SD and number of patients in intervention and control groups, pooled using inverse-variance models with random effects
			CS	1	0.77	0.47 to 1.07	-0.27	0.17 to 0.38	CS of Fowler 2008 [345], described in results for body mass index; MD was standardised using mean, SD and number of patients in exposure and comparator groups
Jonker 2020 [248]	Folic acid	Mean birthweight	RCT	3	0.56	0.15 to 0.97	0.20	0.08 to 0.31	Meta-analysis of RCTs shown in Figure 2; MD was standardised using mean, SD and number of patients in intervention and control groups, pooled using inverse-variance models with random effects

Picasso 2019 [256]	Vegetarian diet	Triglycerides	RCT	5	0.11	-0.10 to 0.31	0.08	-0.07 to 0.23	Meta-analysis of RCTs shown in Figure 6, Analysis 5.1.1; MD was standardised using mean, SD and number of patients in intervention and control groups, pooled using inverse-variance models with random effects
Picasso 2019 [256]	Vegetarian diet	HDL-cholesterol	RCT	4	-0.06	-0.17 to 0.04	-0.13	-0.29 to 0.04	Meta-analysis of RCTs shown in Figure 8, Analysis 7.1.1; MD was standardised using mean, SD and number of patients in intervention and control groups, pooled using inverse-variance models with random effects
					SMD	95% CI	MD	95% CI	
Te Morenga 2013 [260]	Low dietary sugars	Body mass index	RCT	3	0.09	-0.14 to 0.32	-0.06	-0.15 to 0.04	Meta-analysis of RCTs (n=5) shown in Figure 6 with SMDs; we included only RCTs reporting the outcome body mass index (Ebbeling 2006 [346], James 2004 [347], Paineau 2008 [348]; MDs were calculated using mean, SD and number of patients in intervention and control groups; pooled using inverse-variance models with random effects
					SMD	95% CI	SMD & MD	95% CI	
Jonker 2020 [248]	Folic acid	Mean birthweight	CS	2	0.31	0.18 to 0.43	SMD 0.36 MD 0.14	0.08 to 0.65 0.05 to 0.24	Meta-analysis of non-randomised studies shown in Figure 2; we included only the two prospective studies on folic acid supplementation (Chaudhary 2012 [277], Wang 2016 [280]; MD and SMDs calculated and pooled using inverse-variance models with random effects

					MD	95% CI	MD & SMD	95% CI	
Picasso 2019 [256]	Vegetarian diet	Triglycerides	CS	2	0.06	-0.22 to 0.35	MD 0.07 SMD 0.03	-0.39 to 0.53 -0.19 to 0.26	Meta-analysis of CSs shown in Figure 6, Analysis 5.1.2; we included only the prospective cohort study (Brestrich 1996 [349]; MD was standardised using mean, SD and number of patients in intervention and control group, pooled using inverse-variance models with random effects
Picasso 2019 [256]	Vegetarian diet	HDL-cholesterol	CS	1	0.16	-0.24 to 0.55	MD -0.03 SMD -0.09	-0.10 to 0.04 -0.32 to 0.13	Meta-analysis of CSs shown in Figure 8, Analysis 7.1.2; we included only the prospective cohort study (Brestrich 1996 [349]; MD was standardised using mean, SD and number of patients in intervention and control group, pooled using inverse-variance models with random effects
Thorne-Lyman 2012 [225]	Vitamin D	Mean birthweight	RCT	5	58.07	-63.87 to 180.00	MD 0.06 SMD 0.13	-0.06 to 0.18 -0.12 to 0.39	Meta-analysis of observational studies shown in Figure S1 (Bolus or daily dosing and mean birthweight); MD was converted from gram to kilogram; MD was standardised using mean, SD and number of patients in intervention and control groups, pooled using inverse-variance models with random effects
			CS	2	52.66	3.54 to 101.77	MD 0.05 SMD 0.11	0.003 to 0.10 0.01 to 0.21	Meta-analysis of observational studies shown in Figure S1 (Observational studies of intake); MD was converted from grams to kilograms; MD was standardised using mean, SD and number of patients in intervention and control groups, pooled using inverse-variance models with random effects
					Correlation				
Miller 2014 [253]	Low calorie sweeteners	Body weight	CS	4	0.02	-0.01 to 0.06	-	-	Impossibility to convert weighted group mean correlation (CSs) to weighted group MD (RCTs)

Setien-Suero 2016 [259]	Homocysteine	Mini Mental State Examination Score	CS	3	-0.11	-0.18 to -0.05	-	-	Impossibility to convert β (CSs) to SMD (RCTs)
Zhang 2018 [230]	Vitamin D	Fasting plasma glucose	CS	13	-0.09	-0.20 to 0.01	-	-	Impossibility to convert correlation coefficient (CSs) to SMD (RCTs)

ACR=assumed control risk; BoE=bodies of evidence; CHD=coronary heart disease; CI=confidence interval; CS=cohort study; CVD=cardiovascular disease; HDL=high density lipoprotein; HPFS=Health Professionals Follow-Up Study; HR=hazard ratio; MD=mean difference; NHS=Nurses' Health Study; OR=odds ratio; RCT=randomised controlled trial; RR=risk ratio; SD=standard deviation; SMD=standardised mean difference

Supplementary Table S6 Characteristics of included BoE from randomised controlled trials

Reference	Intervention	Outcome	Studies, n	Sample size, n	Cases, n	Description of population	Age (years)	Description of intervention	Description of comparator	Description of outcome	Study design	Follow-up
Aburto 2013 [2]	Low sodium	All-cause mortality	4	3,595	69	Male and female population/ mostly healthy with normal blood pressure (3x); partly with hypertension (1x)/ partly moderate overweight (1x)	20-54 (range)	Dietary and behavioural counselling to reduce sodium intake/ partly with weight loss intervention (1x)	No or unspecific counselling/ usual care (1x); no intervention (1x) or counselling for generally healthy diet (1x); weight loss alone (1x)	All-cause mortality	Parallel, factorial	0.5-3 years
Aburto 2013 [2]	Low sodium	Cardiovascular disease	2	720	93	Male and female population/ with hypertension, but otherwise healthy/ partly obese	66.5 (mean) or >50	Dietary and behavioural counselling to reduce sodium intake	No or unspecific counselling/ general advice on subjects unrelated to CVD, blood pressure, nutrition (1x); no intervention (1x)	Cardiovascular disease	Parallel, factorial	2-2.5 years
Afshin 2014 [240]	Nuts	Coronary heart disease	1	7,447	69	Male and female population/ without CVD, but with either T2D or ≥ 3 major cardiovascular risk factors	55-80 (range); 66.7-67.3 (mean)	Nut consumption/ intake of tree nuts (e.g. walnut, almond, hazelnuts, pecan, cashew, and pistachio) and peanuts/ dose-response: per 4 weekly servings (28.4g)	Dose-response	Total incident ischemic heart disease (fatal and nonfatal)	Parallel	4.8 years
Afshin 2014 [240]	Nuts	Stroke	1	7,447	90	Male and female population/ without CVD, but with either T2D or ≥ 3 major cardiovascular risk factors	55-80 (range); 66.7-67.3 (mean)	Nut consumption/ intake of tree nuts (e.g. walnut, almond, hazelnuts, pecan, cashew, and pistachio) and peanuts/ dose-response: per 4 weekly servings (28.4g)	Dose-response	Total stroke (ischemic and haemorrhagic)	Parallel	4.8 years
Afshin 2014 [240]	Nuts	Type 2 diabetes	1	3,541	273	Male and female population/ without CVD and T2D, but with ≥ 3 major cardiovascular risk factors	55-80 (range); 66.2-67.2 (mean)	Nut consumption/ intake of tree nuts (e.g. walnut, almond, hazelnuts, pecan, cashew, and pistachio) and peanuts/ dose-response: per 4 weekly servings (28.4g)	Dose-response	Type 2 diabetes	Parallel	4.1 years
Aguilar-Cordero 2020 [218]	Vitamin D	Pre-eclampsia	7	1,660	160	Female population/ pregnant/ without specific previous pathologies	24.8-32.4 (mean)	Vitamin D supplementation/ 600-4,400IU/day or 50,000IU every 2 weeks (2x) or single dose of 60,000IU or 2-4 doses of 120,000IU	No or low vitamin D supplementation/ placebo (2x); routine care (2x); low vitamin D (400IU/day; 2x); no intervention (1x)	Gestational hypertension with proteinuria and/or other maternal organ dysfunction and/or utero-placental dysfunction/ ≥ 20 weeks of gestation	Parallel	NR
Alexander 2017 [219]	Omega-3 fatty acids	Coronary heart disease	18	93,633	NR	Male and female population/ with or without CHD/ partly with low left ventricular systolic function or T2D or peripheral arterial disease (3x)/ non-hospitalised	>18	EPA+DHA intake or supplementation/ food, ethyl esters, fish oil, fatty fish or EPA+DHA enriched margarine/ 0.38-5.04g/day	Mostly oils not containing EPA+DHA (olive, corn or sunflower oil, 11x)/ partly gelatine (1x); aluminium hydroxide (1x); dietary advice (2x); no supplementation (2x)	Fatal and nonfatal myocardial infarction, coronary death, sudden cardiac death, angina	Parallel	0.5-7 years

Amegah 2017 [231]	Vitamin D	Gestational length	2	525	N/A	Female population/ pregnant/ with singleton pregnancy	25.19-27.4 (mean)	Vitamin D supplementation/ 2,000 or 4,000IU/day/ partly with 400mg/day ferrous sulfate and 600mg/day calcium	No or low vitamin D supplementation/ 400IU/day (1x); routine care with 400mg/day ferrous sulfate and 600mg/day calcium (1x)	Gestational length (weeks)	Parallel	≤16 or ≤20 weeks of gestation till delivery
Azad 2017 [220]	Nonnutritive sweeteners	Body mass index	3	242	N/A	Male and female population/ with mild hypertension (2x); overweight (1x), but otherwise healthy (3x)	32-52 (mean)	Stevioside capsules: 3.8-15mg/kg per day (1x); 1,500mg/day (1x); 250ml/day artificially sweetened beverage (1x)	Placebo (2x); water (1x)	Change in body mass index (kg/m ²) at latest follow-up	Parallel	0.5-2 years (mean)
Bolland 2015 [221]	Calcium	All fractures	20	58,573	6,725	Male and female population/ mostly living in the community (17x); partly in residential care (3x)	56-86 (mean)	Calcium supplementation/ 500-1,600mg/day/ alone (7x); combined with vitamin D (400-1,000IU/day or 300,000IU i.m. or UV exposure, 13x)/ partly with alendronate (1x)	Placebo (10x); not further described (8x)/ partly with vitamin D (1x); alendronate (1x)	Fracture at any site	Parallel, factorial, cluster	1-7 years
Chowdhury 2012 [241]	Omega-3 fatty acids	Cerebrovascular disease	2	31,181	978	Male and female population/ without CVD, but at risk of CVD (with hyperlipidaemia or dysglycaemia)	61-63.6 (mean)	Omega-3 fatty acids supplementation/ 1.8g/day EPA (1x); 1g/day EPA+DHA/ partly with Pravastatin 10mg/day or Simvastatin 6mg/day (1x)	Placebo containing olive oil (1x); Pravastatin 10mg/day or Simvastatin 5mg/day (1x)	Fatal or non-fatal ischaemic stroke or haemorrhagic stroke, cerebrovascular accident, or transient ischaemic attack	Parallel, factorial	4.6-6.2 years (mean)
Chowdhury 2014a [234]	α-linolenic acid	Coronary disease	4	18,866	419	Male and female population/ mostly with CHD (3x); partly general healthy (1x)	40-80 (range) or <70	α-linolenic acid supplementation or dietary oils; dose: 2-5.5g/day margarine supplemented with α-linolenic acid (2x); flaxseed oil (1); linseed oil (1x)	General dietary advice (1x); sunflower oil (2x); placebo margarine (1x)	Fatal or nonfatal myocardial infarction, CHD, coronary insufficiency, coronary death, angina, angiographic coronary stenosis	Parallel	1-3.4 years
Chowdhury 2014a [234]	Omega-6 fatty acids	Coronary disease	8	14,476	974	Male and female population/ mostly male/ with CHD (5x); at high risk of CHD (3x)	30-88 (range) or <66 or <70	Diet rich of linolenic acid; 8-72% linolenic acid intake of total daily fat or 40.5-85g/day of polyunsaturated fatty acids; mixed polyunsaturated intervention with linoleic acid as the primary fatty acid (6x); linolenic acid-specific supplementation (2x)	Usual diet/ care (2x); unspecific dietary advice (3x); low linolenic acid diet (1x); low linoleic acid diet with high proportion of other fats (2x)	Fatal or nonfatal myocardial infarction, CHD, coronary insufficiency, coronary death, angina, angiographic coronary stenosis	Parallel, factorial, crossover, cluster	1-8 years
Chowdhury 2014b [3]	Vitamin D	All-cause mortality	22	30,716	5,114	Male and female population/ with or without pre-existing chronic disease (CVD, metabolic disease, cancer)/ community dwelling or institutionalised	56-85 (IQR)	Vitamin D supplementation/ mostly oral; partly with injections/ vitamin D3: 10-6,000IU/day; vitamin D2: 208-4,500IU/day	Placebo (18x); no treatment (4x)	Deaths from any cause	Parallel, factorial, cluster	0.38-6.8 years (mean)

Chung 2011 [242]	Vitamin D	Colorectal cancer	1	2,686	55	Male and female general population/ elderly/ community dwelling	65-85 (range); 75 (mean)	Vitamin D supplementation/ 100,000IU every 4 months	Placebo	Colorectal cancer incidence (fatal outcomes included)	Parallel	5 years (duration + follow-up)
Chung 2011 [242]	Vitamin D	Breast cancer	1	649	8	Female general population/ elderly/ community dwelling	65-85 (range); 75 (mean)	Vitamin D supplementation/ 100,000IU every 4 months	Placebo	Breast cancer incidence (fatal outcomes included)	Parallel	5 years (duration and follow-up)
Ding 2017 [243]	Dairy	Systolic blood pressure	8	735	N/A	Male and female population/ partly healthy (3x); partly with overweight or obesity (2x); prehypertension (1x); metabolic disease (2x)	20-85 (range)	High dairy intake/ 3-5 servings/day (6x); 400-650g/day (2x)/ partly reduced fat products (4x)	Low or no dairy intake/ usual diet (2x); 1-2 servings/day (2x); diet without milk or dairy (1x); control products equivalent in macro-nutrients and sodium (1x); fruit juice and biscuits (1x); NR (1x)	Systolic blood pressure (mmHg)	Parallel, crossover	1-12 months (duration)
Feng 2015 [4]	Folic acid	Congenital heart defects	1	4,862	30	Female population/ in childbearing age/ planning to get pregnant	NR	Folic acid supplementation/ periconceptional/ capsule including 12 vitamins, four minerals and three trace elements), 3 month before pregnancy/ 0.8mg/day	Placebo-like trace element control group (three trace elements only)	Congenital heart defects/ overall or any one of the subtypes in infants (e.g. conotruncal defects, septal defects)	Parallel	3 month before conception till delivery
Filippini 2020 [244]	Green Tea	Prostate cancer	3	201	32	Male population/ at high risk of prostate cancer / with high-grade prostate intraepithelial neoplasia (2x) and/or atypical small acinar proliferation (1x)	30-80 (range)	Green tea supplementation/ capsules with green tea extract or green tea catechins: 300mg/day (2x); 400mg/day of epigallocatechin-3-gallate (1x)	Placebo	Prostate cancer	Parallel	1 year (duration)
Filippini 2020 [244]	Green Tea	Endometrial cancer	1	1,075	1	Female population/ postmenopausal/ without history of cancer, but with high mammography density	50-70 (range)	Green tea supplementation/ capsules with green tea extract, that is 1,315mg ($\pm$ 116mg) of total catechins/day, including 843mg/day ($\pm$ 44mg/day) of epigallocatechin-3-gallate	Placebo	Endometrial cancer	Parallel	1 year (duration)
Fu 2021 [245]	Vitamin C	Lung cancer	2	22,237	177	Male and female population/ with or at high risk of CVD/ partly with history of CVD, T2D or cancer	60.4-64.3 (mean)	Vitamin C supplementation/ 500mg/day/ with or without vitamin E and/or β -carotene	Placebo/ with or without vitamin E and/or β -carotene	Confirmed diagnosis of lung cancer	Factorial	8-9.4 years (mean)

Fu 2021 [245]	Vitamin C	Breast cancer	1	7,627	257	Female population/ with or at high risk for CVD/ partly with history of cancer or T2D	60.4 (mean)	Vitamin C supplementation/ 500mg/day/ with or without vitamin E and/or β -carotene	Placebo/ with or without vitamin E and/or β -carotene	Confirmed diagnosis of breast cancer	Factorial	9.4 years (mean)
Fu 2021 [245]	Vitamin C	Colorectal cancer	2	22,147	206	Male and female population/ with or at high risk of CVD/ partly with history of CVD, T2D or cancer	60.4-64.3 (mean)	Vitamin C supplementation/ 500mg/day/ with or without vitamin E and/or β -carotene	Placebo/ with or without vitamin E and/or β -carotene	Confirmed diagnosis of colorectal cancer	Factorial	8-9.4 years (mean)
Gayer 2019 [246]	Apples	Body weight	5	393	N/A	Male and female population/ healthy or at risk of CVD or metabolic disease/ partly with hypertension and/or hyper-/ dyslipidaemia	36.2-56.7 (mean)	Apple intake/ 170-900g/day apple or 500-2,250ml/day apple juice or 75g/day dried apple or 22mg/day apple pomace	No or low apple intake: controlled beverage (2,250ml/day); dried plums (100g/day); oatmeal cookie (180g); only restricted diet allowed; green kiwi fruit (360g/day)	Body weight change (kg)	Parallel, crossover	1-12 months
Grosso 2015 [222]	Mediterranean diet	Cardiovascular disease	4	12,293	590	Male and female population/ partly with previous myocardial infarction (2x); partly at risk of CVD (2x)	48-67.3 (mean)	Advice to follow Mediterranean dietary pattern: Indo-Mediterranean diet (1x); Mediterranean diet (3x)	Control interventions: prudent diet (2x); general health advice (1x); low-fat diet (1x)	CVD/ including CVD mortality, myocardial infarction and stroke incidence	Parallel, factorial	2-5 years
Jiang 2019 [5]	Vitamin E	Age-related cataract	6	92,243	5,219	Male and female population/ mostly male (4x)/ without (ascertained) cataract	50-80 (range); ≥ 45	Vitamin E supplementation/ 50-334mg/day or 267-400mg every other day/ alone (4x)/ partly with β -carotene (20mg/day) (2x)	Placebo	Age-related cataract	Parallel, factorial	4-9.7 years (mean/ median)
Jiang 2019 [5]	β -carotene	Age-related cataract	4	88,266	3,415	Male and female population/ mostly male (3x)/ without (ascertained) cataract	40-84 (range); ≥ 45	β -carotene supplementation/ alone (2x); with 50mg/day vitamin E (20mg/day) (2x); 50mg every other day	Placebo	Age-related cataract	Factorial	2.1-12 years (median/ mean)
Johnston 2019 [247]	Low red meat	Cardiovascular disease	1	48,835	NR	Female population/ postmenopausal/ mostly overweight or obese (>70%); partly hypertensive	50-79 (range)	Behavioural support to reduce total dietary fat to 20% and to increase intake of vegetables and fruit and grains	Habitual diet/ general dietary advice by educational materials	Cardiovascular mortality	Parallel	<13.8 years
Johnston 2019 [247]	Low red meat	Type 2 diabetes	1	45,887	NR	Female population/ postmenopausal/ without T2D/ mostly overweight or obese (>70%); partly hypertensive	50-79 (range); 62.2 (mean)	Behavioural support to reduce total dietary fat to 20% and to increase intake of vegetables and fruit and grains	Habitual diet/ general dietary advice by educational materials	Type 2 diabetes	Parallel	8.1 years

Johnston 2019 [247]	Low red meat	Colorectal cancer	1	48,835	NR	Female population/ postmenopausal/ mostly overweight or obese (>70%); partly hypertensive	50-79 (range)	Behavioural support to reduce total dietary fat to 20% and to increase intake of vegetables and fruit and grains	Habitual diet/ general dietary advice by educational materials	Colorectal cancer incidence	Parallel	<12.3 years
Jonker 2020 [248]	Folic acid	Mean birthweight	3	1,503	N/A	Female population/ pregnant/ in low- and middle-income countries	NR	Folic acid supplementation/ 0.4-5mg/ with vitamin A (100µg) (1x); with formulated food multimix (1x); with 120mg iron (1x)	No folic acid supplementation: diet + placebo (1x); vitamin A (1x); placebo (1x)	Mean birthweight (kg)	Parallel, cluster	<12 months before pregnancy or 20 weeks of gestation till delivery; NR (1x)
Jonker 2020 [248]	Folic acid	Low birthweight	1	3,084	1,313	Female population/ pregnant/ low- and middle-income countries	NR	Folic acid supplementation/ 400µg with vitamin A (100µg)	No folic acid supplementation: vitamin A alone (100µg)	Incidence of low birthweight; <2500g; measured within 72 hours of birth	Cluster	<12 months before pregnancy
Jonker 2020 [248]	Folic acid	Small for gestational age	1	2,254	1,313	Female population/ pregnant/ low- and middle-income countries	NR	Folic acid supplementation/ 400µg with vitamin A (100µg)	No folic acid supplementation: vitamin A alone, 100µg	Infants whose weight is less than the 10th percentile for gestational age	Cluster	<12 months before pregnancy
Kastorini 2011 [6]	Mediterranean diet	Metabolic syndrome	2	1,404	NR	Male and female population/ at high risk of CVD (1x); with metabolic syndrome (1x)	44 or 67 (mean)	High adherence to Mediterranean diet: at least 250–300g fruits, 125–150g vegetables, 25–50g walnuts, 400g whole grains, increase consumption of olive oil/ carbohydrates: 50%–60%, protein: 15%–20%, fat: 30%, SF: 10%, cholesterol: 300g/day (1x); instructions to increase the 14-item Mediterranean score and free provision of olive oil and/or nuts (1x)	Low adherence to Mediterranean diet; control diet, i.e. low-fat diet or usual care	Metabolic syndrome	Parallel	1-2 years
Kastorini 2011 [6]	Mediterranean diet	HDL-cholesterol	27	3,734	N/A	Male and female population/ healthy and/or at risk of CVD or with CVD, diabetes or metabolic syndrome	18-80 (range); 23-61 (mean)	High adherence to Mediterranean diet: <30-40.1% from fat, mostly diets rich in olive oil and/or nuts and/or fish	Low adherence to Mediterranean diet	HDL-cholesterol (mg/dl)	Parallel, crossover	1 month-5 years
Kastorini 2011 [6]	Mediterranean diet	Triglycerides	27	3,734	N/A	Male and female population/ healthy and/or at risk of CVD or with CVD, diabetes or metabolic syndrome	18-80 (range); 23-61 (mean)	High adherence to Mediterranean diet: <30-40.1% from fat, mostly diets rich in olive oil and/or nuts and/or fish	Low adherence to Mediterranean diet	Triglycerides (mg/dl)	Parallel, crossover	1 months-5 years

Kastorini 2011 [6]	Mediterranean diet	Systolic blood pressure	14	3,068	N/A	Male and female population/ healthy and/or at risk of CVD or with CVD, diabetes or metabolic syndrome	18-80 (range); 44-58 (mean)	High adherence to Mediterranean diet: <30-40.1% from fat, mostly diets rich in olive oil and/or nuts and/or fish	Low adherence to Mediterranean diet	Systolic blood pressure (mmHg)	Parallel, crossover	1 months-5 years
Kim 2018 [223]	Multivitamins/ Minerals	Cardiovascular disease	1	14,641	829	Male population/ generally healthy/ partly with T2D or history of CVD	64.3 (mean)	Multivitamin/ Mineral supplementation: 1 tablet daily containing 13 vitamins and 17 minerals	Placebo	CHD mortality and stroke mortality	Factorial	11.2 years
Kim 2018 [223]	Multivitamins/ Minerals	Coronary heart disease	2	27,658	NR	Male population/ generally healthy/ partly with T2D or history of CVD	48.2-64.3 (mean)	Multivitamin/ Mineral supplementation: 1 tablet/day with 13 vitamins and 17 minerals (1x); combination of 120mg ascorbic acid, 30mg vitamin E, 6mg β -carotene, 100 μ g selenium, 20mg zinc (1x)	Placebo	Incident CHD events/ cardiac revascularization and fatal and nonfatal myocardial infarction	Parallel, factorial	7.5-11.2 years (mean)
Kim 2018 [223]	Multivitamins/ Minerals	Stroke	1	14,641	786	Male and female population/ generally healthy/ partly with T2D or history of CVD	64.3 (mean)	Multivitamin/ Mineral supplementation/ 1 tablet/day with 13 vitamins and 17 minerals	Placebo	Incident ischemic and haemorrhagic stroke (fatal and nonfatal)	Factorial	11.2 years
Kong 2014 [249]	Vitamins	Gastric cancer	11	152,848	661	Male and female population/ healthy or at high risk for gastric cancer	28-84 (range)	Vitamin supplementation/ vitamin A, B1, B2, B6, B12, C, E, E in different compositions	Placebo	Gastric cancer occurrence	Parallel, factorial	3-14.7 years
Lin 2019 [250]	Vitamin E	Bladder cancer	3	78,590	491	Male population/ mostly general population; partly with (history of) cancer/ partly smokers	56.9-67 (median/ mean)	Vitamin E supplementation/ 50mg/day (1x); 400IU/day α -tocopherol (1x); 400IU/day all-rac- α -tocopherylacetate (1x)	Low vitamin E intake/ control groups, placebo	Bladder cancer incidence	Parallel, factorial	6.1-10.3 years
Lin 2020 [251]	Vitamin D	Nephrolithiasis	7	14,652	220	Male and female population/ without prior nephrolithiasis/ partly with history of low-trauma osteoporotic fracture, knee osteoarthritis, adenomas or prediabetes/ partly obese/ overweight; partly postmenopausal	21-84 (range); 57-78 (mean)	Vitamin D supplementation/ 400-2,000IU/day or 20,000-40,000IU/week or 100,000IU/ month	Placebo/ partly with 500mg/day Calcium (1x)	Nephrolithiasis incidence	NR	1-5 years

Lin 2020 [251]	Calcium	Nephrolithiasis	4	2,934	41	Male and female population/ without prior nephrolithiasis/ partly with adenomas; partly postmenopausal	56-66.7 (mean)	Calcium supplementation/ 600-1,600mg/day	Placebo	Nephrolithiasis incidence	NR	2-4 years
Martinez-Gonzalez 2014b [252]	Olive oil	Coronary heart disease	1	7,447	75	Male and female population/ without CVD, but with T2D or ≥ 3 cardiovascular risk factors or family history of premature CHD	55-80 (range); 67 (mean)	Mediterranean diet supplemented with extra-virgin olive oil; per +25g/day of consumption (total: 29.3g/day)	Advice to reduce dietary fat (total: 2.6g d)	Myocardial infarction	Parallel	4.8 years
Martinez-Gonzalez 2014b [252]	Olive oil	Stroke	1	7,447	107	Male and female population/ without CVD, but with T2D or ≥ 3 cardiovascular risk factors or family history of premature CHD	55-80 (range); 67 (mean)	Mediterranean diet supplemented with extra-virgin olive oil; per +25g/day	Advice to reduce dietary fat	Stroke (ischaemic or haemorrhagic)	Parallel	4.8 years
Miller 2014 [253]	Low calorie sweeteners	Body weight	13	1,715	N/A	Male and female population/ generally healthy	4-65 (range); children (3x); adults (10x)	Low calorie sweeteners: low-calorie sweetened beverages (10.8-38oz/day) (7x); aspartame (285-2,700mg/day) (3x); isomalt (30g/day) (1x); aspartame (480-670mg/day) + acesulfame potassium + cyclamate + saccharin (1x); NR (1x)	Sugar sweetened beverages (5x); sugar sweetened food and beverages (1x); assigned to follow a habitual (usual) diet that contained sugar sweetened beverages and sugar sweetened foods (4x); energy-reduced diet (similar to the intervention) that did not include low calorie sweeteners; with table sugar for sweetener (1x); with lactose capsules (1x); without any aspartame- or saccharin-sweetened products (1x)	Body weight change	Parallel, crossover	3-78 weeks
Moazzen 2018 [254]	Folic acid	Colorectal cancer	4	31,192	346	Male and female population/ mostly with history of CVD (3x) or at high risk of CVD; partly with T2D	62.6-68.9 (mean)	Folic acid supplementation/ 2-2.5mg/day/ with vitamin B6 (50mg/day) and/or B12 (1mg/day)	Placebo	Colorectal cancer incidence	Parallel, factorial	3-7.3 years
Mocellin 2017 [255]	Vitamin B6	Cancer	9	34,911	2,539	Male and female population/ mostly with CVD (8x); partly with chronic renal failure (1x)	46.3-68.9 (mean)	Vitamin B6 supplementation/ 3-100mg/day/ partly in combination with vitamin B12 and folate; or high dose of micronutrients	Placebo (8x); low doses of micronutrients (1x)	Incidence of cancer (any site)	Parallel, factorial	2-7.3 (mean)

Morze 2021 [233]	Mediterranean diet	Cancer mortality	1	605	7	Male and female population/ mostly men/ with previous myocardial infarction	54 (mean)	Mediterranean diet/ instruction to follow a Mediterranean-type of diet	Control diet/ close to the step 1 American Heart Association prudent diet	Cancer mortality	Parallel	4 years
Morze 2021 [233]	Mediterranean diet	Breast cancer	1	4,152	35	Female population/ postmenopausal/ without history of cancer or CVD, but at risk of CVD	60-80 (range)	Mediterranean diet/ instruction to follow a Mediterranean-type of diet supplemented with nuts or extra-virgin olive oil	Control diet/ advice on a low-fat diet	Breast cancer	Parallel	4.8 years
Picasso 2019 [256]	Vegetarian diet	Triglycerides	5	689	N/A	Male and female population/ mostly female/ mostly overweight or obese/ partly with T2D (1x)	25.4-57.9 (mean)	Vegetarian diet/ dietary counselling to adhere to vegetarian diet/ emphasise consumption of plant foods like legumes, vegetables, grains and fruits, exclude meats/ partly excluding also animal derived foods (2x)	Omnivorous diet/ conventional diet, consuming meat, poultry or fish	Triglyceride levels (mmol/l)	Parallel, factorial	12-74 weeks
Picasso 2019 [256]	Vegetarian diet	HDL-cholesterol	4	596	N/A	Male and female population/ mostly female/ mostly overweight or obese/ partly with T2D (1x)	45.2-57.9 (mean)	Vegetarian diet/ dietary counselling to adhere to vegetarian diet/ emphasise consumption of plant foods like legumes, vegetables, grains and fruits, exclude meats/ partly excluding also animal derived foods (2x)	Omnivorous diet/ conventional diet, consuming meat, poultry or fish	HDL-cholesterol (mmol/l)	Parallel, factorial	12-74 weeks
Sayehmiri 2018 [257]	Selenium	Prostate cancer	9	45,638	23,994	Male population/ without history of prostate cancer/ partly at high risk of prostate cancer or with history of skin cancer or basal cell or squamous cell carcinoma	63-65.4 (mean); >55; >40	Selenium supplementation/ 200-400µg/day/ as selenised yeast or selenomethionine	Placebo	Prostate cancer	Parallel, factorial	NR
Schwingshackl 2015 [237]	Mediterranean diet	Type 2 diabetes	1	3,541	273	Male and female population/ without CVD, but with CVD risk factors	55-80 (range)	Highest adherence to a Mediterranean diet (in primary study: Intervention promoting the Mediterranean diet supplemented with olive oil or mixed nuts)	Lowest adherence to a Mediterranean diet (in primary study: advice on a low-fat diet)	Type 2 diabetes	Parallel	4.1 years
Schwingshackl 2017 [258]	Olive oil	Type 2 diabetes	1	2,301	181	Male and female population/ without T2D or CVD, but with CVD risk factors	55-80 (range); 67.2 (mean)	Highest olive oil intake (in primary study: Intervention promoting the Mediterranean diet supplemented with extra-virgin olive oil)	Lowest olive oil intake category (in primary study: advice on a low-fat diet)	Type 2 diabetes	Parallel	4.1 years

Setien-Suero 2016 [259]	Homocysteine	Mini mental state examination score	2	1,109	N/A	Male and female general population/ elderly	73.4-79.1 (mean)	Vitamin supplementation to reduce Homocysteine levels/ 1 tablet/day/ with folate (1mg) + B12 (0.5mg) + B6 (10mg) (1x); folate (0.4mg) + B12 (0.5mg) + D3 (0.015mg) (1x)	Placebo	Cognition assessed via mini mental state examination score	Parallel	2 years
Stratton 2011 [224]	Multivitamins	Prostate cancer	1	5,034	103	Male population/ generally healthy/ without severe health problems	51.3 (mean)	Multivitamin supplementation/ 1 tablet/day/ composition: 120mg vitamin C, 30mg α -tocopherol, 6mg β -carotene, 100 μ g selenium, 20mg zinc	Placebo	Prostate cancer incidence	Parallel	8.8 years (mean)
Stratton 2011 [224]	Vitamin E	Prostate cancer	3	39,318	1,969	Male population/ partly smokers/ partly with PSA <4.0ng/dl and normal rectal examination (1x)	50-69 (range); ≥ 50	Vitamin E supplementation/ 400IU/day (1x); 400IU every other day (1x); 50mg/day (α -tocopherol) (1x)	Placebo	Prostate cancer incidence	Factorial	5-8 years (mean)
Stratton 2011 [224]	Vitamin C	Prostate cancer	1	7,000	1,008	Male population/ partly with history of cancer or myocardial infarction or stroke	≥ 50	Vitamin C supplementation/ 500mg/day/ with Placebo vitamin E	Placebo	Prostate cancer incidence	Factorial	8 years (mean)
Stratton 2011 [224]	β -carotene	Prostate cancer	1	14,569	147	Male population/ generally healthy/ smokers (≥ 5 cigarettes/day)	50-69 (range)	β -carotene supplementation/ 20mg/day	Placebo	Prostate cancer incidence	Factorial	5-8 years/ 6.1 (median)
Te Morenga 2013 [260]	Low dietary sugars	Weight gain	10	382	N/A	Male and female population/ mostly healthy (7x); partly at risk of CVD (2x); with metabolic disease (1x)/ partly overweight	18-55 (range); NR (2x)	High-sugar diet with sugar-rich food, drinks or both/ sugar in foods: 80- >100g/day or 28% of total energy/ sugar sweetened beverages: 1l or 1,135g/day/ partly: with additional fibre depletion (1x); low fat (1x)	Diet with high amount of complex carbohydrates, artificial sweeteners or low to moderate sugars (10-40g/day); partly with fibre depletion (1x); low fat (1x)	Change in body weight (kg)	Parallel, crossover	1-6 months

Te Morenga 2013 [260]	Low dietary sugars	Body mass index	3	1,627	N/A	Male and female general population/ primary schoolchildren (2x); adolescents (1x)	7-18 (range)	Dietary intervention to reduce sugar intake/ counselling to reduce sugar-sweetened beverages, home deliveries of non-caloric beverages (1x); nutrition education to reduce fat and sugar intake and increase intake of complex carbohydrates (1x); behavioural classroom intervention to reduce intake of sugar sweetened beverages (1x)	General dietary advice/ nutrition education without specific advice regarding reduction of sugars or no intervention	Change in Body mass index (kg/m ²)	Parallel, cluster	6-8 months (duration)
Thorne-Lyman 2012 [225]	Vitamin D	Mean birthweight	5	932	N/A	Female general population/ pregnant/ without pre-existing diseases	18-45 (range); 25-27 (mean)	Vitamin D supplementation/ 400-4,000IU/day or single dose of 200,000IU or 2 doses of 600,000IU/ postconceptional	Nil (3x); placebo (1x); low dose (400IU/day) (1x)	Mean birthweight (g)	Parallel	<16 -28 weeks of gestation till delivery
Trikalinos 2012 [239]	Vitamin E	Cardiovascular disease	14	111,481	NR	Male and female population/ mostly with (history of) CVD (8x); partly at risk of CVD (3x); partly with T2D (1x); partly general population (2x)	55-69 (mean)	Vitamin E supplementation/ 50-1,200IU/day/ mostly 300-800IU/day (12x)	Placebo (8x); placebo and/or vitamin A (1x); placebo and/or vitamin C (1x); nil (3x); soybean oil (1x)	Cardiac or cardiovascular death (as defined in the primary studies)	Parallel, factorial	NR
Vinceti 2018a [7]	Selenium	Any cancer	5	21,860	2,332	Male and female population/ with diverse health status/ partly healthy, partly at high risk for prostate cancer, with resected non-small-cell lung cancer, BRCA1+ mutation carriers or history of skin cancer	62.3-66 (mean/ median); NR (1x)	Selenium supplementation/ 200-400µg/day as selenised yeast (2x); 200µg/day as L-selenomethionine (1x); 250µg/day as sodium selenite (1x)	Placebo	Incidence of any cancer	Parallel	3-13 years
Vinceti 2018a [7]	Selenium	Cancer mortality	2	18,698	359	Male and female population/ mostly male/ partly healthy, partly with history of skin cancer	62.3-63.4 (mean/ median)	Selenium supplementation/ 200µg/day as selenised yeast (1x); 200µg/day as L-selenomethionine (1x)	Placebo	Death from cancer	Parallel	8-13 years
Vinceti 2018b [236]	Selenium	Type 2 diabetes	5	22,053	1,603	Male and female population/ mostly men/ at risk of prostate cancer, with history of skin cancer, with resected non-small-cell lung cancer or removed colorectal adenomas	63-66 (mean)	Selenium supplementation/ 200µg/day as selenised yeast (4x); 200µg/day as L-selenomethionine (1x)	Placebo	Type 2 diabetes	Parallel	3-13 years (follow-up)

Wien 2012 [226]	Folic acid	Cancer	10	38,233	3,515	Male and female population/ mostly with CVD or at high risk of CVD; partly with history of colorectal adenoma or chronic renal disease	57-69 (mean)	Folic acid supplementation/ $\geq 0.4\text{mg/day}$ / alone (1x); with aspirin (2x); vitamin B12 + B6 (6x); vitamin B12 (1x)	Placebo (9x); low dose (20 $\mu\text{g/day}$; 1x)/ alone (5x); with aspirin (2x); vitamins B12 + B6 (1x); vitamin B6 (2x)	Total cancer incidence	Parallel, factorial	1.7-7.3 years (mean)
Wolf 2017 [8]	Multivitamins	Preterm birth	1	353	17	Female population/ pregnant/ ≤ 13 weeks of gestation/ high-income country, low-income multi-ethnic population	28 (mean)	Multivitamin and mineral supplementation/ postconceptional	Placebo	Preterm birth/ delivery <37 weeks of gestational age	Parallel	First trimester till delivery
Wolf 2017 [8]	Multivitamins	Stillbirth	1	261	2	Female population/ with previous neural tube defect affected pregnancy/ planning to have another child/ in high-income country	31.3 (mean)	Multivitamin supplementation/ 3 tablets/day/ pre- and periconceptional (at least two months before conception and until the date of the third missed period)	Folic acid supplementation/ 360 $\mu\text{g/day}$	Stillbirth	Parallel	>2 month pre-conception till delivery
Yang 2016 [235]	Folic acid	Gestational hypertension	1	1,078	89	Female population/ pregnant/ HIV-infected/ low-income country	24.7 (mean)	Folic acid supplementation/ postconceptional/ with other multivitamins	Placebo	Gestational hypertension	Factorial	Till 3 months postpartum
Yang 2020 [227]	Calcium	Cardiovascular disease	4	3,996	NR	Male and female population/ generally healthy	58-75.1 (mean); ≥ 55	Calcium supplementation/ 1,000-1,200 mg/day / as calcium carbonate (1x); elemental calcium (2x); lactogluconatecarbonate (100 g/day , 1x)	Placebo	Cardiovascular disease incidence and mortality	Parallel	4-5 years
Yao 2019 [228]	Vitamin D	Any fracture	11	34,243	2,843	Male and female population/ partly institutionalised (3x) or with previous fractures (5x)	61.8-85 (mean)	Vitamin D supplementation/ 400-833IU/day (3x); 20,000-100,000IU/week (2x); multiple bolus of 100,000-500,000IU (6x)/ oral (10x); i.m. injection (1x)	Placebo (10x); nil (1x)	Any fracture/ fractures that occurred at any site	Parallel, factorial	0.8-5 years
Yu 2021 [229]	Folic acid	Pre-eclampsia	3	5,108	441	Female population/ pregnant/ partly at high risk of pre-eclampsia (2x)	26-31 (mean)	Folic acid supplementation/ 4-5 mg/day / postconceptional	Placebo/ partly with low dose (200 $\mu\text{g/day}$; 1x)	Pre-eclampsia	Parallel	1-4 years
Zhang 2016 [9]	Selenium	Cardiovascular disease	9	36,511	NR	Male and female population/ mostly generally healthy (6x); partly with CVD (3x)	46.6-76.2 (mean)	Selenium supplementation/ 100-200 $\mu\text{g/day}$ / alone (3x); combined with other antioxidants (6x)	Placebo	Cardiovascular disease incidence and mortality	Parallel, factorial	0.25-12 years

Zhang 2018 [230]	Vitamin D	Gestational diabetes	4	968	87	Female population/ pregnant/ partly healthy (2x); partly with Vitamin D insufficiency (2x)	25.2-29.5 (mean); NR (1x)	Vitamin D supplementation/ 4,000-5,000IU/day (2x); 50,000IU/2 weeks (1x); 60,000-480,000IU total depending on 25(OH)D level (1x) mostly alone (3x); partly with 200mg/day ferrous sulfate and 600mg/day calcium (1x)	Low dose (400IU/day; 2x); nil (1x); routine care with 200mg/day ferrous sulfate and 600mg/day calcium (1x)	Gestational diabetes mellitus (hyperglycaemia first recognised at any stage of pregnancy)	Parallel	12/ <20weeks of gestation to 34-36 weeks of gestation or delivery
Zhang 2018 [230]	Vitamin D	Fasting plasma glucose	16	NR	N/A	Female population/ pregnant/ mostly with gestational diabetes (13x); partly at risk of pre-eclampsia (3x) or with Vitamin D insufficiency (2x)	NR	Vitamin D supplementation/ 10-5,000IU/day or 50,000IU/2 weeks or 50,000IU/month or 150,000-700,000IU total/ alone (14x); via primrose oil (1x); with calcium (1x)/ oral substitution (12x); intramuscular injection (4x)	Placebo (5x); nil (3x); low dose (400IU/day; 2x); low dose vitamin D + insulin (2x); insulin (2x); dietary intervention (1x); yogurt (1x)	Fasting plasma glucose level (difference)	NR	4-16 weeks (8x); 12 weeks of gestation to 34-36 weeks of gestation or delivery (3x); once (4x);NR(1x)
Zhao 2014 [261]	Multivitamins/ Minerals	Nuclear cataract	2	3,161	362	Male and female general population/ partly at high risk of chronic nutrient deficiencies or with early cataract	45-75 (range)	Multivitamin and mineral supplementation	Placebo	Nuclear cataract incidence	Parallel, factorial	5-9 years
Zhao 2014 [261]	Multivitamins/ Minerals	Cortical cataract	2	3,161	960	Male and female general population/ partly at high risk of chronic nutrient deficiencies or with early cataract	45-75 (range)	Multivitamin and mineral supplementation	Placebo	Cortical cataract incidence	Parallel, factorial	5-9 years
Zhao 2014 [261]	Multivitamins/ Minerals	Posterior subcapsular cataract	2	3,161	156	Male and female general population/ partly at high risk of chronic nutrient deficiencies or with early cataract	45-75 (range)	Multivitamin and mineral supplementation	Placebo	Posterior subcapsular cataract incidence	Parallel, factorial	5-9 years
Zhou 2017 [232]	Vitamin D	Preterm birth	5	1,633	74	Female population/ pregnant/ generally healthy/ without HIV infection/ partly with gestational diabetes (1x)	25.2-31.1 (mean); NR (1x)	Vitamin D supplementation/ 1,000-4,000IU/day or multiple bolus dosage (60,000-480,000IU in 1-4 doses)	Placebo (4x); routine antenatal care (200mg/day ferrous sulfate + 600mg/day calcium) (1x)	Delivery of a liveborn neonate before 37 weeks of gestation	Parallel	14-28 weeks of gestation till delivery

BoE=bodies of evidence; CHD=coronary heart disease; CVD=cardiovascular disease; DHA=docosahexaenoic acid; EPA=eicosapentaenoic acid; HDL=high density lipoprotein; HIV=human immunodeficiency virus; IQR=interquartile range; N/A=not applicable; NR=not reported; PSA=prostate-specific antigen; SF=saturated fatty acids; T2D=type 2 diabetes mellitus; UV=ultraviolet

Supplementary Table S7 Characteristics of included BoE from cohort studies

Reference	Exposure	Outcome	Studies, n	Sample size, n	Cases, n	Description of population	Age (years)	Description of exposure	Description of comparator	Description of outcome	Study design	Study length/ follow-up
Aburto 2013 [2]	Low sodium	All-cause mortality	2	18,184	NR	Male and female general population	24-74 (range); >30	High sodium intake	Low sodium intake	All-cause mortality	Prospective cohort	8.7-22 years
Aburto 2013 [2]	Low sodium	Cardiovascular disease	3	76,914	NR	Male and female general population	25-79 (range); >30	High sodium intake	Low sodium intake	Cardiovascular disease incidence and mortality	Prospective cohort	8.7-22 years
Afshin 2014 [240]	Nuts	Coronary heart disease	6	284,683	8,712	Male and female general population/ mostly healthy; partly adults at high risk of CVD (1x)	34-84 (range); 25+	Nut consumption/ dose-response: per 4 weekly servings (28.4g) of nuts	Dose-response	Total incident ischemic heart disease (fatal and nonfatal)	Cohort	4.8-30 years
Afshin 2014 [240]	Nuts	Stroke	2	148,238	5,454	Male and female population/ generally healthy/ healthcare professionals	30-86 (range)	Nut consumption/ dose-response: per 4 weekly servings (28.4g) of nuts	Dose-response	Total stroke (ischemic and haemorrhagic)	Cohort	21.1-26 years
Afshin 2014 [240]	Nuts	Type 2 diabetes	4	226,675	13,035	Male and female population/ generally healthy	35-87 (range)	Nut consumption/ dose-response: per 4 weekly servings (28.4g) of nuts	Dose-response	Type 2 diabetes	Cohort	4.6-23 years
Aguilar-Cordero 2020 [218]	Vitamin D	Pre-eclampsia	7	11,902	4,658	Pregnant women/ general healthy without specific previous pathologies	NR	Low vitamin D status/ deficiency/ 25(OH)D level in maternal blood <50nmol/l	Normal vitamin D status/ 25(OH)D level in maternal blood >50nmol/l	Gestational hypertension with proteinuria and/or other maternal organ dysfunction and/or utero-placental dysfunction/ ≥20 weeks of gestation	Cohort, nested case-control	10-36.9 weeks of gestation till delivery
Alexander 2017 [219]	Omega-3 fatty acids	Coronary heart disease	17	687,166	NR	Male and female general population (6x male; 3x female; 8x both)/ mostly generally healthy (16x); partly with coronary artery disease (1x)	>18	High intake of omega-3 fatty acids/ EPA and DHA/ ≥0.15-≥0.62g/day EPA and 0.22-5.18g/day DHA	Low intake of omega-3 fatty acids/ <0.04 - ≤0.39g/day EPA and DHA ≤1.48g/day	Fatal or nonfatal myocardial infarction, CHD incidence, coronary death, sudden cardiac death, angina	Cohort	4.8-40 years
Amegah 2017 [231]	Vitamin D	Gestational length	1	401	N/A	Pregnant women/ with singleton pregnancy before 16 weeks of gestation	29.3 (mean)	Low vitamin D status/ insufficiency/ 25(OH)D level in maternal blood <28nmol/l at 28-32 weeks of gestation	Normal vitamin D status/ 25(OH)D level in maternal blood ≥28nmol/l	Gestational length (weeks)	Prospective cohort	1.5 years/ <16 weeks of gestation till delivery

Azad 2017 [220]	Nonnutritive sweeteners	Body mass index	1	2,222	N/A	Male and female general population (58% female)/ mean body mass index at baseline: 27	44 (mean)	High intake of nonnutritive sweeteners/ ≥ 22 servings/week of artificially sweetened beverages (including sodas and other beverages, e.g. coffee, tea)	Low intake of nonnutritive sweeteners intake/ nonusers	Change in body mass index (kg/m ²)	Prospective cohort	8 years
Bolland 2015 [221]	Calcium	All fractures	5	116,199	11,194	Male and female general population/ mostly female	50-79 (range); 56-77 (mean)	Calcium supplementation/ partly with vitamin D (1x)	No calcium supplementation	Fractures at any site	Prospective cohort	3-8.4 years
Chowdhury 2012 [241]	Omega-3 fatty acids	Cerebrovascular disease	10	301,023	4,197	Male and female general population/ healthy	16-84 (range)	High omega-3 fatty acid intake/ 0.11-2.6g/day	Low omega-3 fatty acid intake	Fatal or non-fatal ischaemic stroke, haemorrhagic stroke, cerebrovascular accident, transient ischaemic attack	Prospective cohort	4-28 years
Chowdhury 2014a [234]	α -linolenic acid	Coronary disease	7	157,258	7,431	Male and female population/ mostly healthy (6x); partly population with high CVD risk (1x)	30-84 (range)	High α -linolenic acid intake	Low α -linolenic acid intake	Fatal or nonfatal myocardial infarction, CHD, coronary insufficiency, coronary death, angina, angiographic coronary stenosis	Prospective cohort	5-23 years
Chowdhury 2014a [234]	Omega-6 fatty acids	Coronary disease	8	206,376	8,155	Male and female population/ mostly generally healthy (7x); partly with pre-existing high CVD risk (1x)	20-75 (range)	High omega-6 fatty acid intake	Low omega-6 fatty acid intake	Fatal or nonfatal myocardial infarction, CHD, coronary insufficiency, coronary death, angina, angiographic coronary stenosis	Prospective cohort	5-23 years
Chowdhury 2014b [3]	Vitamin D	All-cause mortality	68	840,908	64,636	Male and female population/ with or without pre-existing chronic disease	29.2-80 (range); 63-63.4 (mean/ median)	Low vitamin D status/ 25(OH)D level <10-29ng/ml in serum or plasma	High vitamin D status/ 25(OH)D level ≥ 30 ng/ml in serum or plasma	Deaths from any cause	Cohort	0.3-29 years
Chung 2011 [242]	Vitamin D	Colorectal cancer	9	7,402	3,136	Male and female general population/ partly smokers (2x)	40-79 (range); 57-69.2 (mean)	Vitamin D status/ dose-response per 10nmol/l increase in 25(OH)D concentration	Dose-response	Colorectal cancer incidence and mortality	Nested case-control	7 months- 17 years
Chung 2011 [242]	Vitamin D	Breast cancer	4	5,995	2,986	Female population/ partly postmenopausal	30-85 (range); 57-70 (mean)	Vitamin D status/ dose-response per 10nmol/l increase in 25(OH)D concentration	Dose-response	Breast cancer incidence and/or death	Nested case-control	3.9-15 years
Ding 2017 [243]	Dairy	Systolic blood pressure	27	171,426	N/A	Male and female general population	1-84 (range); NR (3x)	Dairy intake/ dose-response per serving/day increase	Dose-response	Systolic blood pressure (mmHg)	Prospective cohort	2-27 years

Feng 2015 [4]	Folic acid	Congenital heart defects	1	6,112	81	Female population/ women in childbearing age	27.4 (mean)	Folic acid supplementation/ 0.6mg/day/ periconceptional: 1 months before through 3 months after conception/ combined with other multivitamins/ minerals	No folic acid supplementation/ non-user	Congenital heart defects/ overall or any one of the subtypes in infants	Matched cohort	1 month before conception till delivery
Filippini 2020 [244]	Green Tea	Prostate cancer	5	122,110	1,179	Male population/ without history of cancer (2x); without history of prostate cancer (2x); atomic-bomb survivors without prostate cancer (1x)	40-79 (range); NR (2x)	High green tee intake	Low green tea intake/ <1 cup or time/day	Prostate cancer	Prospective cohort	8-21 years
Filippini 2020 [244]	Green Tea	Endometrial cancer	1	53,724	117	Female general population/ without history of cancer	40-69 (range)	Highest green tee exposure/ ≥5 cups/day	Lowest green tea exposure/ ≤4 cups/week	Endometrial cancer	Prospective cohort	5 years
Fu 2021 [245]	Vitamin C	Lung cancer	3	144,263	1,406	Male and female general population/ partly elderly (1x)	50-84 (range)	Vitamin C supplementation/ >66.6- >322mg/day or 500mg/day	No vitamin C supplementation/ non-users	Lung cancer incidence	Prospective cohort	4.05-10.6 years
Fu 2021 [245]	Vitamin C	Breast cancer	6	215,581	8,050	Female population/ postmenopausal (3x)	40-84 (range)	Vitamin C supplementation/ >64mg/day - >1,000IU/day or ever supplement use (1x)	No vitamin C supplementation/ non-users	Breast cancer incidence	Prospective cohort, nested case-control	5-10.6 years
Fu 2021 [245]	Vitamin C	Colorectal cancer	2	50,700	246	Male and female general population/ partly residents of a retirement community (1x)	57.1 (mean); NR (1x)	Vitamin C supplementation/ ≥360mg/day (1x)	No vitamin C supplementation/ non-users	Colorectal cancer incidence	Prospective cohort	4.5-7 years
Gayer 2019 [246]	Apples	Body weight	1	133,468	N/A	Male and female population/ healthy or at risk for CVD or cardio-metabolic disease/ without T2D or existing CVD	41.65 (mean)	Increased intake of apple per servings/day	Dose-response, per serving/day	Body weight change (kg)	Prospective cohort	4 years
Grosso 2015 [222]	Mediterranean diet	Cardiovascular disease	20	888,257	22,987	Male and female population/ partly with previous coronary artery disease (1x); with diabetes (1x); with cardiovascular disease (1x)	20-90 (range); 38-68.6 (mean)	High adherence to Mediterranean diet	Low adherence to Mediterranean diet	CVD incidence (including deaths)	Prospective cohort	2-40 years
Jiang 2019 [5]	Vitamin E	Age-related cataract	6	42,147	NR	Male and female population/ mostly female/ without (ascertained) cataract	43-84 (range); ≥43; ≥49	High vitamin E intake/ 8.4-90.8mg/day	Low vitamin E intake/ 3-6.7mg/day	Age-related cataract	Longitudinal and non-longitudinal cohort	5-15 years
Jiang 2019 [5]	β-carotene	Age-related cataract	7	154,449	NR	Male and female population/ without (ascertained) cataract	43-84 (range); ≥45; ≥49	High β-carotene intake/ 1.94-12.11mg/day	Low β-carotene intake/ 0.47-3mg/day	Age-related cataract	Longitudinal and non-longitudinal cohort	5-15 years

Johnston 2019 [247]	Low red meat	Cardiovascular disease	25	858,554	NR	Male and female population/ generally healthy	33-74.1 (mean)	Low adherence to diet rich in red and processed meat or non-vegetarian diet	High adherence to diet rich in red and processed meat or non-vegetarian diet	Cardiovascular mortality	Prospective cohort	4-26 years
Johnston 2019 [247]	Low red meat	Type 2 diabetes	14	378,788	NR	Male and female population/ generally healthy	36-61.7 (mean); NR (3x)	Low adherence to diet rich in red and processed meat or non-vegetarian diet	Higher adherence to diet rich in red and processed meat or non-vegetarian diet	Type 2 diabetes incidence	Cohort	2-34 years
Johnston 2019 [247]	Low red meat	Colorectal cancer	16	840,980	NR	Male and female population/ generally healthy	NR	Low adherence to diet rich in red and processed meat or non-vegetarian diet	Higher adherence to diet rich in red and processed meat or non-vegetarian diet	Colorectal cancer incidence	Prospective cohort	5-26 years
Jonker 2020 [248]	Folic acid	Mean birthweight	2	2,892	N/A	Female population/ pregnant/ in low- and middle-income countries	NR	Folic acid supplementation/ alone (1x); with iron (1x)	No folic acid supplementation	Mean birthweight (kg)	Cohort	NR
Jonker 2020 [248]	Folic acid	Low birthweight	5	26,270	5,114	Female population/ pregnant/ in low- and middle-income countries	NR	Folic acid supplementation/ mostly with iron supplementation (4x)	No folic acid supplementation	Incidence of low birthweight	Cohort	NR
Jonker 2020 [248]	Folic acid	Small for gestational age	4	239,862	23,666	Female population/ pregnant/ in low- and middle-income countries	NR	Folic acid supplementation/ alone (2x); with iron (1x); with dietary folate intake (1x)	No folic acid supplementation/ non-users	Infants whose weight is <10th percentile for gestational age	Cohort	NR
Kastorini 2011 [6]	Mediterranean diet	Metabolic syndrome	2	4,495	NR	Male and female population/ without CVD risk factors (1x); without diabetes mellitus or metabolic syndrome (1x)	20-90 (range); 54 (mean)	High adherence to Mediterranean diet	Low adherence to Mediterranean diet	Metabolic syndrome	Prospective cohort	6.2-7 years
Kastorini 2011 [6]	Mediterranean diet	HDL-cholesterol	1	2,563	N/A	Male and female population/ without CVD risk factors	20-90 (range)	High adherence to Mediterranean diet	Low adherence to Mediterranean diet	HDL Cholesterol (mg/dl)	Prospective cohort	6.2 years
Kastorini 2011 [6]	Mediterranean diet	Triglycerides	1	2,563	N/A	Male and female population/ without CVD risk factors	20-90 (range)	High adherence to Mediterranean diet	Low adherence to Mediterranean diet	Triglycerides (mg/dl)	Prospective cohort	6.2 years
Kastorini 2011 [6]	Mediterranean diet	Systolic blood pressure	1	2,563	N/A	Male and female population/ without CVD risk factors	20-90 (range)	High adherence to Mediterranean diet	Low adherence to Mediterranean diet	Systolic blood pressure (mmHg)	Prospective cohort	6.2 years
Kim 2018 [223]	Multivitamins/ Minerals	Cardiovascular disease	9	587,638	NR	Male and female general population/ without history of CVD and cancer; without medical conditions with a predicted survival of ≤3 years (1x)	50.6-63.9 (mean)	Multivitamin/ Mineral supplementation/ >3 vitamin and mineral ingredients	No multivitamin/Mineral supplementation/ non-users	CHD mortality and stroke mortality	Prospective cohort	5-19 years

Kim 2018 [223]	Multivitamins/ Minerals	Coronary heart disease	6	370,085	NR	Male and female general population/ without history of CVD and cancer; without medical conditions with a predicted survival of ≤ 3 years (1x)	30-55y (range); 50-63.9 (mean)	Multivitamin/ Mineral supplementation/ >3 vitamin and mineral ingredients	No multivitamin/ Mineral supplementation/ non-users	Incident CHD events/ cardiac revascularization and fatal and nonfatal myocardial infarction	Prospective cohort	7.9-16.2 years
Kim 2018 [223]	Multivitamins/ Minerals	Stroke	3	217,529	NR	Male and female general population/ without history of CVD and cancer (2x); without medical conditions with a predicted survival of ≤ 3 years (1x)	52.9-63.9 (mean)	Multivitamin/ Mineral supplementation/ >3 vitamin and mineral ingredients	No multivitamin/ Mineral supplementation/ non-users	Incident ischemic and haemorrhagic stroke (fatal and nonfatal)	Prospective cohort	7.9-16.2 years
Kong 2014 [249]	Vitamins	Gastric cancer	7	1,037,967	1,606	Male and female population	40-83 (range); >40	High vitamin intake/ in different compositions: ACE + folate (2x); ACE (1x); A + folate (1x); E (1x); A (1x); NR (1x)	Low vitamin intake	Gastric cancer occurrence	Prospective cohort	6.3-14 years
Lin 2019 [250]	Vitamin E	Bladder cancer	8	497,011	2,774	Male and female general population	30-84 (range)	High vitamin E intake	Low vitamin E intake	Bladder cancer incidence	Prospective cohort	6-20 years
Lin 2020 [251]	Vitamin D	Nephrolithiasis	3	192,126	6,449	Male and female population/ without prior nephrolithiasis	25-75 (range)	High vitamin D supplementation	Low vitamin D supplementation	Nephrolithiasis incidence	Prospective cohort	11.3-12.1 years
Lin 2020 [251]	Calcium	Nephrolithiasis	2	187,976	2,087	Female population/ without prior nephrolithiasis	25-55 (range)	High calcium supplementation	Low calcium supplementation	Nephrolithiasis incidence	Prospective cohort	8-12 years
Martinez-Gonzalez 2014b [252]	Olive oil	Coronary heart disease	3	93,760	1,367	Male and female population/ healthy (2x); without CVD or cancer (1x)	20-86 (range); 49-50 (median)	Olive oil intake/ dose-response per increase of 25g/day	Dose-response	CHD events (fatal and non-fatal)/ acute myocardial infarction, revascularisation, angina and other	Prospective cohort	7.85-10.4 years
Martinez-Gonzalez 2014b [252]	Olive oil	Stroke	2	31,226	543	Male and female population/ without history of stroke (1x); free of CVD and cancer (1x)	73.8 (mean); $<55 = 58\%$, $55-64 = 23\%$, $\geq 65 = 19\%$	Olive oil intake/ dose-response per increase of 25g/day	Dose-response	Stroke (ischaemic or haemorrhagic)	Prospective cohort	5.25-10.6 years
Miller 2014 [253]	Low calorie sweeteners	Body weight	4	85,353	N/A	Male and female population/ mostly female/ generally healthy	3x adults: 18-64 (range); 1x preschool children: 2-5 (range)	Low calorie sweeteners intake/ saccharin (2x); low-calorie sweetened beverages (2x)	Mean correlation	Body weight change (kg)	Prospective cohort	0.5-4 years
Moazzen 2018 [254]	Folic acid	Colorectal cancer	3	1,926,520	10,514	Male and female general population/ without colorectal cancer	50-64.5 (mean)	High folic acid supplementation/ $>800-939\mu\text{g/day}$	Low folic acid supplementation/ $<242-300\mu\text{g/day}$ (2x)	Colorectal cancer or adenoma incidence	Cohort	9.1-16 years

Mocellin 2017 [255]	Vitamin B6	Cancer	29	NR	10,825	Male and female general population	NR	High vitamin B6 intake	Low vitamin B6 intake	Incidence of cancer (any site)	Prospective cohort, nested case-control	NR
Morze 2021 [233]	Mediterranean diet	Cancer mortality	18	1,660,652	71,138	Male and female population	20-90 (range); 38 (mean)	High adherence to Mediterranean diet	Low adherence to Mediterranean diet	Cancer mortality	Prospective cohort	6.25-40 years
Morze 2021 [233]	Mediterranean diet	Breast cancer	12	982,723	35,338	Male and female population/ mostly women	18-104 (range)	High adherence to Mediterranean diet:	Low adherence to Mediterranean diet	Breast cancer	Prospective cohort	3-18 years
Picasso 2019 [256]	Vegetarian diet	Triglycerides	1	302	N/A	Population in inpatient rehabilitation centre/ with cardiovascular disease/ partly overweight	54.5 (mean)	Adherence to lacto-ovo-vegetarian diet	Adherence to omnivorous diet	Triglyceride levels (mmol/l)	Prospective cohort	NR
Picasso 2019 [256]	Vegetarian diet	HDL-cholesterol	1	302	N/A	Population in inpatient rehabilitation centre/ with cardiovascular disease/ partly overweight	54.5 (mean)	Adherence to lacto-ovo-vegetarian diet	Adherence to omnivorous diet	HDL-cholesterol (mmol/l)	Prospective cohort	NR
Sayehmiri 2018 [257]	Selenium	Prostate cancer	2	45,370	672	Male population/ without history of prostate cancer (1x); smokers without history of cancer(1x)	50-76 (range)	High selenium intake and supplementation/ >50 >111.05µg/day	Low selenium intake/ nil (1x); <71.52µg/day (1x)	Prostate cancer incidence	Prospective cohort, trial-based cohort	9-10 years
Schwingshackl 2015 [237]	Mediterranean diet	Type 2 diabetes	8	119,269	NR	Male and female population/ mostly without CVD (6x); partly with myocardial infarction (1x) or prior gestational diabetes mellitus (1x)	20-90 (range)	High adherence to Mediterranean diet	Low adherence to Mediterranean diet	Type 2 diabetes	Prospective cohort	3.2-20 years
Schwingshackl 2017 [258]	Olive oil	Type 2 diabetes	4	183,370	21,688	Male and female general population/ without T2D	36-55 (mean); <50	High olive oil intake	Low olive oil intake	Type 2 diabetes	Prospective cohort	5.7-22 years
Setien-Suero 2016 [259]	Homocysteine	Mini mental state examination score	3	1,625	N/A	Male and female general population/ elderly	67-75.4 (mean)	Homocysteine status	Linear regression	Cognition/ Mini mental state examination score	Prospective cohort	3-7 years
Stratton 2011 [224]	Multivitamins	Prostate cancer	1	295,344	1,331	Male general population/ without cancer	50-71 (range)	Multivitamin supplementation	No multivitamin supplementation/ non-users	Prostate cancer incidence	Prospective cohort	5 years
Stratton 2011 [224]	Vitamin E	Prostate cancer	5	457,136	17,541	Male general population	40-76 (range)	Vitamin E supplementation	No vitamin E supplementation	Prostate cancer incidence	Prospective cohort	4-10 years
Stratton 2011 [224]	Vitamin C	Prostate cancer	1	29,361	1,338	Male population/ without history of prostate, colon, or lung cancer	55-74 (range)	Vitamin C supplementation	No vitamin C supplementation	Prostate cancer incidence	Prospective cohort	4.2 years

Stratton 2011 [224]	β-carotene	Prostate cancer	1	25,016	1,338	Male population/ without history of prostate, colon, or lung cancer	55-74 (range)	β-carotene supplementation	No β-carotene supplementation	Prostate cancer incidence	Prospective cohort	4.2 years
Te Morenga 2013 [260]	Low dietary sugars	Weight gain	4	149,305	N/A	Male and female population/ generally healthy/ younger (university graduates, nurses) and older adults/ without: diabetes (3x); severe diseases (3x)	21-74 (range)	High intake of sugar sweetened beverages/ including: sugar sweetened soft drinks, fruit drinks and fruit juices	Low intake of sugar sweetened beverages	Weight difference (kg)	Prospective cohort	4-6 years
Te Morenga 2013 [260]	Low dietary sugars	Body mass index	4	4,094	N/A	Male and female general population/ children and adolescents	2-14 (range)	Intake of dietary sugars/ dose-response per 1 serving/day or 100g/day intake increase of sugar sweetened beverages (soft drinks, fruit juice)	Dose-response	Change in body mass index (kg/m ²)	Prospective cohort, prospective cohort nested in RCT	1-10 years
Thorne-Lyman 2012 [225]	Vitamin D	Mean birthweight	2	1,497	N/A	Pregnant woman and their children/ without serious non-obstetric problems (1x); without multiple gestations or gestational age >22 weeks at enrolment (1x)	22-32.5 (mean)	Highest categories of vitamin D intake/ including dietary intake and supplementation: >658IU (1x); >535IU/day (1x)	Lowest categories of vitamin D intake/ <446IU (1x); <185IU/day (1x)	Mean birthweight (g)	Prospective cohort	NR (till birth)
Trikalinos 2012 [239]	Vitamin E	Cardiovascular disease	8	137,237	NR	Male and female population/ with or without history of cardiovascular disease	30-93 (range); 50-88 (mean)	High vitamin E intake	Lowest intake category of vitamin E	Cardiac or cardiovascular death (as defined in the primary studies)	Prospective cohort	6-15 years
Vinceti 2018a [7]	Selenium	Any cancer	7	76,239	1,940	Male and female general population/ without cancer	20-74 (range); ≥15; NR (2x)	High selenium exposure/ mostly in serum (6x); partly in plasma and serum (1x)/ 60-136µg/l (1x) or ≥78-240µg/l	Low selenium exposure/ ≤48.9-≤142µg/l	Incidence of any cancer	Nested case-control, cohort-subcohort controlled	5-19 years
Vinceti 2018a [7]	Selenium	Cancer mortality	1	133,957	2,603	Male and female general population/ without history of cancer	40-74 (range)	High selenium intake/ men: ≥31.92µg per 1,000kcal/day; women: ≥33.36µg per 1,000kcal/day	Low selenium intake/ men: <19.36µg/1,000; women: <19.05µg/1,000kcal/day	Cancer mortality	Prospective cohort	8.4-13.9 years
Vinceti 2018b [236]	Selenium	Type 2 diabetes	1	7,182	253	Female population/ healthy without history of cancer or T2D	35-70 (range)	High dietary selenium intake/ >65.9µg/day	Low selenium exposure/ ≤47µg/day	Type 2 diabetes	Cohort	16 years
Wien 2012 [226]	Folic acid	Cancer	5	372,799	2,547	Male and female general population	51.7-67 (mean)	Folic acid supplementation/ ≥0.4mg/day	No or low folic acid supplementation/ <0.2mg for women or <0.3mg for men (1x)	Breast (2x); pancreatic (2x) or lung (1x) cancer incidence	Prospective cohort	4-14 years
Wolf 2017 [8]	Multivitamins	Preterm birth	4	42,592	2,280	Female population/ pregnant/ in high-income countries	NR	Multivitamin supplementation/ ≥3 vitamins or minerals/ peri- and postconceptional	No multivitamin use	Preterm birth/ delivery <37 weeks of gestational age	Prospective cohort, matched cohort	3-10 years

Wolf 2017 [8]	Multivitamins	Stillbirth	2	39,845	208	Female population/ pregnant/ in high-income countries	NR	Multivitamin supplementation/ ≥3 vitamins or minerals/ periconceptual	No vitamin use	Stillbirth: fetal death at gestational age ≥28 weeks and/or fetus weigh >1,000g (1x); fetal death at gestational age >20 weeks (1x)	Prospective cohort, matched cohort	3-6 years
Yang 2016 [235]	Folic acid	Gestational hypertension	3	201,265	18,894	Female general population/ pregnant (2x); preparing for marriage or becoming pregnant (1x)	24.9-31.2 (mean)	Folic acid supplementation/ periconceptual/ partly with folate intake	No or low folic acid supplementation (2x) or intake (1x)	Gestational hypertension	Prospective cohort	NR (till delivery)
Yang 2020 [227]	Calcium	Cardiovascular disease	13	823,293	30,117	Male and female general population	17-79 (range)	High calcium intake	Low calcium intake	Cardiovascular disease incidence and mortality	Prospective cohort	9-28 years
Yao 2019 [228]	Vitamin D	Any fracture	10	39,141	6,278	Male and female population/ mostly female/ not living in institutions/ baseline blood 25(OH)D level: 53.5 - 81.0nmol/l	52.6-76.7 (mean) or 68.6 (mean of means)	Vitamin D status/ 25(OH)D level in blood	Dose-response	Any fracture/ fractures at any site	Prospective cohort, nested case-control, case-cohort	5-15 years
Yu 2021 [229]	Folic acid	Pre-eclampsia	13	154,268	NR	Female population/ pregnant/ generally healthy	14-44 (range); 27.8-33.1 (mean); or >18	Folic acid supplementation/ 0.4mg/day/ alone (12x); combined with dietary intake (1x)/ pre- or postconceptional	No folic acid supplementation	Pre-eclampsia	Prospective cohort	2-6 years
Zhang 2016 [9]	Selenium	Cardiovascular disease	14	34,109	3,749	Male and female general population	20-90 (range); ≥35; ≥65; 48	High selenium status/ 101.5µg/l (median)/ serum (11x); plasma (1x); erythrocyte (1x)	Low selenium status/ 53.7µg/l	Cardiovascular disease incidence or mortality	Prospective cohort, nested case-control studies	3-15 years
Zhang 2018 [230]	Vitamin D	Gestational diabetes	24	NR	NR	Female population/ pregnant/ prevalence of Vitamin D sufficiency in populations 3.3 - 73.33% (range); NR (4x)	NR	Low vitamin D status/ 25(OH)D level in blood 25- 50nmol/l and/or 50 - 75nmol/l	Normal vitamin D status/ 25(OH)D level in blood >50 or 75nmol/l	Gestational diabetes mellitus (hyperglycaemia first recognised at any stage of pregnancy	Cohort, nested case- control	NR
Zhang 2018 [230]	Vitamin D	Fasting plasma glucose	13	NR	N/A	Female population/ pregnant/ prevalence of Vitamin D sufficiency in populations 5.8 - 58.7% (range); NR (3x)	NR	Vitamin D status/ 25(OH)D level in blood	Correlation	Fasting plasma glucose level	Cohort, nested case- control	NR
Zhao 2014 [261]	Multivitamins/ Minerals	Nuclear cataract	5	13,637	NR	Male and female population/ general (2x); clinic-based population (2x); nondiabetic women (1x)	43-97 (range); ≥40; ≥45	Multivitamin/ Mineral supplementation/ longest duration	No multivitamin/ Mineral supplementation	Nuclear, age-related cataract incidence	Prospective cohort	4.8-15 years

Zhao 2014 [261]	Multivitamins/ Minerals	Cortical cataract	4	12,887	NR	Male and female population/ general (2x); clinic-based population (1x); nondiabetic women (1x)	43-97 (range); ≥45	Multivitamin/ Mineral supplementation/ longest duration	No multivitamin/ Mineral supplementation	Cortical, age- related cataract incidence	Prospective cohort	6.3-15 years
Zhao 2014 [261]	Multivitamins/ Minerals	Posterior subcapsular cataract	3	8,772	NR	Male and female population/ general (1x); clinic-based population (1x); nondiabetic women (1x)	43-86 (range); ≥45	Multivitamin/ Mineral supplementation/ longest duration	No multivitamin/ Mineral supplementation	Posterior subcapsular, age- related cataract incidence	Prospective cohort	5-15 years
Zhou 2017 [232]	Vitamin D	Preterm birth	12	15,184	2,347	Female population/ pregnant women/ without HIV infection	NR	Low vitamin D status/ 25(OH)D level <50nmol/l	High vitamin D status/ 25(OH)D level >50nmol/l	Delivery of a liveborn neonate before 37 weeks of gestation	Prospective cohort, case- cohort, nested case-control	NR (till birth)

BoE=bodies of evidence; CHD=coronary heart disease; CVD=cardiovascular disease; DHA=docosahexaenoic acid; EPA=eicosapentaenoic acid; HDL=high density lipoprotein; HIV=human immunodeficiency virus; N/A=not applicable; NR=not reported; T2D=type 2 diabetes mellitus

Supplementary Table S8 Methodological quality assessment of the included systematic reviews (AMSTAR 2)

Reference	1	2	3	4	5	6	7	8	9 (RCTs)	9 (CSs)	10	11 (RCTs)	11 (CSs)	12	13	14	15	16	Overall confidence	Search period	
Aburto 2013 [2]	Y	N	Y	Y	Y	Y	N	Y	Y	Y	N	Y	Y	Y	Y	Y	Y	Y	Critically low	08.2011	
Afshin 2014 [240]	Y	N	N	Y	N	Y	N	PY	N	N	N	Y	N	N	N	N	Y	Y	Critically low	25.12.2013	
Aguilar-Cordero 2020 [218]	Y	N	N	N	N	Y	N	N	Y	PY	N	Y	Y	N	N	N	Y	Y	Critically low	01.2013 – 02.2019	
Alexander 2017 [219]	Y	N	Y	PY	Y	N	Y	PY	Y	PY	N	Y	Y	N	N	N	Y	N	Critically low	01.01.1947 – 02.11.2015	
Amegah 2017 [231]	Y	N	Y	PY	N	Y	N	Y	Y	PY	N	Y	Y	Y	N	Y	Y	Y	Critically low	06.2015	
Azad 2017 [220]	Y	PY	Y	PY	Y	Y	N	Y	Y	PY	Y	Y	Y	Y	Y	Y	Y	Y	Low	01.2016	
Bolland 2015 [221]	Y	N	Y	N	N	N	N	Y	Y	N	Y	Y	No MA	Y	Y	Y	Y	Y	Critically low	09.2014	
Chowdhury 2012 [241]	Y	N	Y	PY	Y	Y	N	PY	PY	Y	N	Y	Y	N	N	Y	Y	Y	Critically low	09.2012	
Chowdhury 2014a [234]	Y	N	Y	Y	N	Y	N	PY	Y	PY	N	Y	Y	Y	Y	N	Y	Y	Critically low	01.07.2013	
Chowdhury 2014b [3]	Y	N	Y	PY	Y	Y	N	Y	Y	PY	N	Y	Y	Y	N	N	Y	Y	Critically low	01.08.2013	
Chung 2011 [242]	Y	PY	Y	N	N	N	N	Y	N	N	N	N	N	N	N	N	N	Y	Critically low	07.2011	
Ding 2017 [243]	Y	N	N	N	N	N	N	PY	N	N	N	Y	Y	N	N	N	Y	Y	Critically low	NR	
Feng 2015 [4]	N	N	N	N	N	Y	N	N	PY	PY	N	No MA	No MA	No MA	N	Y	No MA	Y	Critically low	10.10.2014	
Filippini 2020 [244]	Y	Y	Y	PY	Y	Y	Y	Y	Y	PY	Y	Y	Y	Y	Y	Y	Y	Y	Moderate	01.2019	
Fu 2021 [245]	N	N	Y	N	Y	Y	N	PY	N	N	N	Y	Y	N	N	N	N	Y	Critically low	18.02.2021	
Gayer 2019 [246]	Y	N	N	PY	Y	N	N	Y	Y	PY	Y	N	N	N	N	N	N	Y	Critically low	1946 – 08.2019	
Grosso 2015 [222]	Y	N	Y	N	Y	N	N	PY	N	PY	N	Y	Y	N	N	Y	Y	N	Critically low	01.2000 – 06.2014	
Jiang 2019 [5]	Y	N	N	PY	Y	Y	N	Y	Y	PY	N	Y	Y	N	N	Y	Y	Y	Critically low	06.2018	
Johnston 2019 (Vernooij 2019) [247, 350]	Y	Y	Y	Y	Y	Y	N	N	N/A	Y	Y	No MA	Y	Y	Y	Y	N	N	Y	Critically low	04.2019

Johnston 2019 (Zeraatkar 2019) [247, 351]	Y	Y	Y	Y	Y	Y	N	PY	Y	N/A	Y	Y	No MA	Y	Y	No MA	No MA	Y	Low	04.2019
Jonker 2020 [248]	Y	PY	N	N	Y	N	N	PY	Y	PY	Y	N	N	N	N	N	Y	Y	Critically low	1990 – 2017
Kastorini 2011 [6]	N	N	N	N	N	Y	N	N	N	N	N	N	N	N	N	N	Y	Y	Critically low	30.04.2010
Kim 2018 [223]	Y	N	N	PY	Y	Y	PY	PY	N	N	N	Y	N	N	N	N	Y	Y	Critically low	01.1970 – 08.2016
Kong 2014 [249]	Y	N	Y	N	Y	Y	N	PY	PY	PY	N	Y	Y	N	N	Y	Y	Y	Critically low	02.02.2014
Lin 2019 [250]	N	N	N	PY	N	Y	N	PY	N	N	N	Y	Y	N	N	Y	Y	Y	Critically low	01.01.2016
Lin 2020 [251]	Y	N	Y	N	Y	Y	N	Y	Y	PY	N	Y	Y	N	N	Y	Y	Y	Critically low	05.2019
Martinez-Gonzalez 2014b [252]	Y	N	Y	PY	N	Y	Y	Y	N	PY	N	No MA	Y	N	N	N	N	Y	Critically low	12.2013
Miller 2014 [253]	Y	N	Y	N	N	N	N	Y	N	N	N	N	N	N	N	N	Y	Y	Critically low	16.09.2013
Moazzen 2018 [254]	N	N	Y	N	N	N	N	PY	PY	PY	N	N	N	N	N	N	Y	Y	Critically low	01.2000 – 09.2016
Mocellin 2017 [255]	Y	N	Y	PY	N	Y	N	PY	Y	PY	N	Y	Y	Y	N	Y	Y	N	Critically low	01.2016
Morze 2021 [233]	Y	Y	Y	PY	Y	N	Y	Y	Y	Y	Y	Y	Y	N	N	N	Y	Y	Low	01.2017 – 04.2020
Picasso 2019 [256]	Y	N	N	N	Y	Y	N	PY	Y	PY	N	Y	Y	Y	Y	Y	Y	Y	Critically low	24.05.2017
Sayehmiri 2018 [257]	N	N	N	N	Y	Y	N	N	N	N	N	Y	Y	N	N	N	N	Y	Critically low	2016
Schwingshackl 2015 [237]	Y	N	N	PY	Y	Y	Y	PY	PY	PY	N	Y	Y	Y	Y	Y	Y	Y	Low	02.04.2014
Schwingshackl 2017 [258]	Y	N	N	PY	Y	N	N	PY	Y	N	N	Y	Y	Y	N	Y	Y	Y	Critically low	08.2016
Setien-Suero 2016 [259]	N	N	N	N	N	N	PY	PY	N	N	N	Y	Y	N	N	N	N	N	Critically low	01.2005 – 12.2015
Stratton 2011 [224]	Y	N	N	N	Y	N	N	Y	N	N	N	N	N	N	N	N	N	Y	Critically low	02.2010
Te Morenga 2013 [260]	Y	N	Y	N	N	Y	Y	Y	Y	N	Y	Y	Y	Y	Y	Y	N	Y	Critically low	12.2011
Thorne-Lyman 2012 [225]	Y	N	Y	N	N	N	N	PY	Y	N	N	Y	Y	N	N	N	N	Y	Critically low	06.2011
Trikalinos 2012 [239]	N	N	Y	N	N	N	N	PY	N	N	N	N	N	N	N	Y	N	Y	Critically low	07.10.2010
Vinceti 2018a [7]	Y	Y	Y	Y	Y	N	Y	Y	Y	PY	N	Y	Y	Y	Y	Y	Y	Y	Moderate	02.2017

Vinceti 2018b [236]	N	N	Y	N	N	N	N	PY	N	N	N	Y	No MA	N	N	Y	N	N	Critically low	11.06.2018
Wien 2012 [226]	Y	PY	N	PY	Y	Y	PY	Y	Y	N	N	Y	N	N	Y	Y	Y	Y	Low	06.05.2010
Wolf 2017 [8]	Y	N	N	PY	Y	Y	Y	PY	Y	PY	N	No MA	N	Y	Y	N	Y	Y	Critically low	17.06.2016
Yang 2016 [235]	N	N	N	PY	Y	Y	N	N	PY	PY	N	No MA	N	N	N	Y	Y	Y	Critically low	08.12.2014
Yang 2020 [227]	Y	N	Y	N	Y	Y	N	Y	PY	PY	N	Y	Y	Y	Y	Y	Y	Y	Critically low	03.2019
Yao 2019 [228]	Y	PY	Y	N	Y	Y	PY	PY	Y	Y	N	Y	Y	N	Y	Y	Y	Y	Low	31.12.2019
Yu 2021 [229]	Y	N	N	N	Y	Y	N	PY	Y	PY	N	Y	N	N	N	N	N	Y	Critically low	18.06.2020
Zhang 2016 [9]	Y	N	Y	N	Y	Y	N	PY	N	N	N	Y	Y	N	N	Y	Y	Y	Critically low	15.12.2013
Zhang 2018 [230]	Y	N	Y	PY	N	Y	N	PY	Y	PY	N	Y	Y	Y	N	Y	Y	Y	Critically low	16.05.2017
Zhao 2014 [261]	Y	N	Y	N	N	Y	N	PY	N	PY	N	No MA	Y	Y	N	N	Y	Y	Critically low	01.09.2013
Zhou 2017 [232]	Y	N	Y	N	N	Y	N	PY	Y	PY	N	Y	Y	Y	N	Y	Y	Y	Critically low	06.2016

CS=cohort study; MA=meta-analysis; N=no; N/A=not applicable; NR=not reported; PY=partial yes; RCT=randomised controlled trial; Y=yes

NB: We used AMSTAR 2 (A MeaSurement Tool to Assess systematic Reviews, version 2) [352] to evaluate the methodological quality of the systematic reviews included in our sample. Two reviewers (JB, JS) rated the 16 predefined items independently; discrepancies were discussed with a third reviewer (LS). Based on the presence of critical and noncritical weaknesses, the overall confidence in the systematic review findings was graded as "high", "moderate", "low", or "critically low".

Supplementary Table S9 Ratings of PI/ECO similarity degree for included BoE-pairs

BoE			PI/ECO similarities				
Reference	Intervention/ Exposure	Outcome	Population	Intervention/ Exposure	Comparator	Outcome	Overall
Aburto 2013 [2]	Low sodium	All-cause mortality	2	1	1	1	2
Aburto 2013 [2]	Low sodium	Cardiovascular disease	2	1	1	2	2
Afshin 2014 [240]	Nuts	Coronary heart disease	2	1	1	1	2
Afshin 2014 [240]	Nuts	Stroke	2	1	1	1	2
Afshin 2014 [240]	Nuts	Type 2 diabetes	2	1	1	1	2
Aguilar-Cordero 2020 [218]	Vitamin D	Pre-eclampsia	1	3	3	1	3
Alexander 2017 [219]	Omega-3 fatty acids	Coronary heart disease	2	2	2	1	2
Amegah 2017 [231]	Vitamin D	Gestational length	1	3	3	1	3
Azad 2017 [220]	Nonnutritive sweeteners	Body mass index	2	2	1	1	2
Bolland 2015 [221]	Calcium	All fractures	1	2	1	1	2
Chowdhury 2012 [241]	Omega-3 fatty acids	Cerebrovascular disease	2	2	2	1	2
Chowdhury 2014a [234]	α -linolenic acid	Coronary disease	2	2	2	1	2
Chowdhury 2014a [234]	Omega-6 fatty acids	Coronary disease	2	2	2	1	2
Chowdhury 2014b [3]	Vitamin D	All-cause mortality	2	3	3	1	3
Chung 2011 [242]	Vitamin D	Colorectal cancer	2	3	3	1	3
Chung 2011 [242]	Vitamin D	Breast cancer	2	3	3	1	3
Ding 2017 [243]	Dairy	Systolic blood pressure	2	1	1	1	2
Feng 2015 [4]	Folic acid	Congenital heart defects	1	1	1	1	1
Filippini 2020 [244]	Green Tea	Prostate cancer	3	2	2	1	3
Filippini 2020 [244]	Green Tea	Endometrial cancer	2	2	2	1	2
Fu 2021 [245]	Vitamin C	Lung cancer	2	1	1	1	2
Fu 2021 [245]	Vitamin C	Breast cancer	2	1	1	1	2
Fu 2021 [245]	Vitamin C	Colorectal cancer	2	1	1	1	2
Gayer 2019 [246]	Apples	Body weight	2	1	1	1	2
Grosso 2015 [222]	Mediterranean diet	Cardiovascular disease	2	1	1	1	2
Jiang 2019 [5]	Vitamin E	Age-related cataract	1	2	2	1	2

Jiang 2019 [5]	β-carotene	Age-related cataract	1	2	2	1	2
Johnston 2019 [247]	Low red meat	Cardiovascular disease	2	2	2	1	2
Johnston 2019 [247]	Low red meat	Type 2 diabetes	2	2	2	1	2
Johnston 2019 [247]	Low red meat	Colorectal cancer	2	2	2	1	2
Jonker 2020 [248]	Folic acid	Mean birthweight	1	1	1	1	1
Jonker 2020 [248]	Folic acid	Low birthweight	1	1	1	1	1
Jonker 2020 [248]	Folic acid	Small for gestational age	1	2	2	1	2
Kastorini 2011 [6]	Mediterranean diet	Metabolic syndrome	2	1	1	1	2
Kastorini 2011 [6]	Mediterranean diet	HDL-cholesterol	2	1	1	1	2
Kastorini 2011 [6]	Mediterranean diet	Triglycerides	2	1	1	1	2
Kastorini 2011 [6]	Mediterranean diet	Systolic blood pressure	2	1	1	1	2
Kim 2018 [223]	Multivitamins/ Minerals	Cardiovascular disease	2	1	1	1	2
Kim 2018 [223]	Multivitamins/ Minerals	Coronary heart disease	1	1	1	1	1
Kim 2018 [223]	Multivitamins/ Minerals	Stroke	2	1	1	1	2
Kong 2014 [249]	Vitamins	Gastric cancer	2	2	2	1	2
Lin 2019 [250]	Vitamin E	Bladder cancer	1	2	1	1	2
Lin 2020 [251]	Vitamin D	Nephrolithiasis	1	1	1	1	1
Lin 2020 [251]	Calcium	Nephrolithiasis	1	1	1	1	1
Martinez-Gonzalez 2014b [252]	Olive oil	Coronary heart disease	2	2	2	2	2
Martinez-Gonzalez 2014b [252]	Olive oil	Stroke	2	2	2	1	2
Miller 2014 [253]	Low calorie sweeteners	Body weight	1	1	1	1	1
Moazzen 2018 [254]	Folic acid	Colorectal cancer	2	1	1	1	2
Mocellin 2017 [255]	Vitamin B6	Cancer	2	2	2	1	2
Morze 2021 [233]	Mediterranean diet	Cancer mortality	2	1	1	1	2
Morze 2021 [233]	Mediterranean diet	Breast cancer	2	1	1	1	2
Picasso 2019 [256]	Vegetarian diet	Triglycerides	3	2	2	1	3
Picasso 2019 [256]	Vegetarian diet	HDL-cholesterol	3	2	2	1	3
Sayehmiri 2018 [257]	Selenium	Prostate cancer	2	2	2	1	2
Schwingshackl 2015 [237]	Mediterranean diet	Type 2 diabetes	2	1	1	1	2
Schwingshackl 2017 [258]	Olive oil	Type 2 diabetes	2	1	1	1	2

Setien-Suero 2016 [259]	Homocysteine	Mini mental state examination score	1	3	3	1	3
Stratton 2011 [224]	Multivitamins	Prostate cancer	1	1	1	1	1
Stratton 2011 [224]	Vitamin E	Prostate cancer	1	1	1	1	1
Stratton 2011 [224]	Vitamin C	Prostate cancer	2	1	1	1	2
Stratton 2011 [224]	β-carotene	Prostate cancer	2	1	1	1	2
Te Morenga 2013 [260]	Low dietary sugars	Weight gain	2	2	2	1	2
Te Morenga 2013 [260]	Low dietary sugars	Body mass index	2	2	2	1	2
Thorne-Lyman 2012 [225]	Vitamin D	Mean birthweight	1	2	2	1	2
Trikalinos 2012 [239]	Vitamin E	Cardiovascular disease	2	2	2	1	2
Vinceti 2018a [7]	Selenium	Any cancer	2	3	3	1	3
Vinceti 2018a [7]	Selenium	Cancer mortality	2	2	2	1	2
Vinceti 2018b [236]	Selenium	Type 2 diabetes	2	2	2	1	2
Wien 2012 [226]	Folic acid	Cancer	2	2	2	2	2
Wolf 2017 [8]	Multivitamins	Preterm birth	1	1	1	1	1
Wolf 2017 [8]	Multivitamins	Stillbirth	2	1	1	1	2
Yang 2016 [235]	Folic acid	Gestational hypertension	2	2	2	1	2
Yang 2020 [227]	Calcium	Cardiovascular disease	1	2	2	1	2
Yao 2019 [228]	Vitamin D	Any fracture	2	3	3	1	3
Yu 2021 [229]	Folic acid	Pre-eclampsia	2	2	2	1	2
Zhang 2016 [9]	Selenium	Cardiovascular disease	2	3	3	1	3
Zhang 2018 [230]	Vitamin D	Gestational diabetes	2	3	3	1	3
Zhang 2018 [230]	Vitamin D	Fasting plasma glucose	2	3	3	1	3
Zhao 2014 [261]	Multivitamins/ Minerals	Nuclear cataract	2	1	1	1	2
Zhao 2014 [261]	Multivitamins/ Minerals	Cortical cataract	2	1	1	1	2
Zhao 2014 [261]	Multivitamins/ Minerals	Posterior subcapsular cataract	2	1	1	1	2
Zhou 2017 [232]	Vitamin D	Preterm birth	2	3	3	1	3

BoE= bodies of evidence; HDL=high density lipoprotein; PI/ECO=Population, Intervention/Exposure, Comparator, Outcome

Supplementary Table S10 Analysis of concordance of the included BoE-pairs

Reference	Intervention/ Exposure	Outcome	Summary measure	Direction of effect (RCT)	Significance (RCT)	Direction of association (cohorts)	Significance (cohorts)	Concordance
Aburto 2013 [2]	Low sodium	All-cause mortality	RR	decreasing	not significant	decreasing	not significant	not concordant
Aburto 2013 [2]	Low sodium	Cardiovascular disease	RR	decreasing	not significant	decreasing	not significant	not concordant
Afshin 2014 [240]	Nuts	Coronary heart disease	RR	decreasing	not significant	decreasing	significant	not concordant
Afshin 2014 [240]	Nuts	Stroke	RR	decreasing	significant	decreasing	not significant	not concordant
Afshin 2014 [240]	Nuts	Type 2 diabetes	RR	decreasing	not significant	decreasing	significant	not concordant
Aguilar-Cordero 2020 [218]	Vitamin D	Pre-eclampsia	RR	decreasing	not significant	decreasing	not significant	not concordant
Alexander 2017 [219]	Omega-3 fatty acids	Coronary heart disease	RR	decreasing	not significant	decreasing	significant	not concordant
Amegah 2017 [231]	Vitamin D	Gestational length	SMD	increasing	not significant	increasing	significant	not concordant
			MD	increasing	significant	increasing	significant	concordant
Azad 2017 [220]	Nonnutritive sweeteners	Body mass index	SMD	decreasing	not significant	increasing	significant	not concordant
			MD	decreasing	not significant	increasing	significant	not concordant
Bolland 2015 [221]	Calcium	All fractures	RR	decreasing	significant	increasing	not significant	not concordant
Chowdhury 2012 [241]	Omega-3 fatty acids	Cerebrovascular disease	RR	decreasing	not significant	decreasing	not significant	concordant
Chowdhury 2014a [234]	α -linolenic acid	Coronary disease	RR	decreasing	not significant	decreasing	not significant	not concordant
Chowdhury 2014a [234]	Omega-6 fatty acids	Coronary disease	RR	decreasing	not significant	decreasing	not significant	not concordant
Chowdhury 2014b [3]	Vitamin D	All-cause mortality	RR	decreasing	not significant	decreasing	significant	not concordant
Chung 2011 [242]	Vitamin D	Colorectal cancer	RR	increasing	not significant	decreasing	significant	not concordant
Chung 2011 [242]	Vitamin D	Breast cancer	RR	decreasing	not significant	decreasing	not significant	not concordant
Ding 2017 [243]	Dairy	Systolic blood pressure	MD	decreasing	not significant	decreasing	significant	not concordant
Feng 2015 [4]	Folic acid	Congenital heart defects	RR	decreasing	not significant	decreasing	significant	not concordant
Filippini 2020 [244]	Green Tea	Prostate cancer	RR	decreasing	not significant	increasing	not significant	not concordant
Filippini 2020 [244]	Green Tea	Endometrial cancer	RR	decreasing	not significant	decreasing	not significant	not concordant
Fu 2021 [245]	Vitamin C	Lung cancer	RR	increasing	not significant	increasing	not significant	not concordant
Fu 2021 [245]	Vitamin C	Breast cancer	RR	increasing	not significant	increasing	not significant	not concordant
Fu 2021 [245]	Vitamin C	Colorectal cancer	RR	decreasing	not significant	decreasing	not significant	not concordant
Gayer 2019 [246]	Apples	Body weight	MD	increasing	not significant	decreasing	significant	not concordant

Grosso 2015 [222]	Mediterranean diet	Cardiovascular disease	RR	decreasing	significant	decreasing	significant	concordant
Jiang 2019 [5]	Vitamin E	Age-related cataract	RR	decreasing	not significant	decreasing	not significant	concordant
Jiang 2019 [5]	β -carotene	Age-related cataract	RR	decreasing	not significant	decreasing	significant	not concordant
Johnston 2019 [247]	Low red meat	Cardiovascular disease	RR	increasing	not significant	decreasing	significant	not concordant
Johnston 2019 [247]	Low red meat	Type 2 diabetes	RR	decreasing	not significant	decreasing	significant	not concordant
Johnston 2019 [247]	Low red meat	Colorectal cancer	RR	increasing	not significant	decreasing	not significant	concordant
Jonker 2020 [248]	Folic acid	Mean birthweight	SMD	increasing	significant	increasing	significant	concordant
			MD	increasing	significant	increasing	significant	concordant
Jonker 2020 [248]	Folic acid	Low birthweight	RR	decreasing	not significant	decreasing	significant	not concordant
Jonker 2020 [248]	Folic acid	Small for gestational age	RR	decreasing	not significant	decreasing	not significant	not concordant
Kastorini 2011 [6]	Mediterranean diet	Metabolic syndrome	RR	decreasing	significant	decreasing	not significant	not concordant
Kastorini 2011 [6]	Mediterranean diet	HDL-cholesterol	MD	increasing	significant	increasing	not significant	not concordant
Kastorini 2011 [6]	Mediterranean diet	Triglycerides	MD	decreasing	significant	decreasing	not significant	not concordant
Kastorini 2011 [6]	Mediterranean diet	Systolic blood pressure	MD	decreasing	significant	increasing	not significant	not concordant
Kim 2018 [223]	Multivitamins/Minerals	Cardiovascular disease	RR	decreasing	not significant	increasing	not significant	concordant
Kim 2018 [223]	Multivitamins/Minerals	Coronary heart disease	RR	decreasing	not significant	decreasing	significant	not concordant
Kim 2018 [223]	Multivitamins/Minerals	Stroke	RR	increasing	not significant	decreasing	not significant	not concordant
Kong 2014 [249]	Vitamins	Gastric cancer	RR	increasing	not significant	decreasing	not significant	not concordant
Lin 2019 [250]	Vitamin E	Bladder cancer	RR	increasing	not significant	decreasing	significant	not concordant
Lin 2020 [251]	Vitamin D	Nephrolithiasis	RR	decreasing	not significant	increasing	significant	not concordant
Lin 2020 [251]	Calcium	Nephrolithiasis	RR	increasing	not significant	increasing	significant	not concordant
Martinez-Gonzalez 2014b [252]	Olive oil	Coronary heart disease	RR	decreasing	not significant	decreasing	not significant	not concordant
Martinez-Gonzalez 2014b [252]	Olive oil	Stroke	RR	decreasing	significant	decreasing	significant	concordant
Moazzen 2018 [254]	Folic acid	Colorectal cancer	RR	increasing	not significant	decreasing	not significant	not concordant
Mocellin 2017 [255]	Vitamin B6	Cancer	RR	increasing	not significant	decreasing	not significant	concordant
Morze 2021 [233]	Mediterranean diet	Cancer mortality	RR	decreasing	not significant	decreasing	significant	not concordant
Morze 2021 [233]	Mediterranean diet	Breast cancer	RR	decreasing	significant	decreasing	not significant	not concordant

Picasso 2019 [256]	Vegetarian diet	Triglycerides	SMD	increasing	not significant	increasing	not significant	not concordant
			MD	increasing	not significant	increasing	not significant	not concordant
Picasso 2019 [256]	Vegetarian diet	HDL-cholesterol	SMD	decreasing	not significant	decreasing	not significant	not concordant
			MD	decreasing	not significant	decreasing	not significant	not concordant
Sayehmiri 2018 [257]	Selenium	Prostate cancer	RR	decreasing	not significant	decreasing	not significant	not concordant
Schwingshackl 2015 [237]	Mediterranean diet	Type 2 diabetes	RR	decreasing	significant	decreasing	significant	concordant
Schwingshackl 2017 [258]	Olive oil	Type 2 diabetes	RR	decreasing	significant	decreasing	significant	concordant
Stratton 2011 [224]	Multivitamins	Prostate cancer	RR	decreasing	not significant	increasing	not significant	concordant
Stratton 2011 [224]	Vitamin E	Prostate cancer	RR	increasing	not significant	increasing	not significant	not concordant
Stratton 2011 [224]	Vitamin C	Prostate cancer	RR	increasing	not significant	decreasing	not significant	not concordant
Stratton 2011 [224]	β -carotene	Prostate cancer	RR	increasing	not significant	decreasing	significant	not concordant
Te Morenga 2013 [260]	Low dietary sugars	Body weight change	MD	increasing	significant	increasing	not significant	not concordant
Te Morenga 2013 [260]	Low dietary sugars	Body mass index	MD	decreasing	not significant	decreasing	not significant	concordant
Thorne-Lyman 2012 [225]	Vitamin D	Mean birthweight	SMD	increasing	not significant	increasing	significant	not concordant
			MD	increasing	not significant	increasing	significant	not concordant
Trikalinos 2012 [239]	Vitamin E	Cardiovascular disease	RR	decreasing	not significant	decreasing	significant	not concordant
Vinceti 2018a [7]	Selenium	Any cancer	RR	decreasing	not significant	decreasing	significant	not concordant
Vinceti 2018a [7]	Selenium	Cancer mortality	RR	decreasing	not significant	decreasing	not significant	not concordant
Vinceti 2018b [236]	Selenium	Type 2 diabetes	RR	increasing	significant	increasing	significant	concordant
Wien 2012 [226]	Folic acid	Cancer	RR	increasing	significant	increasing	not significant	not concordant
Wolf 2017 [8]	Multivitamins	Preterm birth	RR	increasing	not significant	decreasing	not significant	not concordant
Wolf 2017 [8]	Multivitamins	Stillbirth	RR	increasing	not significant	decreasing	not significant	not concordant
Yang 2016 [235]	Folic acid	Gestational hypertension	RR	decreasing	significant	increasing	significant	not concordant
Yang 2020 [227]	Calcium	Cardiovascular disease	RR	decreasing	not significant	decreasing	not significant	not concordant
Yao 2019 [228]	Vitamin D	Any fracture	RR	increasing	not significant	decreasing	significant	not concordant
Yu 2021 [229]	Folic acid	Pre-eclampsia	RR	increasing	not significant	decreasing	significant	not concordant
Zhang 2016 [9]	Selenium	Cardiovascular disease	RR	decreasing	not significant	decreasing	significant	not concordant
Zhang 2018 [230]	Vitamin D	Gestational diabetes	RR	decreasing	not significant	decreasing	significant	not concordant
Zhao 2014 [261]	Multivitamins/ Minerals	Nuclear cataract	RR	decreasing	significant	decreasing	significant	concordant

Zhao 2014 [261]	Multivitamins/ Minerals	Cortical cataract	RR	decreasing	not significant	decreasing	significant	not concordant
Zhao 2014 [261]	Multivitamins/ Minerals	Posterior subcapsular cataract	RR	increasing	significant	decreasing	not significant	not concordant
Zhou 2017 [232]	Vitamin D	Preterm birth	RR	decreasing	not significant	decreasing	not significant	not concordant

MD=mean difference; RCT=randomised controlled trial; RR=relative risk; SMD=standardised mean difference

Supplementary Table S11 Subgroup analysis by PI/ECO similarity degree for each domain

	BoE-pairs included	Ratio of risk ratios (95% CI)	Heterogeneity (I² (%); τ^2)	95% prediction interval
Main analysis	66	1.04 (0.99 to 1.10)	59; 0.02	0.77 to 1.41
Stratified by population similarity degree				
More or less identical	15	1.03 (0.94 to 1.13)	55; 0.01	0.79 to 1.35
Similar but not identical	50	1.05 (0.98 to 1.12)	59; 0.03	0.76 to 1.46
Broadly similar	1	0.46 (0.16 to 1.29)	N/A	N/A
Stratified by intervention/exposure similarity degree				
More or less identical	30	0.99 (0.90 to 1.08)	42; 0.03	0.70 to 1.39
Similar but not identical	27	1.05 (0.98 to 1.13)	46; 0.01	0.81 to 1.36
Broadly similar	9	1.20 (1.06 to 1.36)	53; 0.01	0.90 to 1.60
Stratified by comparator similarity degree				
More or less identical	32	0.99 (0.91 to 1.08)	46; 0.03	0.70 to 1.39
Similar but not identical	25	1.07 (0.99 to 1.15)	39; 0.02	0.84 to 1.34
Broadly similar	9	1.20 (1.06 to 1.36)	53; 0.01	0.90 to 1.60
Stratified by outcome similarity degree				
More or less identical	63	1.05 (0.99 to 1.11)	61; 0.02	0.76 to 1.44
Similar but not identical	3	1.02 (0.89 to 1.16)	0; 0	0.42 to 2.46
Broadly similar	0	N/A	N/A	N/A

BoE=bodies of evidence; CI=confidence interval; N/A=not applicable; PI/ECO=Population, Intervention/Exposure, Comparator, Outcome

Supplementary Table S12 Overlap of primary studies in BoE-pairs with highly similar PI/ECO questions – comparison between the present sample and the sample in Schwingshackl 2021 [1]

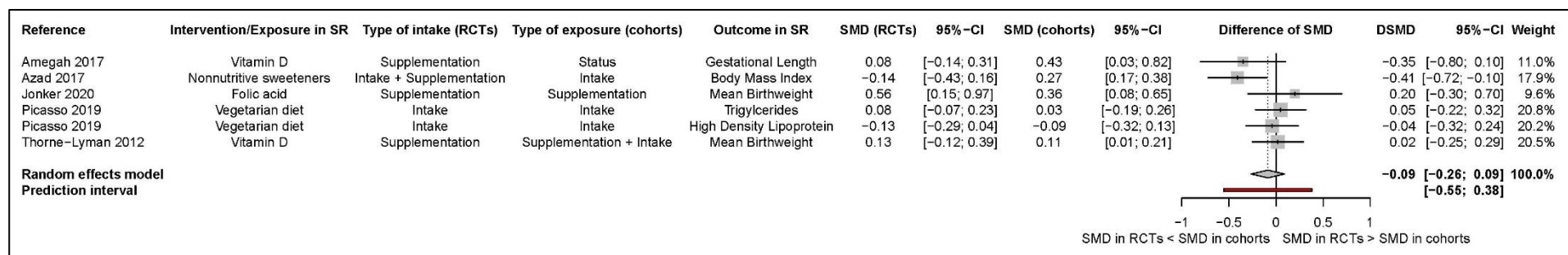
Reference	Intervention/ Exposure	Outcome	Matching BoE-pair in Schwingshackl 2021 [1]	Intervention/ Exposure	Outcome	Overlap of RCTs [#]	Overlap of CSs [#]
Binary outcomes (n=19)							
Aburto 2013 [2]	Low sodium	All-cause mortality	Adler 2014 [353]/ Aburto 2013 [2]	Low sodium	All-cause mortality	4/4	2/2
Aburto 2013 [2]	Low sodium	Cardiovascular disease	Adler 2014 [353]/ Aburto 2013 [2]	Low sodium	Cardiovascular disease	2/2	3/3
			Adler 2014 [353]/ Aburto 2013 [2]	Low sodium	Cardiovascular mortality	1/2	3/3
Aguilar-Cordero 2020 [218]	Vitamin D	Pre-eclampsia	Palacios 2019 [354]/ Yuan 2019 [355]	Vitamin D	Pre-eclampsia	0/7	5/7
Alexander 2017 [219]	Omega-3 fatty acids	Coronary heart disease	Abdelhamid 2018a [356]/ Chowdhury 2014a [234]	Omega-3 fatty acids	Coronary heart disease/ Cardiovascular disease	13/18	7/17
			Abdelhamid 2018a [356]/ Chowdhury 2014a [234]	Omega-3 fatty acids	Cardiovascular mortality/ Coronary heart disease mortality	10/18	N/A
Chowdhury 2014a [234]	α -linolenic acid	Coronary disease	Abdelhamid 2018a [356]/ Wei 2018 [357]	α -linolenic acid	Coronary heart disease	2/4	5/7
Chowdhury 2014a [234]	Omega-6 fatty acids	Coronary disease	Abdelhamid 2018b [358]/ Chowdhury 2014a [234]	Polyunsaturated fat/ Omega-6 fatty acids	Coronary heart disease	4/8	8/8
			Hooper 2018 [359]/ Chowdhury 2014a [234]	Omega-6 fatty acids	Combined cardiovascular events/ Coronary heart disease	4/8	8/8
Chowdhury 2014b [3]	Vitamin D	All-cause mortality	Bjelakovic 2014a [360]/ Chowdhury 2014b [3]	Vitamin D	All-cause mortality	19/22	68/68
Chung 2011 [242]	Vitamin D	Breast cancer	Bjelakovic 2014b [361]/ Hossain 2019 [362]	Vitamin D/ Vitamin D3	Breast cancer	0/1	0/4
Feng 2015 [4]	Folic acid	Congenital heart defects	De-Regil 2015 [363]/ Feng 2015 [4]	Folic acid	Congenital cardiovascular anomalies / Congenital heart defects	0/1	1/1

Grosso 2015 [222]	Mediterranean diet	Cardiovascular disease	Rees 2019 [364]/ Rosato 2019 [365]	Mediterranean diet	Combined cardiovascular events / Cardiovascular disease	1/4	5/20
			Rees 2019 [364]/ Rosato 2019 [365]	Mediterranean diet	Cardiovascular mortality	1/4	4/20
Jiang 2019 [5]	Vitamin E	Age-related cataract	Mathew 2012 [366]/ Jiang 2019 [5]	Vitamin E	Cataract	3/6	6/6
Jiang 2019 [5]	β -carotene	Age-related cataract	Mathew 2012 [366]/ Jiang 2019 [5]	β -carotene	Cataract	2/4	7/7
Vinceti 2018a [7]	Selenium	Any cancer	Vinceti 2018a [7]	Selenium	Cancer	5/5	7/7
Vinceti 2018a [7]	Selenium	Cancer mortality	Vinceti 2018a [7]	Selenium	Cancer mortality	2/2	1/1
Wolf 2017 [8]	Multivitamins	Preterm birth	Keats 2019 [367]/ Wolf 2017 [8]	Micronutrients/ Multivitamins	Preterm birth	0/1	4/4
Yao 2019 [228]	Vitamin D	Any fracture	Avenell 2014 [368]/ Feng 2017 [369]	Vitamin D	Any fracture	8/11	4/10
Zhang 2016 [9]	Selenium	Cardiovascular disease	Rees 2013b [370]/ Xiang 2019 [371]	Selenium	Cardiovascular mortality	2/9	1/14
			Rees 2013b [370]/ Zhang 2016 [9]	Selenium	Combined cardiovascular events / Cardiovascular disease	2/9	14/14
Zhang 2018 [230]	Vitamin D	Gestational diabetes	Palacios 2019 [354]/ Hu 2018 [372]	Vitamin D	Gestational diabetes	3/4	9/24
Zhou 2017 [232]	Vitamin D	Preterm birth	Palacios 2019 [354]/ Tous 2020 [373]	Vitamin D	Preterm birth	0/5	8/12
Continuous outcomes (n=5)							
Ding 2017 [243]	Dairy	Systolic blood pressure	Usinger 2012 [374]/ Soedamah-Muthu 2012 [375]	Fermented milk	Systolic blood pressure/ Hypertension	0/8	2/27
Kastorini 2011 [6]	Mediterranean diet	HDL-cholesterol	Rees 2019 [364]/ Kastorini 2011 [6]	Mediterranean diet	HDL-cholesterol	2/27	1/1

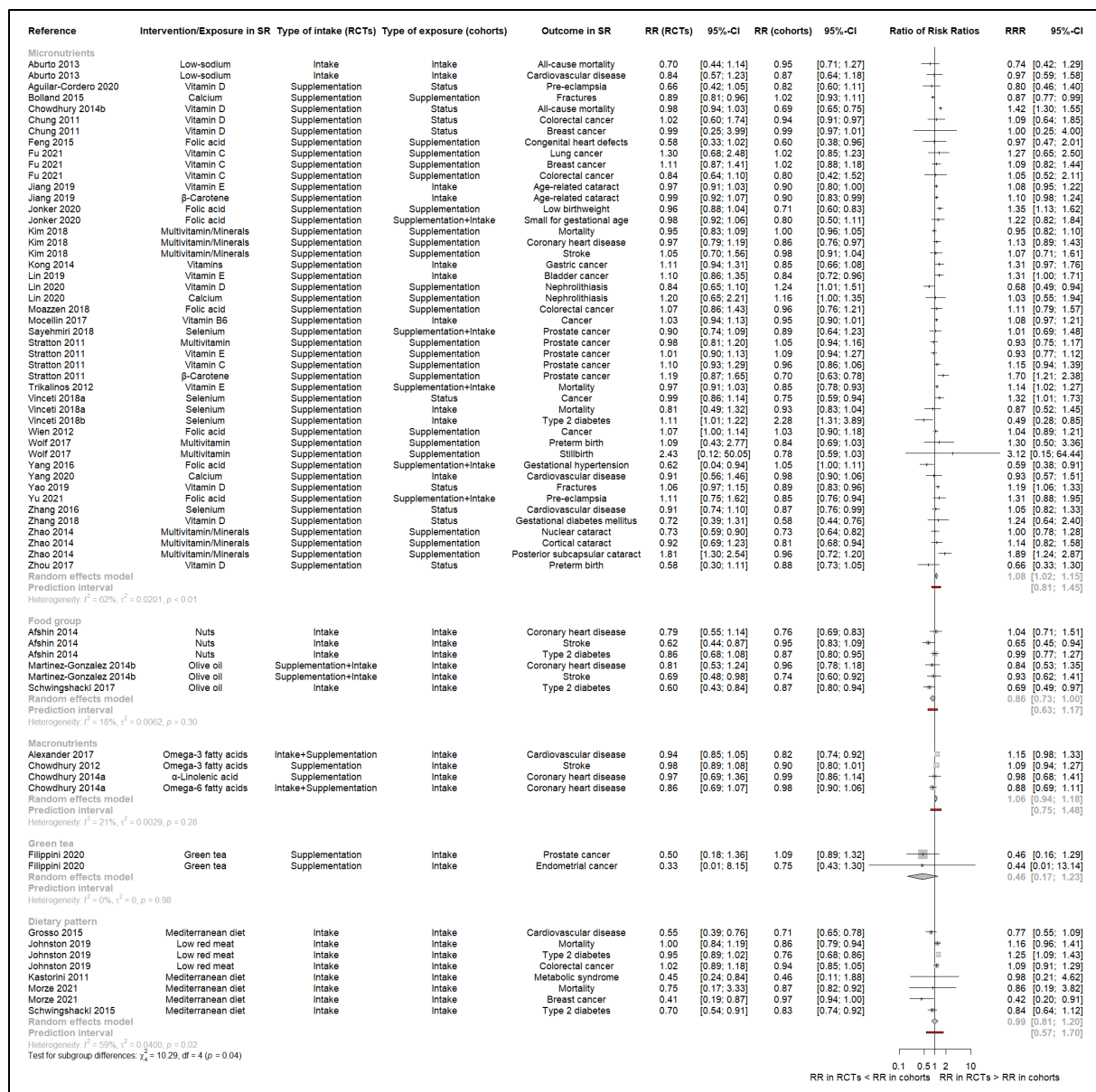
Kastorini 2011 [6]	Mediterranean diet	Triglycerides	Rees 2019 [364]/ Kastorini 2011 [6]	Mediterranean diet	Triglycerides	2/27	1/1
Kastorini 2011 [6]	Mediterranean diet	Systolic blood pressure	Rees 2019 [364]/ Kastorini 2011 [6]	Mediterranean diet	Systolic blood pressure	2/14	1/1
			Rees 2013a [376]/ Kastorini 2011 [6]	Healthy diet/ Mediterranean diet	Systolic blood pressure	0/14	1/1
Thorne-Lyman 2012 [225]	Vitamin D	Birthweight	Palacios 2019 [354]/ Tous 2020 [373]	Vitamin D	Mean birthweight/ Birth weight	1/5	0/2

[#] overlap defined as number of primary studies also included in Schwingshackl et al. 2021 [1], reference: total number of primary studies included in the BoE of the present replication study

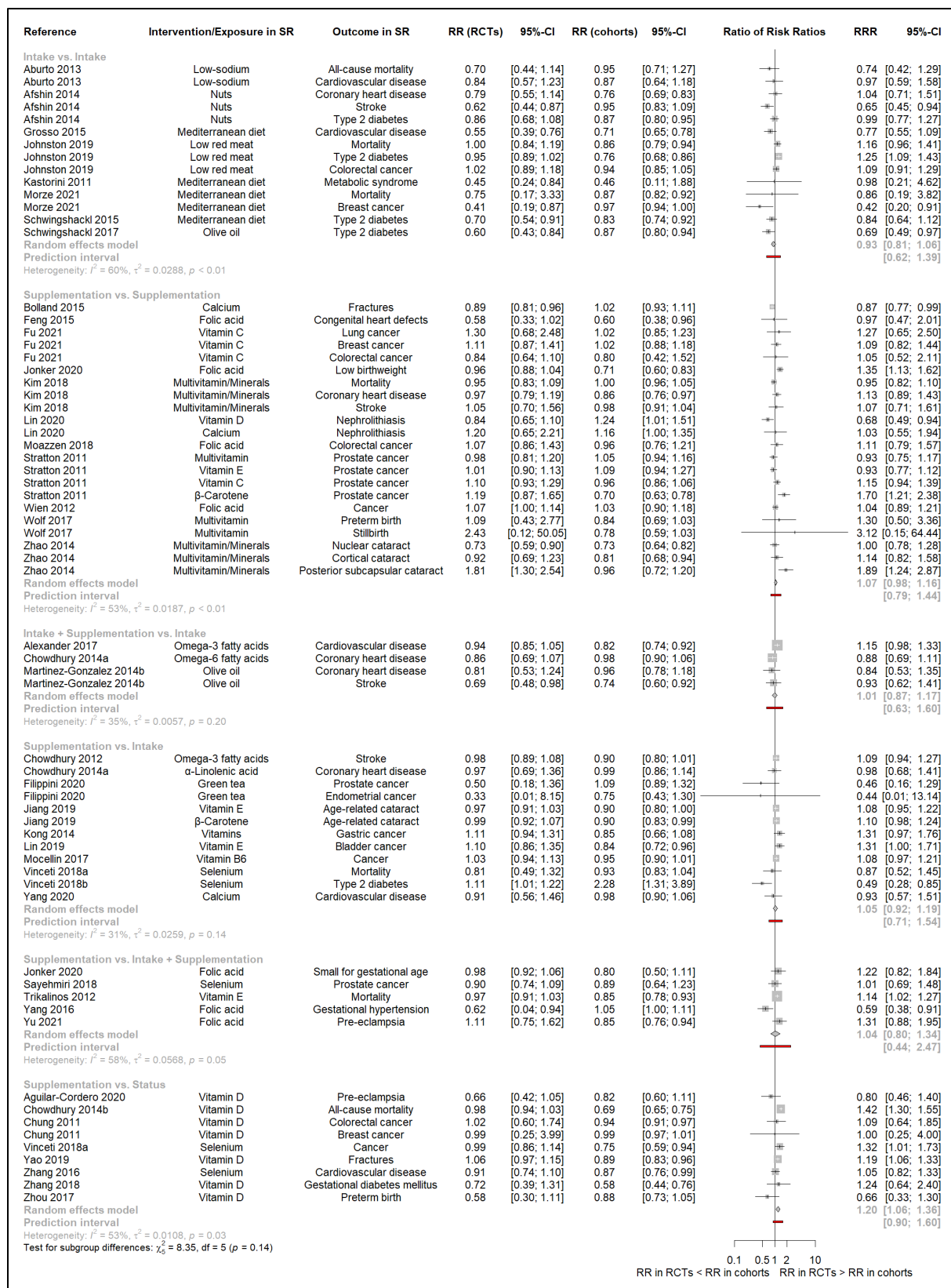
BoE=bodies of evidence; CS=cohort study; HDL=high density lipoprotein; N/A=not applicable; PI/ECO=Population, Intervention/Exposure, Comparator, Outcome; RCT=randomised controlled trial



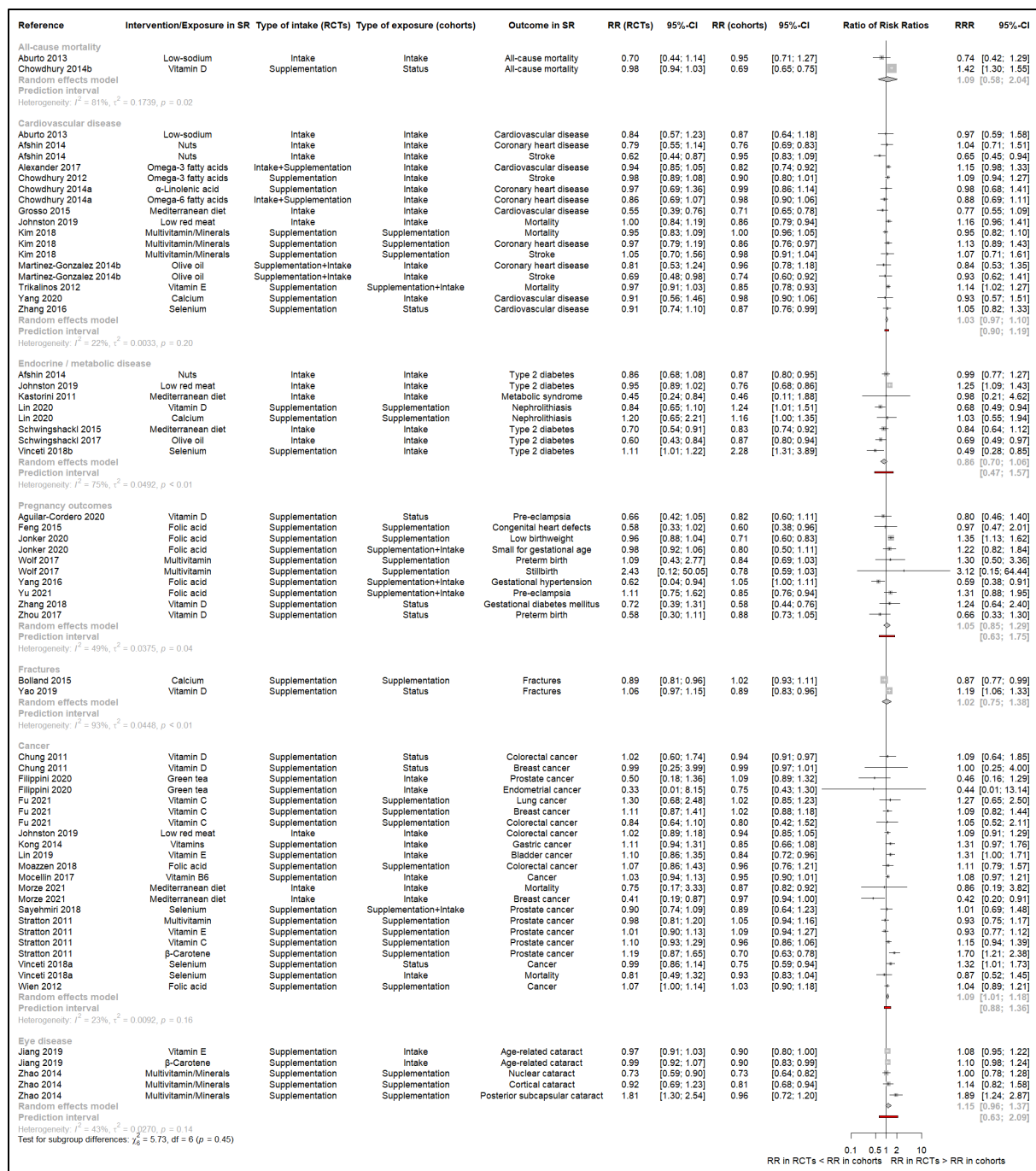
Supplementary Fig. S1 Forest plot, analysis of BoE-pairs with continuous outcomes and standardised mean difference



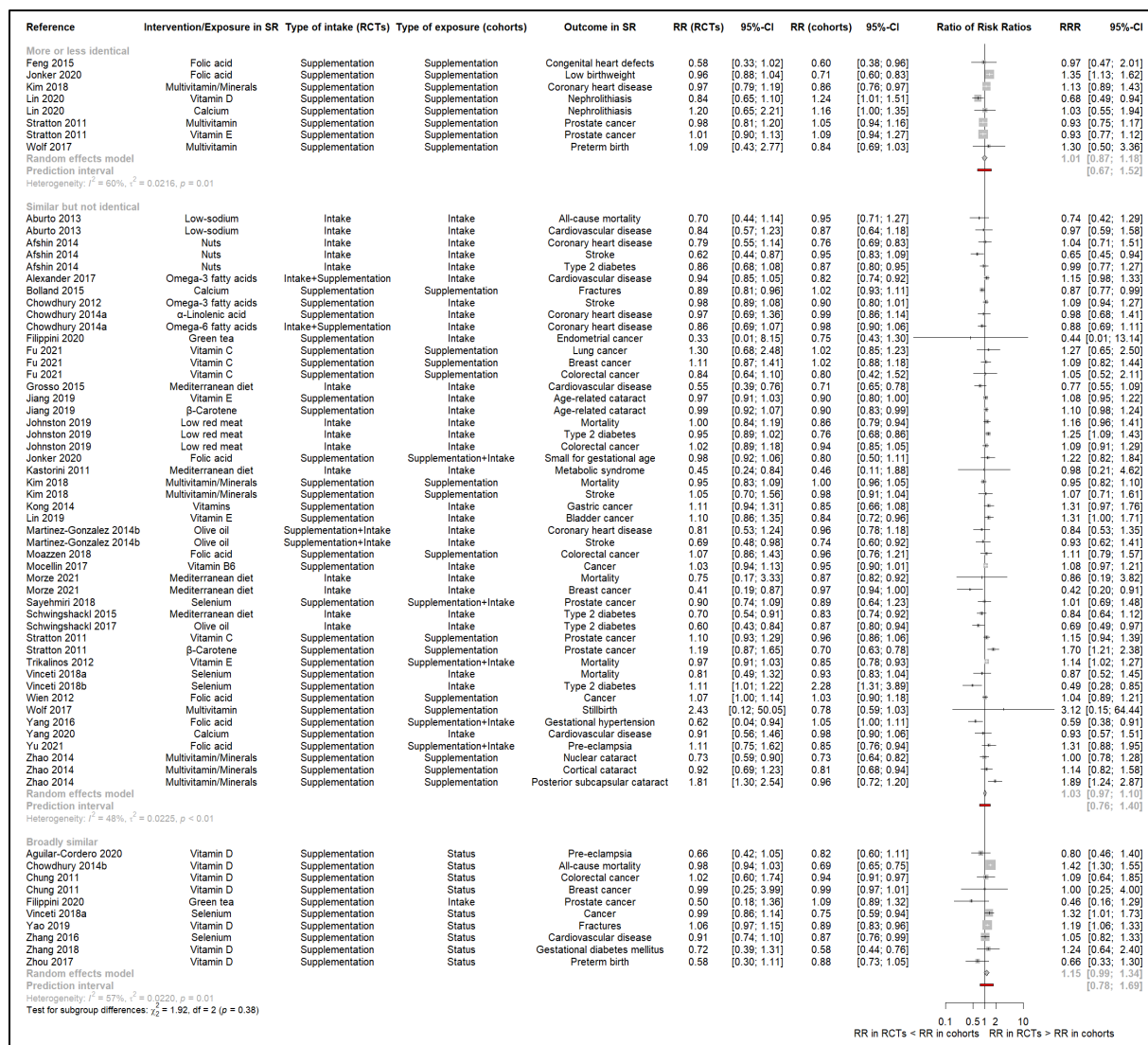
Supplementary Fig. S2 Forest plot, subgroup analysis of BoE-pairs with binary outcomes by type of intervention/exposure



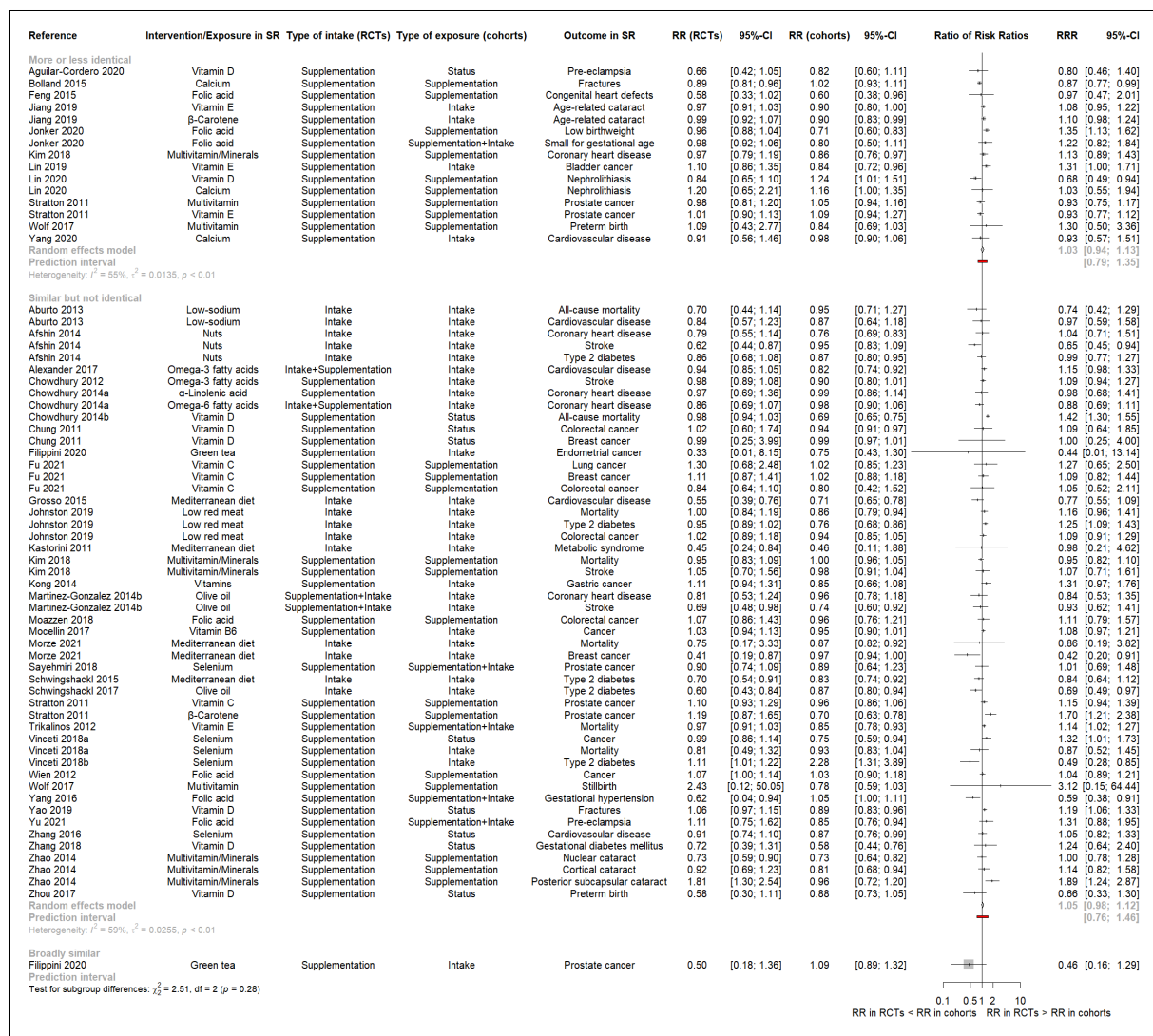
Supplementary Fig. S3 Forest plot, subgroup analysis of BoE-pairs with binary outcomes by type of intake/exposure



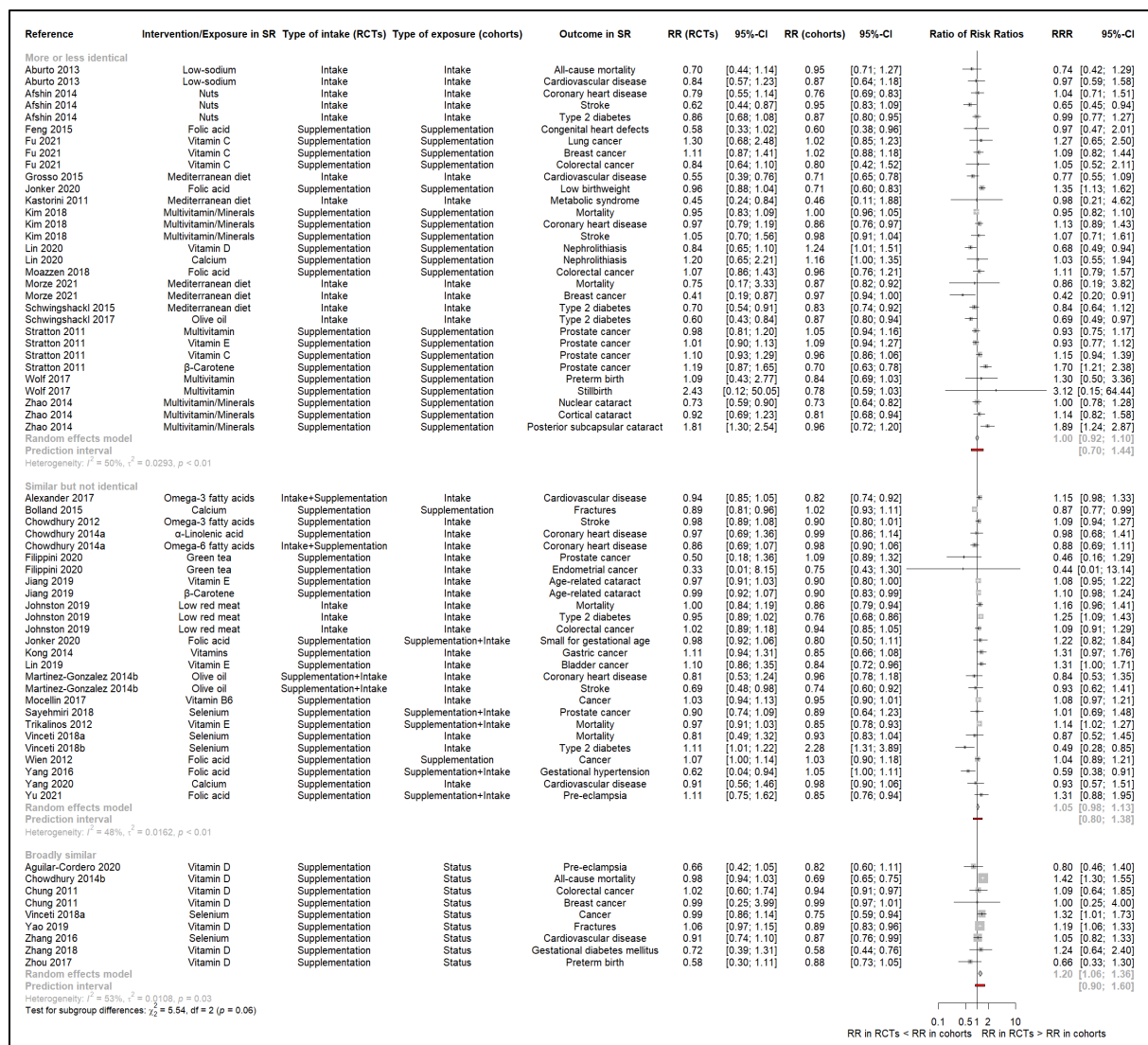
Supplementary Fig. S4 Forest plot, subgroup analysis of BoE-pairs with binary outcomes by type of outcome



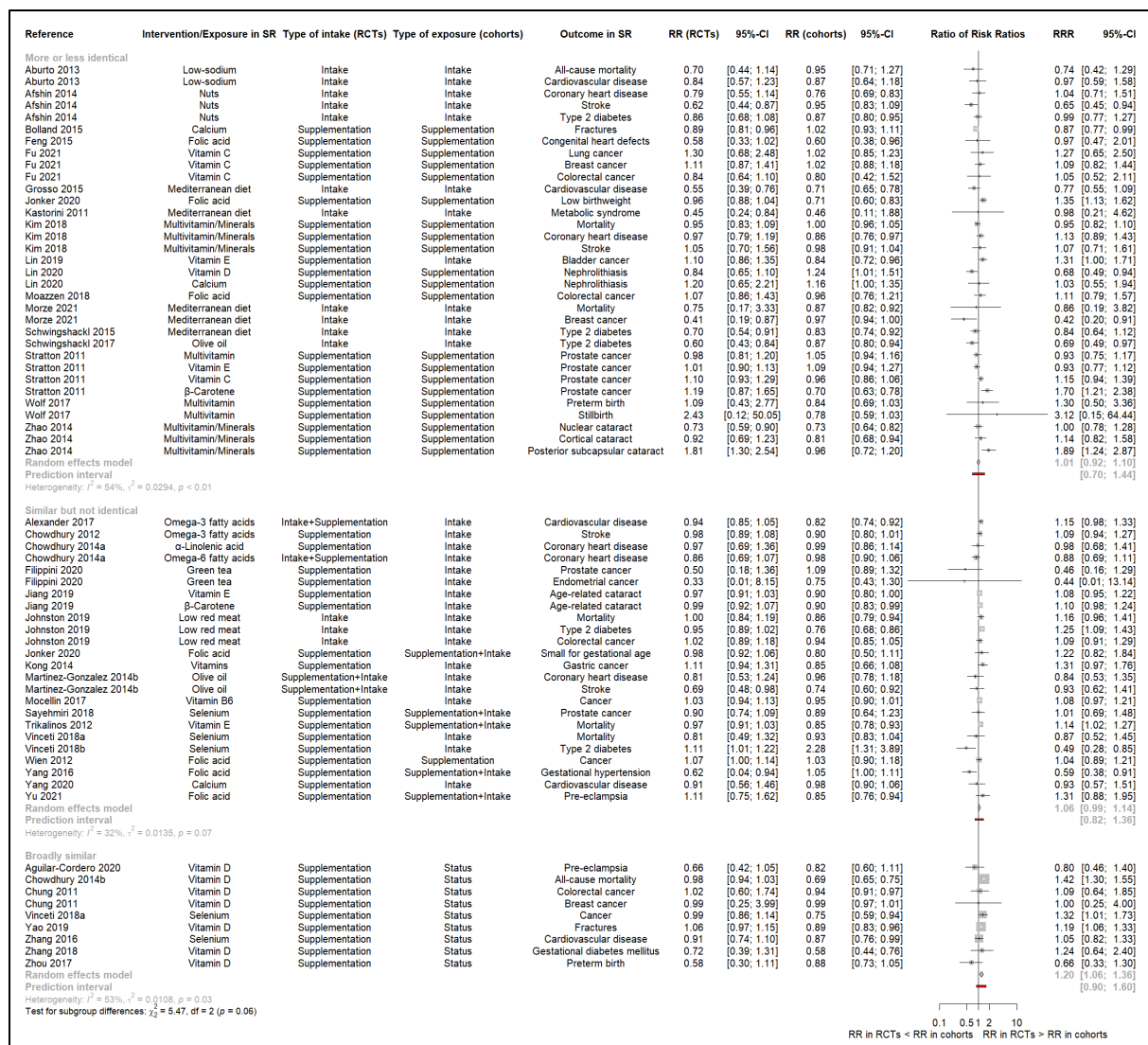
Supplementary Fig. S5 Forest plot, subgroup analysis of BoE-pairs with binary outcomes by overall PI/ECO similarity degree



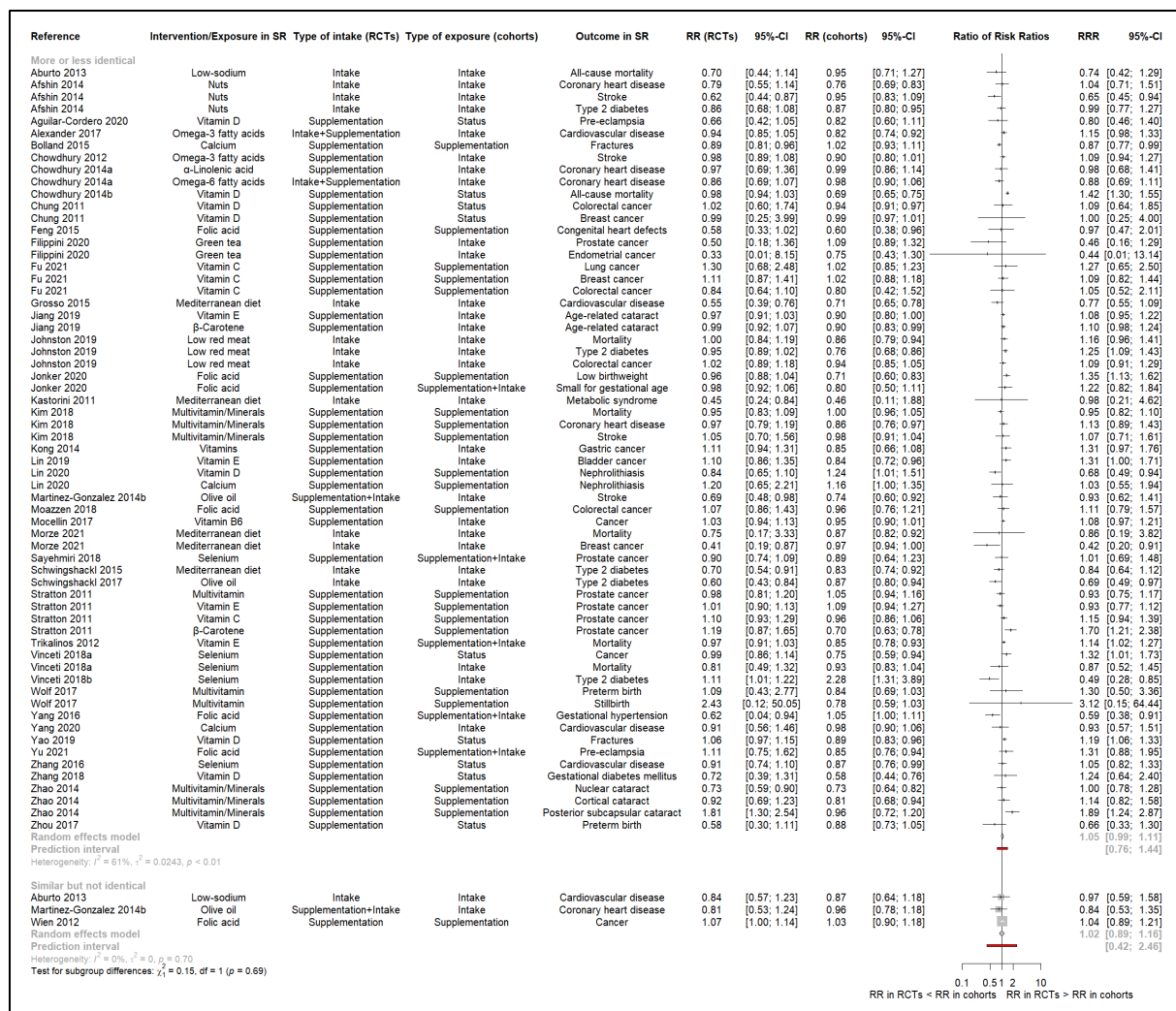
Supplementary Fig. S6 Forest plot, subgroup analysis of BoE-pairs with binary outcomes by population similarity degree



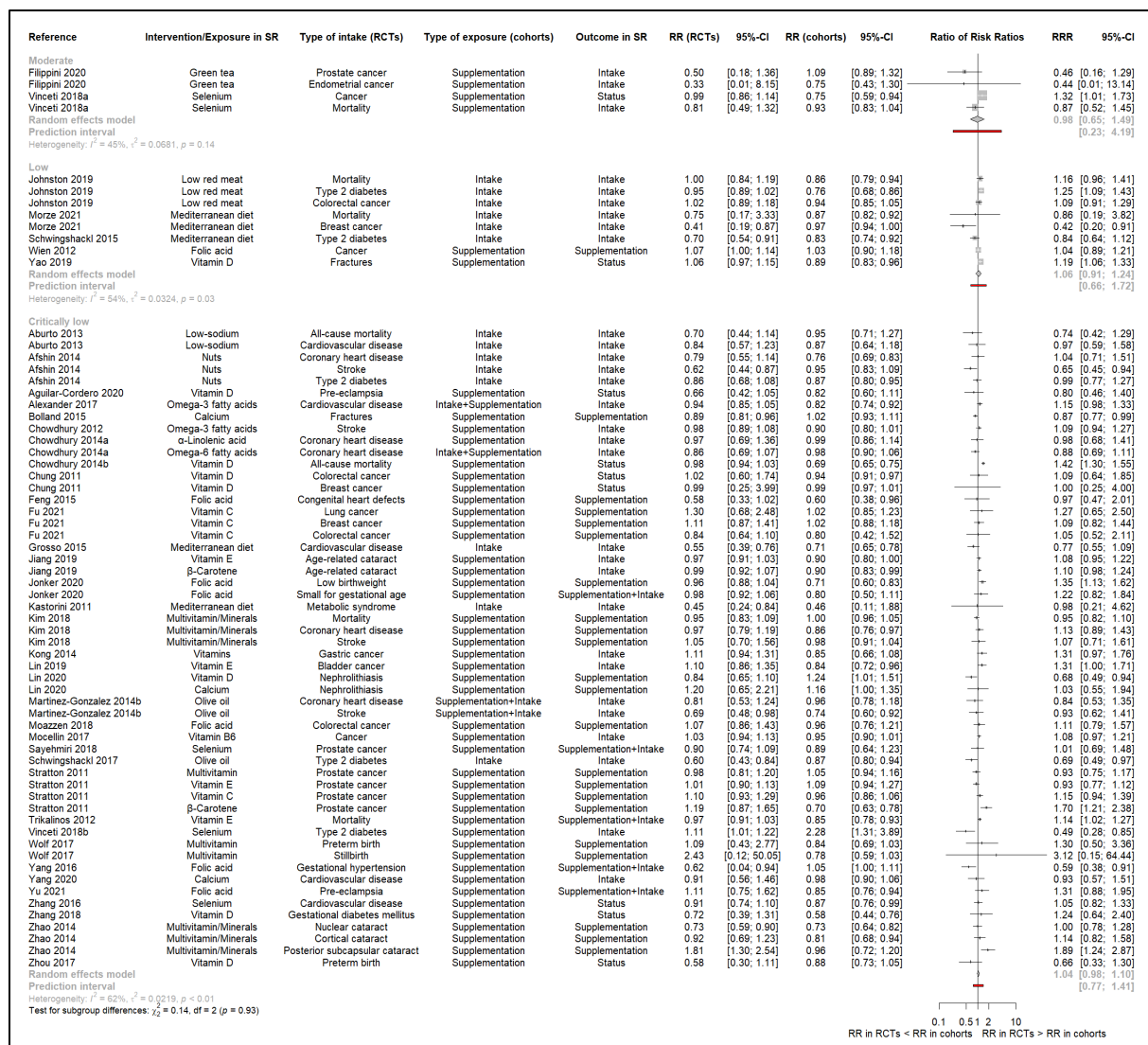
Supplementary Fig. S7 Forest plot, subgroup analysis of BoE-pairs with binary outcomes by intervention/exposure similarity degree



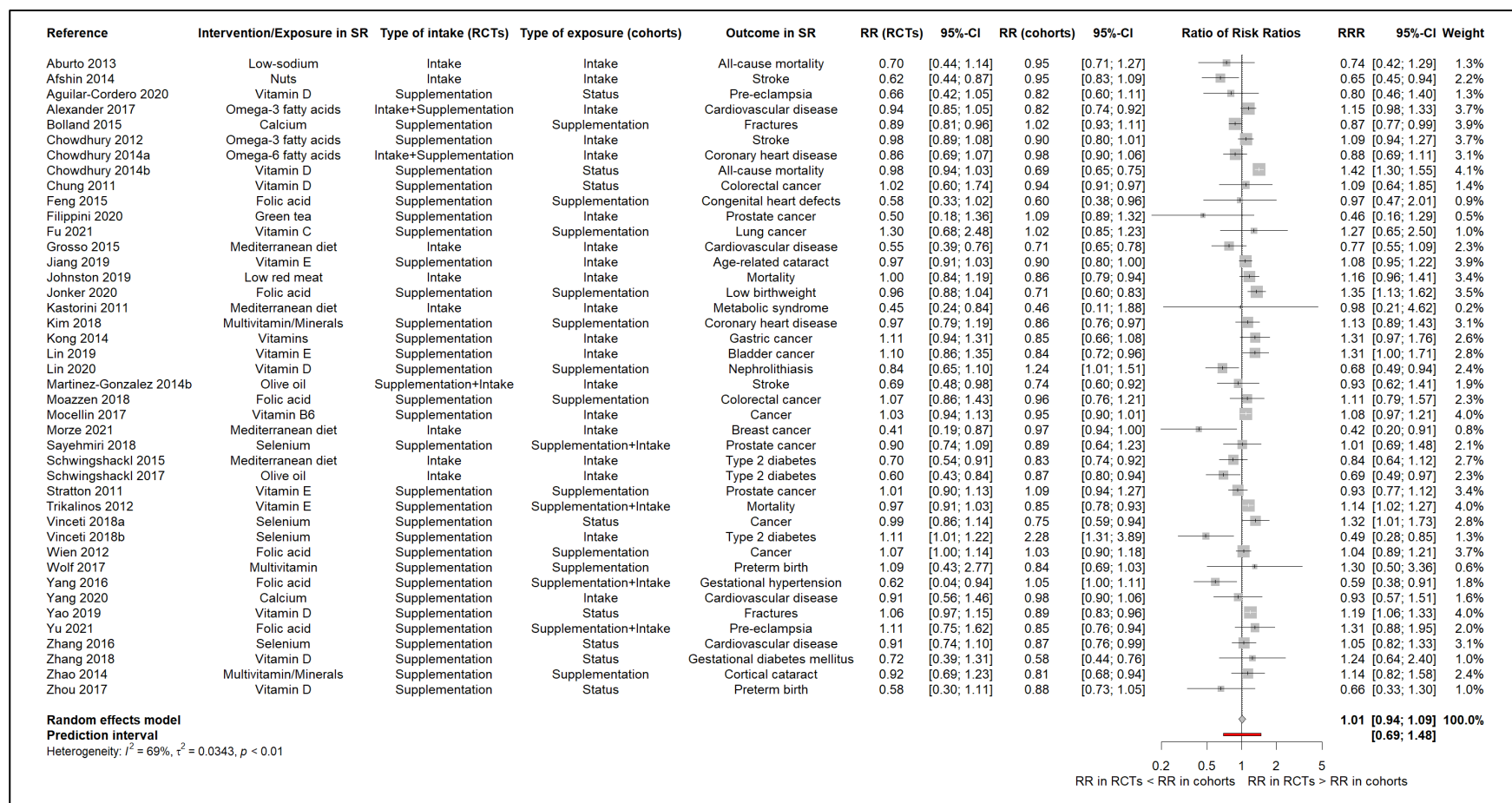
Supplementary Fig. S8 Forest plot, subgroup analysis of BoE-pairs with binary outcomes by comparator similarity degree



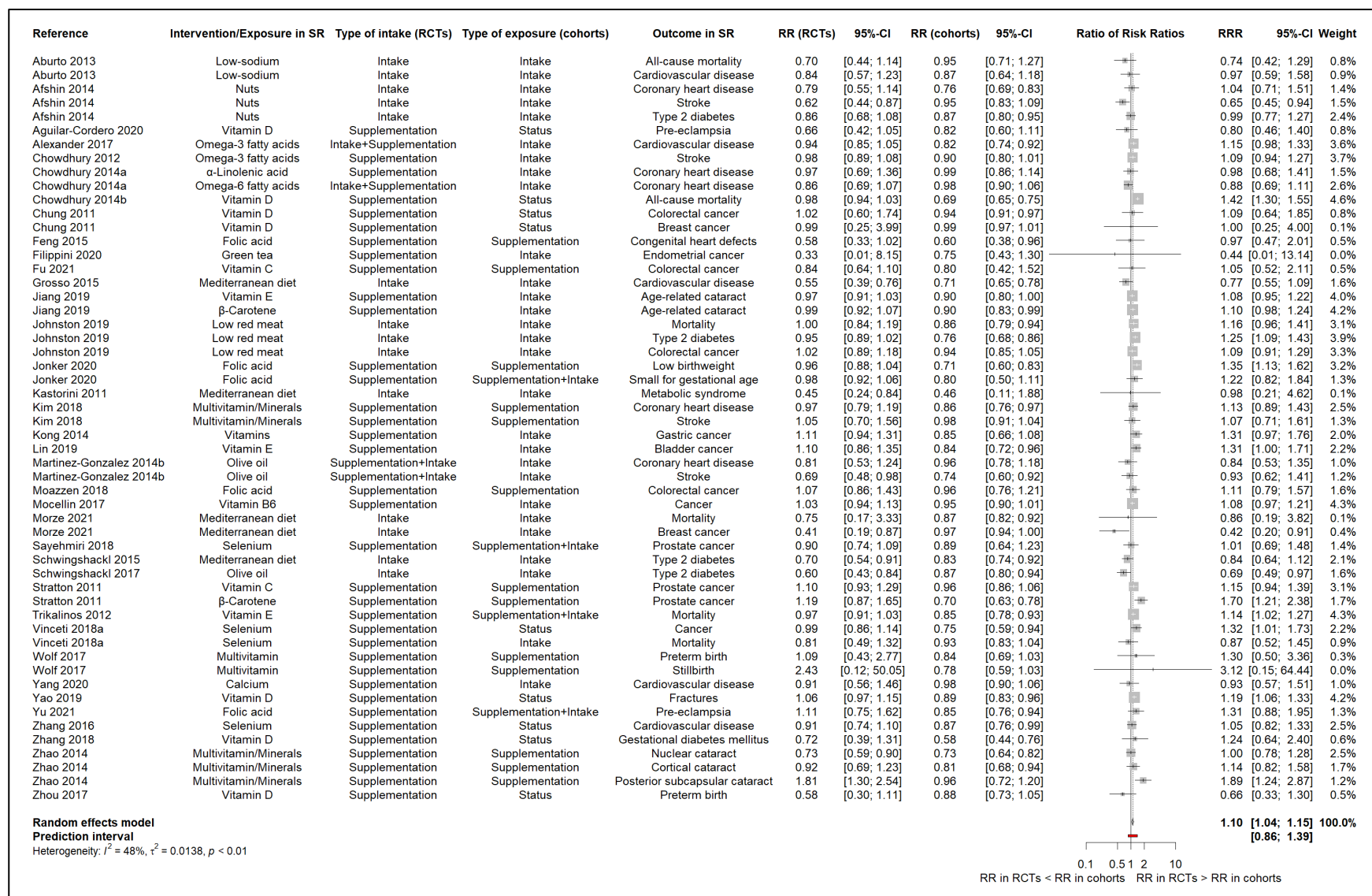
Supplementary Fig. S9 Forest plot, subgroup analysis of BoE-pairs with binary outcomes by outcome similarity degree



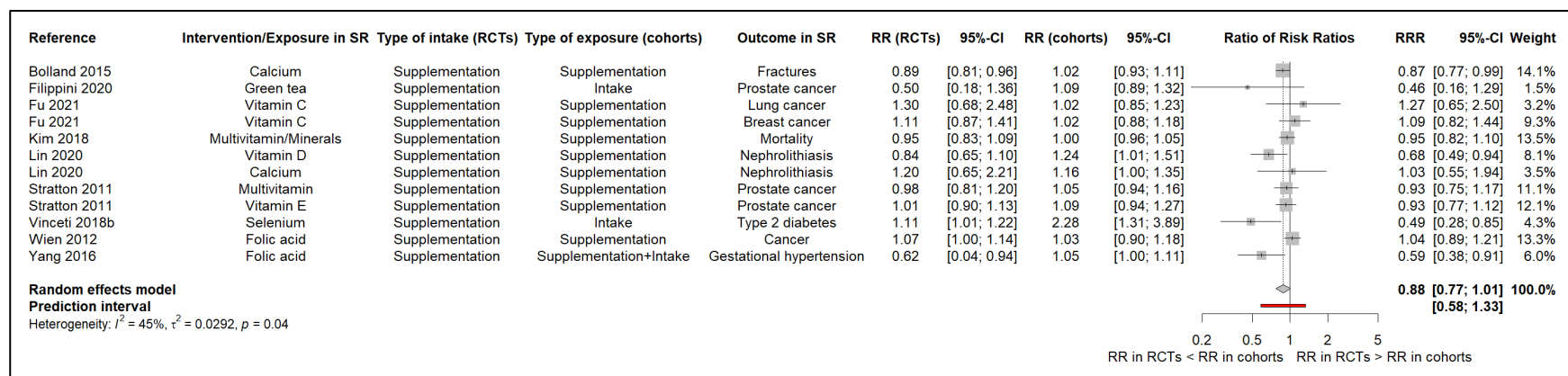
Supplementary Fig. S10 Forest plot, subgroup analysis of BoE-pairs with binary outcomes by AMSTAR 2 rating



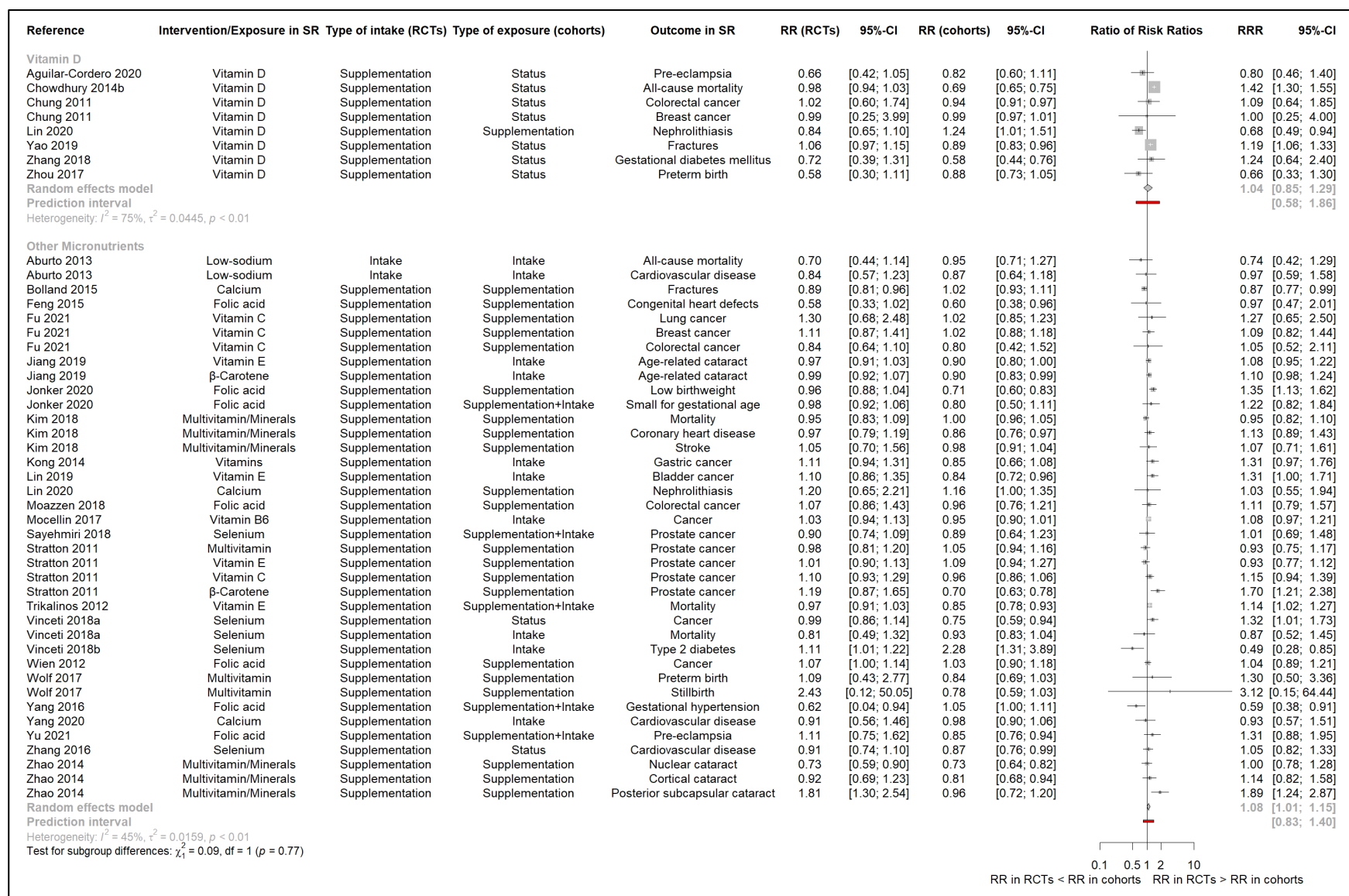
Supplementary Fig. S11 Forest plot, sensitivity analysis including one BoE-pair per systematic review



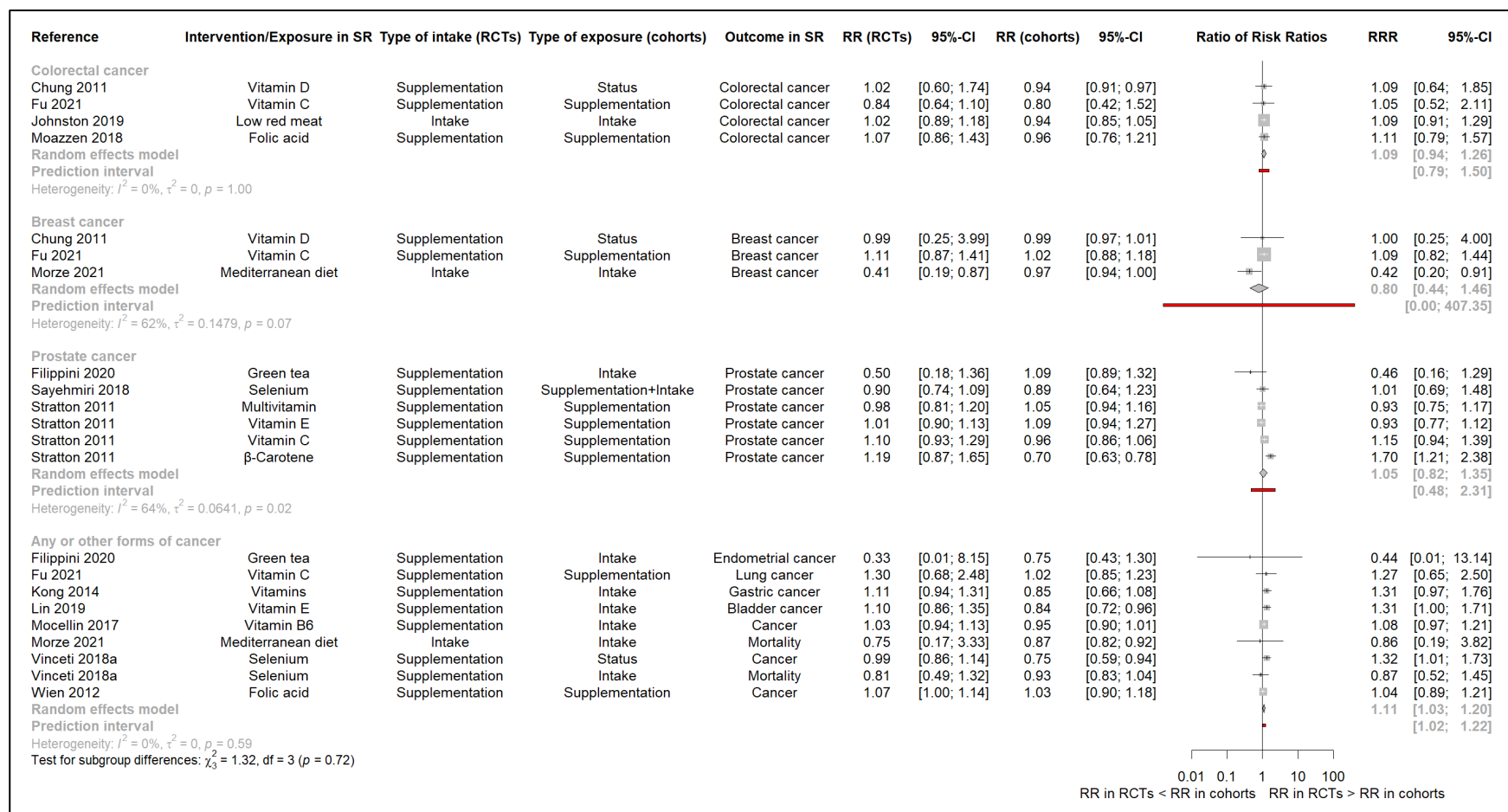
Supplementary Fig. S12 Forest plot, sensitivity analysis by direction of cohort study summary effect estimate (cohort studies with risk ratio [RR] <1)



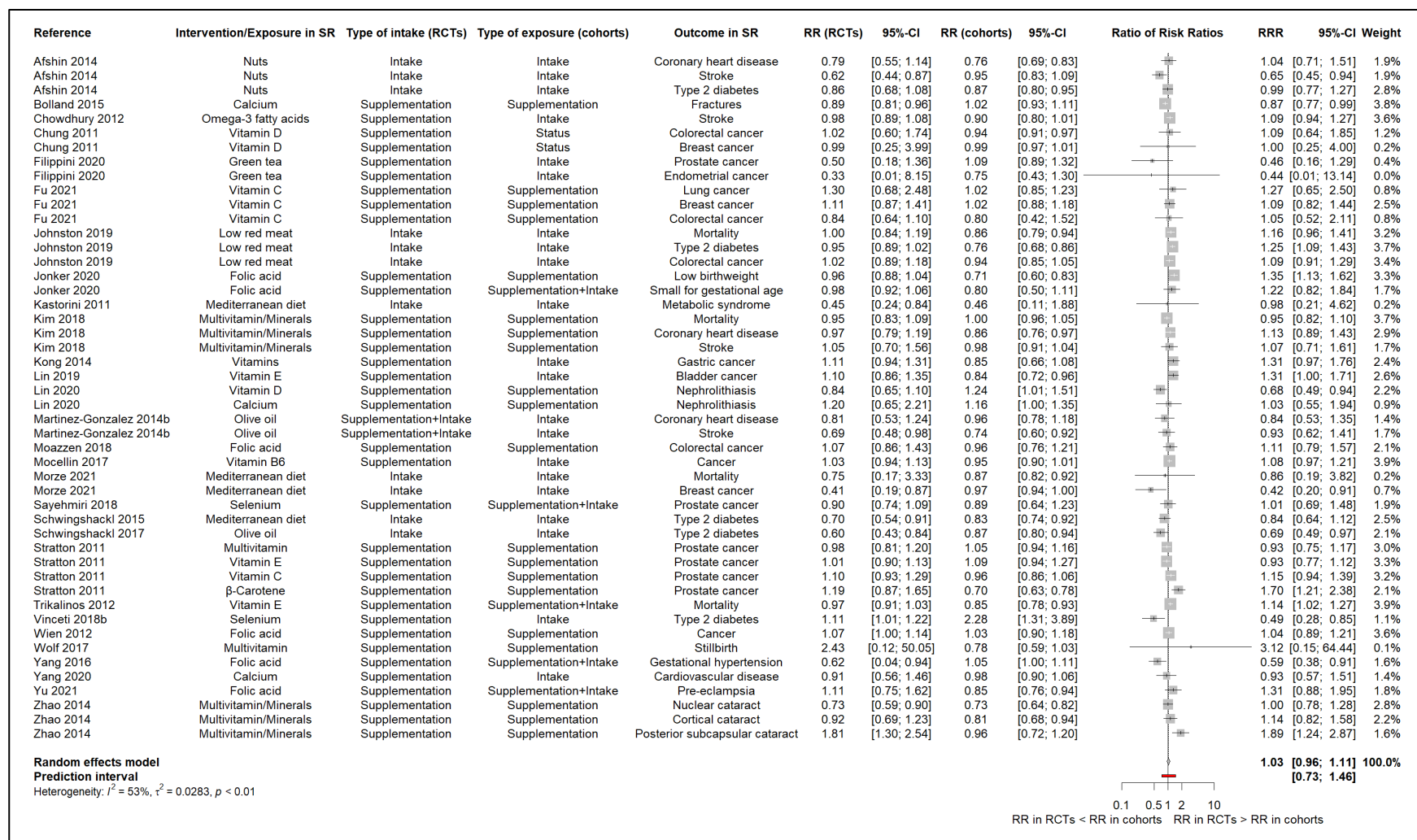
Supplementary Fig. S13 Forest plot, sensitivity analysis by direction of cohort study summary effect estimate (cohort studies with risk ratio [RR] ≥ 1)



Supplementary Fig. S14 Forest plot, sensitivity analysis for interventions with micronutrients



Supplementary Fig. S15 Forest plot, sensitivity analysis for cancer outcomes



Supplementary Fig. S16 Forest plot, sensitivity analysis excluding BoE-pairs with highly similar PI/ECO questions and overlapping primary studies

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