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# **ABBREVIATIONS DEFINITIONS**

BMI: body mass index BP: blood pressure

C.I.: confidence intervals

D.A.S.H.: Dietary Approach to Stop Hypertension

GBD: Global Burden of Disease

IPCC: Intergovernmental Panel on Climate Change

PBD: plant-based diet

PRISMA: Preferred Reporting Items for Systematic Reviews and Meta-Analyses

PROSPERO: International Prospective Register of Systematic Reviews

RCT: randomised controlled trial

SD: standard deviation SE: standard error US: United States

WHO: World Health Organization

# **CONDENSED ABSTRACT**

Vegetarian diets are associated with lower hypertension and cardiovascular risk. However, the effect of less strict plant-based diets (PBDs) on blood pressure (BP) remains unknown. This systematic review and meta-analysis are the most comprehensive synthesis to date on the effect of PBDs on BP. Forty-one clinical trials of various PBDs, including 8,416 participants, were analysed. They included D.A.S.H., Mediterranean, vegan, lacto-ovo vegetarian, Nordic, high fiber and high fruit and vegetables diets. All PBDs reduced BP, with varying potency, the most effective being the D.A.S.H. diet. Any shift along the PBD spectrum provides health benefits, even with the presence of reduced animal products.

The effect of plant-based dietary patterns on blood pressure: a systematic

review and meta-analysis of controlled intervention trials

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**ABSTRACT** 

OBJECTIVES. The consumption of strict vegetarian diets with no animal products is associated with

low blood pressure (BP). It is not clear whether less strict plant-based diets (PBDs) containing some

animal products exert a similar effect. The main objective of this meta-analysis was to assess whether

PBDs reduce BP in controlled clinical trials.

METHODS. We searched Cumulative Index to Nursing and Allied Health Literature, Medline, Embase,

and Web of Science to identify controlled clinical trials investigating the effect of plant-based diets on

BP. Standardised mean differences in BP and 95% C.I. were pooled using a random effects model. Risk

of bias, sensitivity, heterogeneity and publication bias were assessed.

**RESULTS.** Of the 790 studies identified, 41 clinical trials met the inclusion criteria (8,416 participants

of mean age 49.2 yrs). In the pooled analysis, PBDs were associated with lower systolic BP (DASH -5.53

mmHg [95% C.I. -7.95, -3.12], Mediterranean -0.95 mmHg [-1.70, -0.20], Vegan -1.30 mmHg [-3.90,

1.29], Lacto-ovo vegetarian -5.47 mmHg [-7.60, -3.34], Nordic -4.47 mmHg [-7.14, -1.81], high-fiber -

0.65 mmHg [-1.83, 0.53], high fruit and vegetable -0.57 mmHg [-7.45, 6.32]. Similar effects were seen

on diastolic BP. There was no evidence of publication bias and some heterogeneity was detected. The

certainty of the results is high for the lacto-ovo vegetarian and DASH diets, moderate for the Nordic

and Mediterranean diets, low for the vegan diet, and very low for the high fruit and vegetable and

high-fiber diets.

**CONCLUSION.** PBDs with limited animal products lower both systolic and diastolic BP, across sex and

body mass index.

**Keywords**: Plant-based diet, hypertension, blood pressure, Meta-analysis, nutrition

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#### INTRODUCTION

The Global Burden of Disease (GBD) study identified hypertension (high blood pressure) as the global number one risk factor for deaths and disability-adjusted life years (1). Hypertension is accountable for the death of nine million people worldwide every year (2), due to its contribution to a variety of causes of death, including coronary heart disease, stroke, chronic kidney disease, and aneurysms. Hypertension is estimated to contribute 49% of all coronary heart disease and 62% of all stroke events (3). An estimated 1.13 billion people globally have hypertension (4).

The GBD study estimated that increased consumption of whole grains, vegetables, nuts and seeds, and fruit could save 1.7 million, 1.8 million, 2.5 million, and 4.9 million lives per year respectively, through the beneficial effects on cardiovascular risk factors (2). Some epidemiological evidence supports an inverse association between fruit and vegetable consumption and blood pressure (BP)(5-8). There is also evidence of a positive association between meat consumption and hypertension risk (9). Additionally, vegetarian individuals have lower observed rates of ischaemic heart disease than meat and fish eaters (10).

Two meta-analyses have been published in the past few years. One estimated the effect of vegetarian diets on BP (11) in 32 observational studies (totalling 21,604 participants). Consumption of vegetarian diets was associated with a 6.9 mmHg (9.1 to 4.7) lower mean systolic BP and a 4.7 mmHg (6.3 to 3.1) lower mean diastolic BP compared with the consumption of omnivorous diets. For more robust evidence, seven clinical trials (totalling 311 participants) were included in the analysis. In the clinical trials, consumption of vegetarian diets was associated with a 4.8 mmHg (6.6 to 3.1) reduction in mean systolic BP and a 2.2 mmHg (3.5 to 1.0) reduction in mean diastolic BP compared with the consumption of omnivorous diets. A second meta-analysis looked at the BP effect of vegan diets compared to less restrictive diets in 11 randomized controlled trials including 983 participants (12). Vegan diets only reduced BP in participants with a baseline systolic BP >130mmHg (-4.10 mmHg (-8.14 to -0.06) systolic and -4.01 mmHg (-5.97 to -2.05) diastolic BP.

Since then, more controlled trials on the effect of plant-based diets (PBDs) on BP have been published. Therefore, our study undertook a more comprehensive systematic review and a meta-analysis of controlled clinical trials involving not only vegan and vegetarian, but also the Mediterranean diet, Dietary Approaches to Stop Hypertension (DASH) diet, and Nordic diet, plant-based diets that allow limited amount of animal products, to investigate whether complete eradication of animal products is necessary to achieve significant BP lowering effects.

### **METHODS**

This systematic review and meta-analysis is reported in line with the Preferred Reporting Items for Systematic reviews and Meta-analyses (PRISMA) guidelines for randomised controlled trials (13) and is registered with PROSPERO (CRD42019153716).

# Search strategy and selection criteria

We performed a computerised systematic search to identify studies on the effect of plant-based diets on BP. On 14 June 2019 we searched the following electronic databases limited to randomised controlled trials or controlled trials published in the English language since the inception of each database: Cumulative Index to Nursing and Allied Health Literature (CINAHL) (1961-2019), MEDLINE (1964-2019), Embase (1974-2019), and, Web of Science (1900-2019). We used "plant-based diet" terms (plant-based diet OR plant food OR "plant food" OR vegetarian\* OR vegetarian diet OR vegan\* OR vegan diet OR Mediterranean diet OR Nordic diet OR high fiber diet OR DASH diet OR semi-vegetarian OR flexitarian OR pescatarian OR prudent diet OR portfolio diet) in combination with "blood pressure" terms (hypertension OR blood pressure). The electronic search strategy is shown in the supplement (Supplementary Table S1).

#### Inclusion and exclusion criteria

For inclusion, studies had to fulfil the following criteria: (1) original published article; (2) age of participants ≥ 18 years; (3) plant-based diet as an intervention, defined as dietary patterns that

support high consumption of fruits, vegetables, whole grains, legumes, nuts and seeds, and often limit the consumption of most or all animal products; (4) collection of sufficient data to allow calculation of mean differences in systolic/diastolic BP between individuals consuming a plant-based diet and those consuming a referent or control diet (5) Randomised controlled trial or controlled trial study design.

Studies were excluded if (1) multiple interventions were used; (2) study samples overlapped; (3) Plant-based controls were used or uncontrolled; (4) only meeting abstracts or unpublished material available. There were no restrictions regarding sex, race, ethnicity, language, sample size, or publication date. If multiple published reports from the same study were available, we only included the one with the most up to date information regarding the outcome. When data were not readily available from published reports, we wrote to the authors to ask for the data.

#### Data Extraction, Risk of Bias, and Quality Assessment

Two reviewers (J.G. and E.G.) independently extracted the data. Disagreements about the inclusion of studies were resolved by arbitration between co-authors. From a total of 1,238 search records, 790 studies were identified after duplicates had been removed (*Figure 1*). Title and abstract screening were performed using Covidence and resulted in the exclusion of 705 studies. Full-text evaluation of 85 studies identified 41 trials that had data suitable for meta-analysis. Relevant data included, data regarding systolic and diastolic BP and variance measures; first authors surname, year of publication and country of origin; number of participants, study design and duration; baseline characteristics of study population, including mean age, sex (proportion of men), systolic BP, diastolic BP, antihypertensive medication use, body mass index (BMI), alcohol intake, and dietary data (type of intervention and control diets); and outcomes, including adjustment factors used for each analytic model. Mean values for baseline age, the proportion of men, systolic and diastolic BP, BMI, and alcohol intake were calculated. We assessed the risk of bias associated with the method of random sequence generation, allocation concealment, blinding, selective reporting, loss to follow-

up, and completeness of reporting outcome data. We graded the risk of bias as low, unclear, or high according to recognised criteria (14). The certainty of the entire body of evidence was assessed using GRADE methodology (15).

#### Intervention

PBDs were defined as dietary patterns that support high consumption of fruits, vegetables, whole grains, legumes, nuts, and seeds, and often limit the consumption of most or all animal products. Dietary patterns that fall within this umbrella term include vegan, lacto-ovo vegetarian, D.A.S.H., healthy Nordic, Mediterranean, high-fiber, and high fruit and vegetables (*Table 1*)

# **Population**

Our study includes normotensive and hypertensive populations.

#### **Outcome**

The difference in systolic and diastolic BP between plant-based diet and comparator (control) after a period of intervention. Any method of BP measurement was included. The measurements were made by health professionals or by the participants if they were trained on how to do so properly.

# **Data Synthesis and Statistical Analysis**

The mean differences in systolic and diastolic BP between groups consuming plant-based or comparison diets were synthesised, and the standard errors (SEs) were obtained. If the SE of the mean differences was not supplied, it was algebraically computed from the 95% confidence intervals (95% CIs) or standard deviations (SDs). The mean differences for individual studies were pooled, stratified by diet type, using a random-effects model. A sub-group analysis was then carried out, in which only studies with the participants usual/standard diet as the control were included.

Estimates of the overall net change in BP and 95% C.I.s associated with the consumption of each diet type were calculated, and each study was weighted by its inverse variance. The heterogeneity

among studies was quantified by I<sup>2</sup>-statistic. Funnel plots were developed to assess the impact of publication bias. Beggs's test and Egger's regression test were applied to measure funnel plot asymmetry. We conducted a one-study-removed analysis as a sensitivity analysis. This involved omitting one study at a time to assess the impact of each study on the combined effect. Subgroup analyses by mean age, duration of plant-based diet consumption, antihypertensive medication use, baseline hypertensive status, and country/region were performed. Random effects meta-regression was used to determine if age, intervention duration, baseline BMI, or sex were significantly associated with heterogeneity.

#### **RESULTS**

#### **Study selection process**

The search strategy retrieved 1,238 articles. After removing duplicates, the title and abstract screening process identified 85 studies. Full-text assessment led to the exclusion of 44 articles from the systematic review (*Supplementary Table S2*). The remaining 41 articles met the inclusion criteria and had suitable data for meta-analyses (*Figure 1*). Two additional publications were found through reference lists and hand searching.

#### Study characteristics

The 41 included studies were published between 1983 and 2019 (16-56) (*Table 1 and Supplementary Table S3*). The total sample size was 8,416 (4,429 in the intervention groups and 3,987 in the control groups; median sample size 65; range 11-4,717) and the mean age of the participants was 49.2 years (range 25.6-71.0 years). All included studies were controlled trials with a duration range of 1.4 to 208 weeks (median duration 12 weeks). Of the 41 clinical trials, two were not randomised (21, 32). Of the 39 randomised controlled trials (RCTs), 26 reported the method of random generation and 13 failed to describe it (*Supplementary Table S3*). Additionally, seven studies used a crossover design and 33 used a parallel design, of which two of the studies were single-blinded (*Table 1*)(26, 47). Two

of the studies had controlled feeding (23, 33) and all of the studies were free living. As shown in Table 1, 12 studies included participants who were taking antihypertensive medications. Foods were provided to the participants in 20 of the clinical trials. The interventions under investigation in the 41 studies are the D.A.S.H. diet (n = 11), vegan diet (n = 9), Mediterranean diet (n = 8), lacto-ovo vegetarian diet (n = 5), healthy Nordic diet (n = 3), high fiber diet (n = 3), and high fruit and vegetables diet (n = 2) (Table 1). 32 of the clinical trials reported how many BP measurements were taken, of these 31 reported repeated BP measurements. 19 of the studies adjusted for potential confounders (Supplementary Table S3). 32 of the studies reported on the adherence of participants to the dietary interventions (Supplementary Table S3). Of these studies, 26 reported high adherence, four reported fair adherence (32, 38, 39, 44), and two reported poor adherence (37, 48). 60% of included studies indicate a low risk of bias for random sequence generation and allocation concealment. 100% of the studies indicate a high risk of performance bias due to the nature of dietary interventions. 40% of the studies indicate a high risk of detection bias due to the lack of outcome assessor blinding. 90% of the studies indicate a low risk of attrition bias. 42.5% of the studies indicate low risk of reporting bias. Finally, 10% of the studies indicate high risk for funding bias.

### Pooled effects of plant-based diets on blood pressure

Healthy Nordic diet

Compared to reference diets, the healthy Nordic diet involves higher intake of plant foods, fish, egg, and vegetable fat, and lower intake of meat products, dairy products, sweets, desserts, and alcoholic beverages (57). In the three identified randomised controlled trials, consumption of the healthy Nordic diet was associated with a mean reduction in systolic BP (-4.47 mm Hg; 95% CI, -7.14 to -1.81; P = 0.001;  $I^2 = 31\%$ ; P = 0.23 for heterogeneity) (*Figure 2*) and diastolic BP (-2.32 mm Hg; 95% CI, -3.83 to -0.82; P = 0.002;  $I^2 = 0\%$ ; P = 0.39 for heterogeneity) (*Figure 3*) compared with the

consumption of comparator diets. In the one-study-removed analysis, results were mostly unaffected, with BP differences between the healthy Nordic and control groups ranging from -3.75 to -5.64 mm Hg for systolic BP and -1.75 to -3.30 mm Hg for diastolic BP (*Supplementary Table S4*). The certainty of this evidence is moderate. (*Table 3*)

High fruit and vegetable diet

The high fruit and vegetable diet is characterised by increased consumption of fruit and vegetables. To further increase the polyphenolic load, one of the studies included regular dark chocolate intake (47). In the two clinical trials, consumption of the high fruit and vegetables diet was associated with a mean reduction in systolic BP (-0.57 mm Hg; 95% CI, -7.45 to 6.32; P = 0.87;  $I^2 = 65\%$ ; P = 0.09 for heterogeneity) (*Figure 2*) and diastolic BP (-0.96 mm Hg; 95% CI, -3.08 to 1.15; P = 0.37; P = 0.37; P = 0.43 for heterogeneity) (*Figure 3*) compared with the consumption of comparator diets. This subgroup was not suitable for a one-study-removed analysis as it only comprised two studies. Overall, the certainty of this evidence is very low (*Table 3*).

High fiber diet

Fiber is found in varying levels in all plant foods and is most prevalent in whole grains and legumes. For this reason, most high fiber diets focus on increasing wholegrain and legume consumption (36). In the three controlled trials, consumption of the high fiber diet was associated with a mean reduction in systolic BP (-0.65 mm Hg; 95% CI, -1.83 to 0.53; P = 0.28; I<sup>2</sup> = 0%; P = 0.39 for heterogeneity) (*Figure 2*) and diastolic BP (-1.02 mm Hg; 95% CI, -3.86 to 1.82; P = 0.0.48; I<sup>2</sup> = 75%; P = 0.02 for heterogeneity) (*Figure 3*) compared with the consumption of comparator diets. The one-study-removed analysis identified the study of Morenga *et al* (36) as a source of heterogeneity. Removal of this study reduced the diastolic BP effect heterogeneity from 75% to 0% (*Supplementary Table S4*). The mean differences produced by this removal were -0.37 (-1.61, 0.87) and 0.24 (-0.92, 1.40) mm Hg for systolic and diastolic BP respectively (*Supplementary Table S5*). The certainty of this evidence is very low (*Table 3*).

#### Lacto-ovo vegetarian diet

Lacto-ovo vegetarian dietary patterns are defined as those that exclude the consumption of all meat, poultry, and fish but still include the consumption of dairy and eggs (20). The main components of the lacto-ovo vegetarian diets included in this study are fruit, vegetables, whole grains, legumes, and nuts and seeds. In the five clinical trials, consumption of the lacto-ovo vegetarian diet was associated with a mean reduction in systolic BP (-5.47 mm Hg; 95% Cl, -7.60 to -3.34; P < 0.00001;  $I^2 = 0\%$ ; P = 0.58 for heterogeneity) (*Figure 2*) and diastolic BP (-2.49 mm Hg; 95% Cl, -4.17 to -0.80; P = 0.004;  $I^2 = 0\%$ ; P = 0.97 for heterogeneity) (*Figure 3*) compared with the consumption of comparator diets. There was no overall heterogeneity for the lacto-ovo vegetarian results. The certainty of the systolic BP result is high; however, the certainty of the diastolic BP result is moderate (*Table 3*). It is to note that about 50% of the contribution to the overall estimate was weighted in favour of a single study (16).

#### D.A.S.H. diet

The D.A.S.H. diet encourages the consumption of fruits, vegetables, whole grains, nuts and seeds, and low-fat dairy products and limits the intake of sweets, saturated fat, and sodium (58, 59). Two of the studies did not reduce participant sodium intake (32, 51). In the 11 identified clinical trials, consumption of the D.A.S.H. diet was associated with a mean reduction in systolic BP (-5.53 mm Hg; 95% CI, -7.95 to -3.12; P < 0.00001; I² = 84%; P < 0.00001 for heterogeneity) (*Figure 2*) and diastolic BP (-3.78 mm Hg; 95% CI, -5.51 to -2.04; P < 0.0001; I² = 84%; P < 0.00001 for heterogeneity) (*Figure 3*) compared with the consumption of comparator diets. Removal of three studies (21, 35, 44) reduced the systolic BP effect heterogeneity from 84% to 0% and changed the mean reduction in systolic BP to -4.70 (-5.76, -3.63) mm Hg (*Supplementary Table S5*). Removal of five studies (21, 30, 35, 44, 52) also reduced the diastolic BP effect heterogeneity from 84% to 0% and changed the mean reduction in diastolic BP to -3.75 (-4.53, -2.97) mm Hg (*Supplementary Table S5*). The certainty of this evidence is high (*Table 3*). Finally, the D.A.S.H. diet was implemented with either a fixed moderate

sodium consumption (23, 33, 35, 54) or with tips given to participants to reduce sodium consumption (21, 27, 44, 52). We carried out a sensitivity analysis between the two groups of trials and did not detect a significant difference in the estimates of effects on blood pressure (for systolic - 6.45 [-9.34 to -3.55] vs -4.25 [-9.44 to 0.95] mmHg, p for interaction = 0.47; for diastolic -3.95 [-6.64 to -1.26] vs -3.63 [-6.92 to -0.33] mmHg, p for interaction = 0.88, respectively). These results do not detect the well-known additive blood pressure-lowering effect of sodium reduction to the core D.A.S.H. diet (60). This could be due to the fact that simple tips to reduce sodium intake may not have led to a true reduction in consumption, evidence not available in the individual trials as sodium excretion was not measured.

#### Mediterranean diet

The main components of the Mediterranean diet are daily consumption of vegetables, fruit, whole grains, olive oil, weekly consumption of legumes, nuts, fish, dairy, and eggs, and limited intake of meat (48). In the eight clinical trials, consumption of the Mediterranean diet was associated with a mean reduction in systolic BP (-0.95 mm Hg; 95% CI, -1.70 to -0.20; P = 0.01; I² = 38%; P = 0.13 for heterogeneity) (*Figure 2*) and diastolic BP (-0.69 mm Hg; 95% CI, -1.44 to 0.06; P = 0.07; I² = 68%; P = 0.003 for heterogeneity) (*Figure 3*) compared with the consumption of comparator diets. Removal of two studies (26, 55) reduced the systolic BP effect heterogeneity from 38% to 0% and changed the mean reduction in systolic BP to -0.97 (-1.58, -0.36) mm Hg (*Supplementary Table S5*). Removal of two different studies (31, 48) reduced the diastolic BP effect heterogeneity from 68% to 0% and changed the mean reduction in diastolic BP to -0.61 (-0.96, -0.26) mm Hg (*Supplementary Table S5*). The certainty of this evidence is moderate (*Table 3*).

### Vegan diet

Vegan diets consist of plant foods exclusively. No animal flesh or other animal-derived products (including dairy and eggs) are included in the diet. The vegan diets included in this study are mostly low-fat and focus on the consumption of whole plant foods like fruits, vegetables, whole grains,

legumes, and nuts and seeds (50). In the nine controlled trials, consumption of the vegan diet was associated with a mean reduction in systolic BP (-1.30 mm Hg; 95% CI, -3.90 to 1.29; P = 0.33; I<sup>2</sup> = 26%; P = 0.21 for heterogeneity) (*Figure 2*) and diastolic BP (-0.81 mm Hg; 95% CI, -2.91 to 1.28; P = 0.45; I<sup>2</sup> = 51%; P = 0.04 for heterogeneity) (*Figure 3*) compared with the consumption of comparator diets. In the one-study-removed analysis, systolic BP results had some diversity, with systolic BP differences between the vegan and control groups ranging from 0.05 to -2.49 mm Hg (*Supplementary Table S4*). Removal of one study (32) reduced the systolic BP effect heterogeneity from 26% to 0% and changed the mean reduction in systolic BP to 0.05 (-1.94, 2.03) mm Hg (*Supplementary Table S5*). Removal of the same study reduced the diastolic BP effect heterogeneity from 51% to 0% and changed the mean reduction in diastolic BP to 0.08 (-1.23, 1.38) mm Hg (*Supplementary Table S5*). The certainty of this evidence is low (*Table 3*).

# Meta-regression

The meta-regression identified age as a potential source of heterogeneity in the diastolic BP mean differences obtained from the clinical trials investigating the Mediterranean diet ( $\beta$  coefficient, 0.081; P = 0.049) (*Supplementary Table S6*). Intervention duration, baseline BMI, and sex (proportion of men) were not significant sources of heterogeneity for any of the dietary interventions (*Supplementary Table S6*). These results suggest that the mean reduction in diastolic BP associated with the consumption of the Mediterranean diet is less pronounced among older individuals.

#### **Publication Bias**

The Egger's and Begg's statistical tests found no significant funnel plot asymmetry for any of the dietary interventions (*Supplementary Table S7*).

# **Standardised Control Diet Analysis**

We carried out a secondary analysis including only trials that employed the habitual diet of the participants or average diet of the specific population as the control diet, in an attempt to

standardise control groups (*Table 4*). Compared with the consumption of the standardised control diet, consumption of plant-based diets was associated with a mean reduction in systolic BP (-4.29 mm Hg; 95% CI, -6.27 to -2.31; P = < 0.0001;  $I^2 = 87\%$ ; P = < 0.00001 for heterogeneity) and diastolic BP (-2.79 mm Hg; 95% CI, -4.33 to -1.24; P = 0.0004;  $I^2 = 88\%$ ; P = < 0.00001 for heterogeneity).

# **DISCUSSION**

The results of our study show, with varying certainty, that plant-based dietary patterns reduce systolic and diastolic BP.

The Healthy Nordic and Mediterranean diets produce statistically significant reductions in systolic BP. The certainty of this evidence is moderate. This finding is of great significance as it shows that complete eradication of animal products from one's diet is not necessary to produce significant improvements in BP. Therefore, these diets can be considered as achievable lifestyle modifications for those trying to lower their BP.

Our results show with high certainty that both the lacto-ovo vegetarian and DASH diets significantly reduce BP. This confirms the results of a previous meta-analysis of clinical trials and observational studies that found vegetarian dietary patterns are effective at reducing BP (11). Our results are also in accord with another meta-analysis which found that the D.A.S.H., Mediterranean, and Nordic diets are effective at lowering BP (61). These results reinforce the concept that complete eradication of animal products is not necessary for BP reduction, and also add that dietary salt reduction is a powerful tool in adjunct with increased plant-food consumption.

The vegan diet did not significantly reduce BP; however, the certainty of this result is low. This result is in line with a recent meta-analysis suggesting that the changes in BP induced by a vegan diet without caloric restrictions are comparable with those induced by other dietary approaches recommended by medical societies (12). On the other hand, it is likely that the effectiveness of vegan diets at lowering BP has been underestimated by the use BP lowering comparator diets. When

only including the vegan studies with the participants usual diet as the comparator, the overall effect estimate becomes statistically significant, but the certainty remains low.

The results for the high fruit and vegetables and high-fiber diets had a very low certainty, largely due to the small number of studies identified for these interventions. Due to this limitation, it is difficult to determine whether simply increasing fruit and vegetable or whole-grain consumption is sufficient to produce a significant reduction in BP. Since these diets may be the most achievable for the general population to adhere, it is imperative that further controlled trials are conducted to confidently establish the effect of consumption on BP.

Our study shows that the healthy Nordic, ovo-lacto vegetarian, and D.A.S.H. diets are more effective at reducing BP than the Mediterranean diet, since the 95% confidence intervals of all three diets do not overlap with the 95% confidence interval of the Mediterranean diet effect estimate.

Consistent with our findings, an analysis of three prospective cohorts (Nurses' Health Study I, Nurses' Health Study II, and Health Professionals Follow-up Study) totalling 188,518 participants, found a positive association between animal flesh consumption and hypertension risk, independent of fruit, vegetable, and whole-grain consumption (9). Similarly, compared with vegetarians, fish eaters, and meat eaters, vegans had the lowest prevalence of hypertension in a cross-sectional analysis of the European Prospective Investigation into Cancer and Nutrition-Oxford study (11,004 participants) (62). In a calibration sub-study of the Adventist Health Study-2, the BP of habitual vegans, lacto-ovo vegetarians, and non-vegetarians was compared for the first time in the literature (63). The analysis found that vegans and lacto-ovo vegetarians had significantly lower systolic and diastolic BP, as well as significantly lower odds of hypertension (63% and 43%, respectively) when compared with non-vegetarians. This is important since non-vegetarian Seventh Day Adventists often consume less meat than individuals consuming a typical western diet (64).

# **Strengths and Limitations**

This review has six key strengths: 1) it is the first review to have a comprehensive inclusion of all diets with a plant-based component; 2) the standardised control diet analysis allowed us to broadly compare the effect of consuming plant-based diets versus the standard control diet on BP, and to specifically identify which plant-based sub-diets are optimal for lowering BP; 3) the included trials provided a moderately large sample size that promotes confidence in the results; 4) 95% of the included trials were RCTs; 5) there was a lack of detectable publication bias for the included studies; 6) the studies responsible for heterogeneity were identified and the results were largely unaffected by their exclusion.

Some limitations of this review should be noted. Firstly, there was a low number of clinical trials investigating the healthy Nordic diet, high-fiber diet, and high fruit and vegetables diet. This issue was exacerbated when standardising for the control diet. Secondly, this review carried forward the design limitations of the included clinical trials. Most prominent in this regard is small sample sizes. Thirdly, some of the clinical trials were of poor quality mainly due to lack of blinding of study personnel. Due to the nature of dietary interventions, double blinding was not possible in any of the included clinical trials. Fourthly, some of the clinical trials did not adjust the BP outcomes for confounding factors. Finally, the food and nutrient compositions of the diets used in each clinical trial varied, so the effect of individual nutrients could not be identified.

# **Potential Mechanisms**

This review supports a causal relationship between the consumption of plant-based diets and subsequent reduction in systolic and diastolic BP. There are numerous lines of evidence to suggest possible mechanisms. First, PBD eaters have improved endothelial function compared with omnivores (65), due to two possible mechanisms. Animal fat transports bacterial endotoxins into the bloodstream which elicits an inflammatory response (66). This inflammation can impair endothelial function within a few hours of animal fat consumption, thus worsening the ability of blood vessels to dilate (67). A lower fat content can then be contributing to improved endothelial function.

Furthermore, flavonoid-rich fruits and nitrate-rich vegetables can increase plasma nitric oxide concentrations, which improves endothelial function and decreases BP within hours of consumption (68). Second, due to the low energy density of whole plant foods, PBD eaters usually have lower BMIs and lower obesity risk compared with omnivores (69). However, this is unlikely to be the only mechanism responsible for the BP reduction produced by PBDs as trials that maintain body weight still demonstrate a BP-lowering effect (14). Third, PBDs are rich in potassium. Meta-analyses of randomised controlled trials investigating the effect of potassium supplementation on BP found that increased potassium intake reduces BP and risk of strokes (70). High potassium intake may achieve BP reduction through many mechanisms, including, vasodilation, increased GFR, and decreased renin, renal sodium reabsorption, reactive oxygen species production, and platelet aggregation (71). Additional cerebrovascular benefits have also been described in animal experiments, such as increased luminal and outer diameter of cerebral arteries and reduced cerebral infarct size due to potassium supplementation (72). Fourth, PBDs may have a lower sodium content compared with the standard western diet. It is estimated that three-quarters of an individual's sodium intake comes from processed foods (73), therefore, switching one's calorie source to whole plant foods may lead to decreased sodium intake. Alternative potential mechanisms include greater antioxidant and antiinflammatory effects, improved insulin sensitivity, decreased blood viscosity, altered baroreceptors, modifications in both renin-angiotensin, and sympathetic nervous systems, modification of the gut microbiota (74).

# **Implications**

Raised BP is the leading risk factor for mortality globally, accounting for about 12.8% of all deaths (4). The decrease in BP caused by the consumption of PBDs can have important health benefits at the population level. According to McPherson *et al* (75), a 5 mmHg reduction in systolic BP in the population of the United Kingdom would reduce the prevalence of hypertension by an estimated 50% in that country. A systolic BP reduction of this scale is also expected to result in a 7%, 9%, and

14% overall reduction in mortality due to all causes, coronary heart disease, and stroke, respectively (76).

The health benefits of PBDs stretch beyond improved BP. The EAT-Lancet Commission on healthy diets for sustainable food systems highlights the fact that unhealthy diets represent a greater risk of morbidity and mortality than does unsafe sex, and alcohol, drug, and tobacco use combined (77). In an analysis of the PREDIMED study that assigned the diets of the participants with a pro-vegetarian score, the highest scoring group of participants achieved a 41% reduction in mortality compared with the lowest scoring group (78). Similarly, in an analysis of nearly 25,000 participants from the National Health and Nutrition Examination Survey, Mazidi *et al.* found that participants with the lowest carbohydrate intake had the highest risk of overall (32%), CVD (50%), cerebrovascular (51%), and cancer (36%) mortality (79). PBDs are associated with a lower risk of overweight and/or obesity (15%) (80), type 2 diabetes (23%) (81), cardiovascular disease (16%), cardiovascular disease mortality (31-32%), and all-cause mortality (18-25%) (82). Other meta-analyses of clinical trials have found that PBDs significantly reduce HbA1c (83), LDL cholesterol (84), and body weight (85). Therefore, PBDs are a useful tool for disease prevention, and they may also be clinically relevant in the treatment of some noncommunicable diseases, for example coronary artery disease (86), Type 2 diabetes (87), and prostate cancer (88).

Plant-based dietary patterns also play an important role in global food sustainability and security (77). According to the Intergovernmental Panel on Climate Change (IPCC), if we switched to a 100% plant-based food system in 2050, adequate food production could be achieved on less land than is currently used (89). This is not surprising considering that more than half of the world's crops are used to feed animals, not people (90). It is estimated that the livestock sector accounts for 80% of total anthropogenic land use (91). The Livestock sector is also a significant burden on the fresh water supply. Agriculture consumes about 70% of global fresh water (92). Approximately 43,000 L of water is required to produce 1 kg of beef but in contrast, it only takes 1000 L to produce 1 Kg of grain (92).

Therefore, PBDs may play a pivotal role in water conservation. The livestock sector has massive implications on global warming. It is accountable for approximately 18% of global greenhouse gas emission (90). The IPCC reported that the vegan diet is the most powerful diet at mitigating greenhouse gas emission, and estimated that the adoption of a 100% plant-based food system in 2050 would save about 8 Gt CO<sub>2</sub>-eq yr<sup>-1</sup> (88). Recently, Eshel *et al* (93) have estimated that Americans can eliminate land-use for pasture, whilst simultaneously saving 35-50% of their dietrelated needs for cropland, reactive nitrogen, and greenhouse gas emission if all US meat is replaced with plant alternatives.

#### **Barriers**

Whilst our study supports the concept that PBDs are efficacious in lowering blood pressure, the success of a dietary intervention aiming at reducing blood pressure in healthy populations or specific patient groups (effectiveness) depends on a variety of factors related to both individual behaviours and to policy approaches. Socio-demographic factors determine an individual's ability to adopt a PBD. A study using data from 1,890 Finns found that the most important barrier to following a PBD is related to meat appreciation (94). The preference for familiarity and the perceived nutritional necessity of meat contributes greatly to the barrier effect. The association of meat consumption with masculinity and the perceived difficulty of preparing plant-based meals also adds resistance to change. Other barriers preventing people from following a PBD are rural residence, low education, and young age. In another study conducted in the United Kingdom, fruit and vegetable expense was also found as a barrier to increased plant-food consumption (95). To overcome these barriers, we ought to formulate strategies to influence beliefs about PBDs, plant food availability, and cost of plant foods.

PBDs are generally assumed to have lower adherence and acceptability rates than more typical omnivorous therapeutic diets. Evidence from controlled clinical trials, however, suggests a more complex issue. In a randomised trial of 63 overweight and obese patients allocated to a variety of

PBDs compared to an omnivorous diet, there was no significant difference in dietary acceptability and/or adherence between the dietary patterns after six months when validated measures of dietary acceptability and adherence were applied (96). Regardless of the low adherence rates amongst participants, non-adherent vegan and vegetarian participants experienced greater weight loss than non-adherent omnivorous participants. A systematic review found similar results in interventions lasting more than a year (97). Adherence rates ranged from 51% to 61% for vegan and vegetarian diets and 20% to 55% for omnivorous diets. There was no difference in acceptability across diets. The same review also found that the consumption of vegan diets improved quality of life. Individuals prescribed these diets reported weight loss, increased energy, decreased menstrual pain, and improved digestion and sleep. These general improvements in well-being likely influence the acceptability of plant-based diets. Finally, nutritional interventions in treated hypertensives, predominantly based on weight, sodium and alcohol reduction, have been effective in reducing the use of anti-hypertensive medications over a 4-year period (98). Likewise, increasing potassium consumption with food led to the same result over a year in an Italian trial (99). Finally, in large population experiments in Finland (North Karelia) have demonstrated the feasibility of substantive dietary changes with sustained beneficial health effects over decades (100). Longer-term trials on PBDs are nevertheless warranted to explore the generalisability and applicability of this specific dietary approach.

#### **CONCLUSION**

A shift towards healthy diets globally requires focus on environmental sustainability of food production and health consequences of final consumption, requiring multisectoral actions, science and evidence-gathering and a full range of policy changes. A healthy reference diet has been suggested (77). It would largely consist of an increase in plant-based foods with limited or no animal products. Our study provides new comprehensive evidence to support this pledge, indicating that

such diets would significantly lower both systolic and diastolic BP, across sex and body mass index, with likely health benefits on a global scale.

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# **Legend to figures**

Figure 1. PRISMA flow chart.

**Figure 2.** The effects of various plant-based diets on systolic BP. Results are expressed as mean difference (95% confidence interval).

**Figure 3.** The effects of various plant-based diets on diastolic BP. Results are expressed as mean difference (95% confidence interval).

**Table 1.** Study designs and participant characteristics of clinical trials included in the meta-analysis.

Plant-based diet	Principal components
Healthy Nordic diet	higher content of plant foods, fish, egg, and vegetable fat, and lower content of meat products, dairy products, sweets, desserts, and alcoholic beverages
High fruit and vegetable diet	increased consumption of fruit and vegetables. To further increase the polyphenolic load, some studies included regular dark chocolate content
High fiber diet	fiber is found in varying levels in all plant foods and is most prevalent in whole grains and legumes. For this reason, most high fiber diets focus on increasing wholegrain and legume consumption
Lacto-ovo vegetarian diet	defined as those that exclude the consumption of all meat, poultry, and fish but still include the consumption of dairy and eggs. The main components include fruit, vegetables, whole grains, legumes, and nuts and seeds
DASH diet	encourages the consumption of fruits, vegetables, whole grains, nuts and seeds, and low-fat dairy products and limits the intake of sweets, saturated fat, and sodium
Mediterranean diet	the main components are daily consumption of vegetables, fruit, whole grains, olive oil, weekly consumption of legumes, nuts, fish, dairy, and eggs, and limited intake of meat
Vegan diet	consists of plant foods exclusively. No animal flesh or other animal-derived products (including dairy and eggs) are included. It is mostly low-fat and focuses on the consumption of whole plant foods like fruits, vegetables, whole grains, legumes, and nuts and seeds

Abbreviations: BP, blood pressure; BMI, body mass index (weight in kilograms divided by height in meters squared); C, crossover; Cont, control; CT, controlled trial; F&V, fruit and vegetables; HF, high fiber; LF, low-fat; MED, Mediterranean; NR, not reported; O, open label; P, parallel; US, United states; RCT, randomised controlled trial; Veg, vegetarian

 Table 2. Study designs and participant characteristics of clinical trials included in the meta-analysis.

Author	Country	Year	Design	Duration (wks)	No.	Age (yrs)	Men (%)	SBP (mmHg)	DBP (mmHg)	Rx (%)	BMI (kg/m²)	Alcohol intake	Interventio n	Reference diet	Food preparation
Rouse (16)	AUS	1983	RCT, O, C	6	38	40.1	50.0	127.7	76.4	0	23.7	Participants asked to not alter alcohol consumption	Lacto-ovo vegetarian	Habitual diet	Yes (2/d)
Margetts (17)	AUS	1985	RCT, O, P	12	39	49.9	71.8	155.4	99.9	0	27.6	Participants asked to not alter alcohol consumption	Lacto-ovo vegetarian	Habitual diet	Yes (meat substitutes)
Fehily (18)	UK	1986	RCT, O, C	8	201	36.0	73.1	132.1	79.8	1.5	NR	HF/LF, 11.5g/13.6g per day	high fiber	Low fiber	No
Kestin (19)	AUS	1989	RCT, O, C	6	17	44.0	100.0	128.0	79.0	0	25.5	Veg/Cont, 4.2%/4.8% energy	Lacto-ovo vegetarian	Average Australian diet	Yes (major sources of protein and fat)
Hakala (20)	Finland	1989	RCT, O, P	52	73	38.0	24.7	129.9	85.0	0	34.4	2% of energy intake for Int. & Ctrl.	Lacto vegetarian	Habitual diet	No
Little (21)	UK	1990	CT, O, P	8	81	57.6	49.4	139.9	78.0	NR	NR	4.9 units/week	DASH	Habitual diet	No
Sciarrone (22)	AUS	1993	RCT, O, P	6	20	41.0	100.0	134.2	77.2	0	25.3	Individuals using >20 g of ethanol/d excluded	Lacto-ovo vegetarian	Average Australian diet	Yes
Appel (23)	USA	1997	RCT, O, P	11	305	44.3	50.4	131.8	85.1	94.8	28.1	Participants excluded if they consumed >14 drinks/week	DASH	Standard Western Diet	Yes
Nicholson (24)	USA	1999	RCT, O, P	12	11	54.3	45.5	141.3	84.7	81.8	NR	Individuals using alcohol regularly were excluded	Vegan diet	Low-fat	All lunches and dinners
Broekmans (25)	NL	2001	RCT, O, P	4	47	49.3	51.1	127.5	80.5	NR	25.8	NR	High F&V	Low F&V	Yes
Esposito (26)	Italy	2004	RCT, SB, P	104	180	43.9	61.1	135.0	85.5	NR	28.0	participants with active	MED	Prudent	No

												alcohol abuse excluded			
Azadbakht (27)	Iran	2005	RCT, O, P	24	76	41.4	29.0	143.6	85.7	0	29.9	NR	DASH	Weight reducing diet	No
Nowson (28)	AUS	2005	RCT, O, P	12	54	48.0	100.0	135.0	88.4	33.3	30.4	DASH/LF, 12 g/17.6 g per day	DASH	Low-fat	Dairy product of choice was provided once
Barnard (29)	USA	2009	RCT, O, P	74	98	55.6	39.4	123.3	78.0	69.7	34.9	Participants with active alcohol abuse excluded	Vegan diet	ADA diet	No
Nowson (30)	AUS	2008	RCT, O, P	14	95	59.2	0.0	127.6	81.0	36.8	29.6	Participants were excluded if they consumed >30 standard drinks/week	DASH + lean beef	Conventional advice	Some foods
Rallidis (31)	Greece	2009	RCT, O, P	8	82	50.4	52.4	129.9	86.1	NR	32.2	Participants were excluded if they consumed >500 g/week	MED	Conventional advice	Some foods
Ferdowsian (32)	USA	2010	CT, O, P	22	113	44.4	17.7	117.8	79.7	NR	NR	Participants with active alcohol abuse excluded	Vegan diet	Habitual diet	Yes (2/d)
Blumethal (33)	USA	2010	RCT, O, P	16	94	51.8	33.7	137.8	85.8	0	32.9	NR	DASH	Habitual diet	No
Adamsson (34)	Sweden	2011	RCT, O, P	6	86	53.0	59.3	128.8	82.1	NR	26.4	Nord/Cont, 1.7%/2.1% energy	healthy Nordic	Habitual diet	Yes
Azadbakht (35)	Iran	2011	RCT, O, C	8	31	NR	58.0	136.0	81.9	NR	NR	NR	DASH	Average Iranian diet	No
Morenga (36)	NZ	2011	RCT, O, P	8	74	41.9	0.0	125.0	80.0	NR	34.0	5.48 g/day	High fibre	High protein	No
Brooking (37)	NZ	2012	RCT, O, P	24	41	41.4	30.4	126.3	79.7	8.9	34.8	NR	High fibre	Habitual diet	Yes (food basket at start)
Toledo (38)	Spain	2013	RCT, O, P	208	4717	66.9	43.2	149.0	82.5	70.3	29.9	Participants with active alcohol abuse excluded	MED + nuts	Low-fat	Yes (30g of nuts/d)
Mishra (39)	USA	2013	RCT, O, P	18	215	45.2	17.2	127.0	81.9	NR	33.8	Participants with active	Vegan diet	Habitual diet	No

												alcohol abuse excluded			
Uusitupa (40)	Nordic countries	2013	RCT, O, P	18	189	54.4	33.3	130.0	82.0	51.9	31.6	participants were excluded if they consumed >40 g/day	Healthy Nordic	Standard Nordic Diet	Yes (key food groups every 1-2 weeks)
Poulsen (41)	Denmark	2014	RCT, O, P	26	145	42.1	29.3	122.5	81.3	NR	30.3	NND/ADD, 0.02%/0.00% energy	Healthy Nordic	Average Danish Diet	Yes
Macknin (42)	USA	2015	RCT, O, P	4	28	46.3	32.1	123.4	78.8	NR	35.2	NR	Vegan diet	АНА	No
Lee (43)	AUS	2015	RCT, O, C	1.4	24	25.6	0.0	113.2	75.3	NR	23.0	participants with active alcohol abuse excluded	MED	Habitual diet	No
Wong (44)	China	2015	RCT, O, P	52	405	55.1	49.0	145.0	90.2	0	24.2	87.1% non- drinkers, 12.9% current/ex- drinkers	DASH	Conventional	No
Bunner (45)	USA	2015	RCT, O, P	20	34	57.0	44.1	141.9	84.4	NR	36.0	consumption is more than two drinks per day	Vegan diet	Habitual diet	No
Lee (46)	S Korea	2016	RCT, O, P	12	93	57.9	19.4	126.6	76.9	43.0	23.5	NR	Vegan diet	Korean Diabetes Association	No
Noad (47)	UK	2016	RCT, SB, P	8	93	54.8	53.8	141.2	85.0	78.5	30.7	Men excluded if consumed >28 units/week and women if >21 units/week	High F&V	Low F&V	F&V
Davis (48)	AUS	2017	RCT, O, P	24	136	71.0	43.6	124.2	71.0	NR	26.9	NR	Med diet	Habitual diet	Some foods
Wright (49)	NZ	2017	RCT, O, P	12	65	56.0	40.0	132.5	79.5	NR	34.4	Participants with active alcohol abuse excluded	Vegan diet	Conventional	No
Barnard (50)	USA	2018	RCT, O, P	20	22	61.0	46.7	129.5	77.9	NR	33.9	Excluded if alcohol consumption >2 drinks/day	Vegan diet	Portion- controlled diet	No
Kucharska (51)	Poland	2018	RCT, O, P	12	126	59.8	50.8	130.5	84.2	100. 0	32.8	<2 drinks/d = 31%, >2 drinks/d = 4%	DASH	Conventional	No
Lee (52)	Korea	2018	RCT, O, P	8	58	43.2	70.7	134.9	86.4	NR	25.2	Participants were excluded	DASH	Conventional	No

												if they consumed >14 servings/week			
Wade (53)	AUS	2018	RCT, O, C	24	41	60.2	31.7	129.5	87.8	0	30.8	MD/LF, 4.58%/5.65% energy	MED + dairy	Low-fat	Some foods
Hashemi (54)	Iran	2019	RCT, O, P	12	75	NR	38.7	130.0	87.3	NR	NR	Participants excluded if they consumed alcohol	DASH	ADA diet	No
Wade (55)	AUS	2019	RCT, O, C	8	31	61.0	30.3	128.9	76.1	0	30.6	MD/LF, 4.41%/3.54% energy	MED + lean pork	Low-fat	Some foods
Mayr (56)	AUS	2019	RCT, O, P	24	65	61.8	83.1	136.8	82.1	NR	29.9	NR	MED	Low-fat	No

Abbreviations: BP, blood pressure; BMI, body mass index (weight in kilograms divided by height in meters squared); C, crossover; Cont, control; CT, controlled trial; F&V, fruit and vegetables; HF, high fiber; LF, low-fat; MED, Mediterranean; NR, not reported; O, open label; P, parallel; US, United states; RCT, randomised controlled trial; Veg, vegetarian

 Table 3. GRADE summary of findings.

Outcomes	Effect (95% CI)	No of participants (No of studies)	Certainty of the evidence (GRADE)	Comments
Change in Systolic Blood Pressure	SMD <b>4.47 lower</b>	420	$\oplus \oplus \oplus \bigcirc$	
(mmHg) - Healthy Nordic	(7.14 lower to 1.81 lower)	(3)	MODERATE a,b,c,d	
Change in Systolic Blood Pressure	SMD <b>0.57 lower</b>	140	$\oplus\bigcirc\bigcirc\bigcirc$	
(mmHg) - High Fruit & Vegetables	(7.45 lower to 6.32 higher)	(2)	VERY LOW a,e,f	
Change in Systolic Blood Pressure	SMD <b>0.65 lower</b>	316	ФООО	
(mmHg) - High Fiber	(1.83 lower to 0.53 higher)	(3)	VERY LOW a,b,c,g	
Change in Systolic Blood Pressure	SMD <b>5.47 lower</b>	187	$\oplus \oplus \oplus \oplus$	
(mmHg) - Lacto (and ovo) vegetarian	(7.6 lower to 3.34 lower)	(5)	HIGH <sup>a,b,c,h</sup>	
Change in Systolic Blood Pressure	SMD <b>5.53 lower</b>	1400	$\oplus \oplus \oplus \oplus$	One study was not
(mmHg) - DASH	(7.95 lower to 3.12 lower)	(11)	HIGH <sup>a,b</sup>	randomised
Change in Systolic Blood Pressure	SMD <b>0.95 lower</b>	5276	⊕⊕⊕○	
(mmHg) - Mediterranean	(1.7 lower to 0.2 lower)	(8)	MODERATE a,b,i	
Change in Systolic Blood Pressure	SMD 1.30 lower	677	$\oplus \oplus \bigcirc \bigcirc$	One study was not
(mmHg) - Vegan	(3.90 lower to 1.29 higher)	(9)	LOW <sup>a,d,j</sup>	randomised
Change in Diastolic Blood Pressure	SMD <b>2.32 lower</b>	420	$\oplus \oplus \oplus \bigcirc$	
(mmHg) - Healthy Nordic	(3.83 lower to 0.82 lower)	(3)	MODERATE a,b,c,d	
Change in Diastolic Blood Pressure	SMD <b>0.96 lower</b>	140	ФООО	
(mmHg) - High Fruit & Vegetables	(3.08 lower to 1.15 higher)	(2)	VERY LOW a,e,f	
Change in Diastolic Blood Pressure	SMD <b>1.02 lower</b>	316	$\oplus\bigcirc\bigcirc\bigcirc$	
(mmHg) - High Fiber	(3.86 lower to 1.82 higher)	(3)	VERY LOW a,b,c,g	
Change in Diastolic Blood Pressure	SMD <b>2.49 lower</b>	187	$\oplus \oplus \oplus \bigcirc$	
(mmHg) - Lacto (and ovo) Vegetarian	(4.17 lower to 0.8 lower)	(5)	MODERATE a,b,c,h	
Change in Diastolic Blood Pressure	SMD <b>3.78 lower</b>	1400	$\oplus \oplus \oplus \oplus$	One study was not
(mmHg) - DASH	(5.51 lower to 2.04 lower)	(11)	HIGH <sup>a,b</sup>	randomised
Change in Diastolic Blood Pressure	SMD <b>0.69 lower</b>	5276	$\oplus \oplus \oplus \bigcirc$	
(mmHg) - Mediterranean	(1.44 lower to 0.06 higher)	(8)	MODERATE a,b,i	

Change in Diastolic Blood Pressure	SMD 0.81 lower	677	$\Theta\Theta\bigcirc\bigcirc$	One study was not
(mmHg) - Vegan	(2.91 lower to 1.28 higher)	(9)	LOW a,d,j	randomised

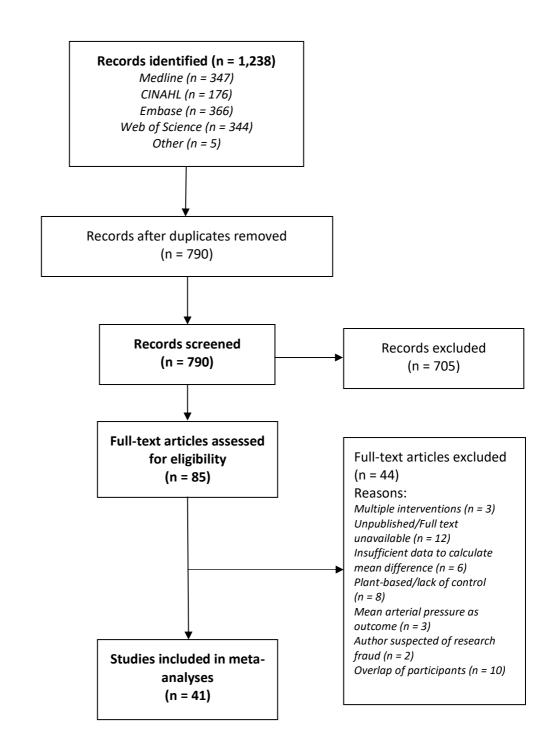
# **Explanations**

- a. Participants were not blinded in all studies
- b. Experimental personnel were not blinded in some studies
- c. Outcome assessor was not blinded in some studies
- d. Selective reporting. The pre-registered protocol did not list BP as an outcome for some studies.
- e. Only two studies, each showing opposite results
- f. Small sample size led to large 95% CIs
- g. Morenga et al found a significantly larger effect compared with the other studies.
- h. Large no. of dropouts and no ITT analysis for some studies
- i. Two of the studies were industry funded
- j. Wide 95% CI that overlaps with no effect

**Table 4.** The effects of various plant-based diets on systolic and diastolic blood pressure when compared to a standardised control diet.

Diet	Studies	Sample	e size	Systolic BP	95% C.I.	Diastolic BP	95% CI
	(n)	Intervention	Control	difference		difference	
				(mmHg)		(mmHg)	
Healthy Nordic Diet	3	234	186	-4.47	-7.14, -1.81	-2.32	-3.83, -0.82
High Fruit & Vegetables	2	70	70	-0.57	-7.45, 6.32	-0.96	-3.08, 1.15
High Fiber	2	59	56	-1.69	-4.61, 1.24	-1.85	-6.15, 2.45
Lacto (and ovo) vegetarian	5	116	71	-5.47	-7.60, -3.34	-2.49	-4.17, -0.80
D.A.S.H.	4	269	242	-8.74	-12.20, -5.28	-6.05	-9.60 <i>,</i> -2.50
Mediterranean	2	94	66	-1.15	-2.04, -0.26	0.29	-1.43, 2.01
Vegan	3	181	181	-2.73	-8.29, 2.83	-2.48	-6.91, 1.94
Pooled	21	1023	872	-4.29	-6.27, -2.31	-2.79	-4.33, -1.24

Results are expressed as mean difference (95% confidence intervals).



# 7. Figure 2\_04062020\_R1.png

	ran Difference	SE.	Total		WHITE	My Kondern, 95% CT Yea	Still Mean Difference N, Random, 1955 CI	ARCDEF
2.1.1 Healthy Medic	7.15	2.64821	344	1.00	24.04	F5514236-100: 201		
Adamsson et al. (2011) Unishipe et al. (2013)	-3	1.0676	89	90	21.0%	-7:5511234,-1:00 201 -2:0035.70,170 201		2202024
Poulsen et al. (2014):	-513	1.5410	95	54	44.4%	-5.131-0.16, -2.10) 201		
Subtrinial (95% Ct)			234	186	100.0%	4.47 [ 7.14, -1.81]	•	The state of the state of the
Heterogeneity $Tau^a = 1.78$ , $Chl^a = 2.9$ Text for overall effect $Z = 3.39$ ( $P = 0.1$		13), P = 31 %					500	
2.1.2 High Fruit & Vegetables	27,100							
Promission at at (2001)	2.8	2.7552	24	23	62.1%	2,8012,60,820,200	-	
issad et al. (2018)	4.23	3.11288	40	47	47.9%	-4.23 (10.33, 1.87) 201		*****
Substituted (95% CD)			70	70	100.0%	0.571-7.45, 6.321		
Helanogenedy Tauf × 16.07, Chif × 2. Test for overall effect Z × 6.16.07 × 0.1		38,7=45%						
2.1.3 High Fiber	W.E.							
Februaret al. (1988)	-2.4	2.6884	201	6	88.3%	-0.405-1.71, 0.912, 198		2200074
Monenga et al. (2011)	-3.1	3.8878	37	37	10.1%	3.1934.60,080) 381		
Brooking et al. (2012)	-011	2.0153	12	19	0.0%	-0.11 (-4.00, 3.94) 201	1	2-2-0-2-0-0
Subtotal (95% CS) Hotorogenede Tauf = 0.00; CnF = 1.9	N. d+2F+53	791, P = 0%	200	-56	100.0%	0.60 [ 1.83, 6.53]	•	
Teat for prevail effect Z = 1.06 (P = 0.)	20							
2, 6,4 Eacle (and evo) vegetation Rouse et al. (1993)	40	1.4286	36.7		17.0%	-0.0010.000.00.100		******
Margartts at at (5 tild)	-5.0	2.416	20	16	10.1%	-6.80 (-12.5); 0.90; 198	The second secon	2202020
Kingtim et al. (1985)	-3	3,1123	17	0	12.2%	-3.001512,310,198		
Hakata et al. (1989)	-3.3	2.5511	31	42	18.2%	3.30 ( 6.30, 1.70) 188		
Sciemone ed al. (1993)	1.5	9.5716	10	10	1.0%	1.50 (-15.30, 18.30), 189		2200020
Subtrical (95% CI)			110	71	100.0%	3.471.7.60, 3.34	•	
Haterogenedy. Tau* = 2.00; Clu* = 2.0 Two: for overall effect. Z = 1.03 (F = 0.1		Mr. P = 0%.					54.51	
2.5.5 DASH								
Little et al. (1998)		3.01057450	41	43	5.4%	(13.70) 21.16, (6.22), 189		000001
Appel et al. (1997)	-5.5	0.3484	151	104	10.7%	-5.50 ) 7.40, -3.80) 199		77.0000
Advisorable et al. (2005) (1)		2.53779476	37	37	11.4	-9.00 ( 17.95, -3.05) 200		
Azadtuskot et al. (2005) (2)	-5	1.02000	77	71	1.0%	-5.00   20.53   10.50   200 -5.50   0.25   1.74   200		
Nowson et al. (2005) Nowson et al. (2005)	-55	1.62052069	46	49	9.5%	-2.90   6.04   0.29   200		1100010
Blumenthal et al. (2010)	-7.8	2.1092537	40	43	8.4%	-7.80 §1211,-3.49 301		*****
Apparitment of all (2011)		0.79392957	31	0	10.9%	-18 50 j 12 86, -8 54, 201		****
Wing et at. (2015)	-0.1	3.1736	294	201	10.2%	-0.10 (-2.40, 2.20) 201		
Hushanika et el. (2018)	-3.79	1.02123	-84	62	10.0%	-3.791-5.79, -1.791 201		220002
Lee et al. (2018)	-1.8	1.9684	30	26	8.0%	-1.001-576, 1.002 221		2202
Hasheni et al. (2019) (3) Hasheni et al. (2019) (6)		5.32618742 4.89678852	13	16	4.1%	-3.00[13:24,7:24] 301 -7.00[16.60,2.60] 301		
Subtremat (199% Ct)		* 1000110004	713		100.0%	5.531 7.95, 3.121	•	
Heteropeneitr Tau* = 13:29; Chi* = 7: Test for overall effect Z = 4.40 (F = 0.)		S 000011, F = 94%				HARROW BOLLOW	2.50	
2.5.6 Meditorraments								
Esposito et al. (2004)	-3	1.0204	95	90	10.0%	-3.00 ( 6.00, -1.00) 200		
Rathitis et al. (2009)	3.8	3 13736	41	41	1.5%	-3.96510.05, 2.20, 200		000707
Turedo et al. (2012)	-0.9	0.4439	2367	2360	27.1%	-0.00 ) 1.77, -0.03 301		
Lee et al. (2015)	7.00	2.6570000	24	0	2.0%	-2.06 ( 0.07, 2.36) 201		220002
Deep et al. (2017):	-1.1	0.4582	70	.06	25.6%	-1.101-2.00,-0.301 301	•	*****
Wade et al. (2018)	-6.2	0.4733	- 41	0	29.5%	-0.2011-01, 0.011-201		
May et al. (2518)	-0.3	4.14057	34	-31	0.0%	-0.3018.43,783 201		110101
Waste et al. (2019) Subbidue (95% Cb)	2.53	2.1220	3688	2524	100.0%	2.53(1.53, 6.60) 201 6.95(1.176, 6.29)		*****
Heteropenety: Tau*+ II 35; Cts*+ 11.		131.7+38%	,,,,,,			most con most	1	
Tent for evenual effect Z = 2.47 (F = 0.1	91)							
2.1.7 Veges	5.5	0.0104	7		7.66	# 865 to 30 to 30 to 300		
hishessis et al (1996) Hamant et al. (2000)	-2.7	2.8963	40	40	15.0%	5.88§10.20,21.20; 199 -3.70§9.20; 1.80; 200		
Ferthrecian et al. (2010)	-6.7	2.4745	44	45	17.7%	-5.70310.55, -0.85, 201		000100
Missing of al. (2012)	0.8	1 2756	90	119	32.6%	8.80 ( 1.72, 3.30) 201		777799
Machinin et al. (2016)	4.02	5.96200694	14	14	6.0%	482 (1878, 510) 201		
Buriner et al (2011)	17.2	0.8369	17	37	25%	-7.20   20.60, 6.30] 201		*****
Les et al. (2016)		3.51007204	46	47	18.9%	2501438,638,221		22
Winger of al. (2017) Damard et al. (2018)	5.5	4.5919 5.8634	33	32	1.0%	-2.00   11.00, 7.00  201 5.50   5.63, 10.00  201		
Subtotal (95% CB)			338		100.00	-1.30 (-3.96, 1.29)	•	
reterogenedy Tau* = 3.79; Cls* = 10.	98, at = 1 at = 0	211.7+26%	1		-		0.57	
Tent for evental effect $Z \approx 0.56 \ (P \approx 0.5)$								
							3a -la a ta	26
							Falcura experimental diel: Favoura control d	SAE.
Eurottophek (1) Ferrosin							filist at bias learns At Pandom sequence perendon (selecti	ne board
C23 MANA							(B) Alteration concentrator (selection bias)	
(3) Male							(C) Strateg of participants and personnel	
(4) Ferroam							(B) Elimping of outcome as a expression of other	
Position of the second							#1 incomplete outcome data catrillon tiral.	
							#) Selective reporting (reporting (ratio)	
							50 Funding tras.	

# 8. Figure 3\_04062020\_R1.png

