

Health Benefits of a Vegan Diet: Current Insights

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Abstract: To assess the health benefits of a vegan diet, observational studies of diabetes (non-insulin dependent; NIDDM), cancer, mortality, gut microbiota, hypertension, lipids, and overweight were examined. Utilizing PRISMA items to identify articles and assess their quality for inclusion, 44 studies were screened into the review. In two separate studies, vegans did not have a reduced risk of diagnosis of “any” cancer, nor of specific cancers when the effects of BMI were adjusted. When data were aggregated, there was a reduced risk of any cancer. Vegans had lower total cholesterol and LDL-C levels compared to omnivores, and in some cases other dietary groups. In the US, there was a reduced risk of a diagnosis of NIDDM and in Western countries, vegans had lower BMI. Research support for reduced risk of diagnosis of female cancers, improved mortality rates, lower blood pressure, lower triglycerides, and a healthier microbiome for vegans compared to omnivores was inconsistent. There was no evidence that reduced specific cancer incidence rates were lower in vegans although inadequate sample sizes had hampered these analyses. In vegans, HDL-C levels were either lower or not significantly different from omnivores. Geographic location was a strong moderator with the most compelling evidence for the health advantages of a vegan diet coming from the US, whereas in Taiwan, India and Vietnam, there was much less evidence of its benefit. In some instances, sex moderated the association between adopting the diet and health outcomes. Adherence, specific content of the diet, and dietary comparison groups utilized in studies may also affect results. Although a vegan diet is associated with some health benefits, the level of support for the benefit varied according to the health outcome being measured, with evidence emerging that BMI is an important mediator and geography and to a lesser extent sex are important moderators.

Keywords: vegan, vegetarian, health

Introduction

In 1944, members of the Vegetarian Society in the United Kingdom first coined the term “vegan.”¹ Veganism is defined as “... the practice of dispensing with all products derived wholly or partly from animals.”² Hence, people identifying as vegan avoid eating meat, poultry, fish, eggs or dairy foods. Although veganism extends beyond food choices, as many adopted the diet, its health benefits began to be investigated.

A review of the health benefits of a vegan diet is timely. One previous review³ was published eleven years ago. Recently there have been more focused reviews and meta-analyses (eg, Yokoyama et al⁴), but a more expansive analysis has not been performed.

Because addressing the health benefits of a vegan diet is a broad topic, further delineation of the parameters of this review was necessary. First, we chose to focus on observational studies (cohort, case control and cross-sectional designs) rather than clinical trials. Second, we decided to forego reviewing the nutrient deficiencies that can result from a poorly planned vegan diet as these have been described elsewhere.⁵ Beyond addressing diet benefits, health conditions and medical/nutritional indicators of health problems were selected based on there being sufficient literature to merit inclusion.

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Furthermore, if findings from included studies showed those following a vegan diet had worse outcomes than those following other diets, these results were reported.

The specific areas targeted in this review are the following: non-insulin dependent diabetes mellitus (NIDDM), cancers, mortality, gut microbiome, hypertension (blood pressure), lipids (cholesterol), and overweight (body mass index (BMI)).

Methodology

In line with recommendations for reviewing studies within a broad topic area, we adopted a semi-systematic approach.⁶ We utilized methods outlined in the Preferred Reporting Items for Systematic Reviews and Meta-Analyses (PRISMA)^{7,8} to identify, evaluate and ultimately select studies for inclusion. Furthermore, we reported in [Table 1–7](#) characteristics of studies in the review including location, cohort, design, sample description, sex, mean age, results/follow-up, adjustment and effect size. However, as the topic areas were diverse (eg, cancer, NIDDM), we could not extract uniform data across topics. Additionally, we included well-conducted meta-analyses and other systematic analyses and reviews in our discussion of findings as this work is informative.

Eligibility criteria for inclusion were the following: 1) English language, 2) Human participants, 3) Focused on health benefits of a vegan diet (see above for specific areas), 4) Inclusion of both vegans and an appropriate comparison group, 5) Age of participants was 18 or older, 6) Vegan diet was defined as consuming fish, meat, poultry, dairy or eggs no more than once monthly, 7) On vegan diet at least 3 weeks, 8) Cohort, cross-sectional or case-controlled design, 9) Dependent variables were well-defined health outcomes, and 10) Full-text article (not just abstract) published in peer-reviewed journals. Exclusion criteria were 1) Vegans combined with another group for study 2) Other diet (eg, vegetarian), and 3) Lack of between-group parametric tests.

Search Plan and Identification of Studies

A systematic search of Pubmed, Cochrane Central Register of Controlled Trials, MEDLINE, and MEDLINE PLUS was performed on June 24, 2019. Then, reference lists of articles with relevant topic areas were reviewed to search for additional studies that were appropriate. In addition, articles published in the first issue of a new journal focused on studies of plant-based diets were considered for the review. The search terms used to locate studies were: “plant-based” OR “vegan” AND “mortality” OR “death” OR “risk assessment” OR “heart disease” OR “cardiovascular” OR “cardiovascular disease” OR

“stroke*” OR “cerebrovascular disease” OR “statin*” OR “cancer*” OR “tumor” OR “diabet*” OR “TRD” OR “DT2” OR “cholester*” OR “gut microbiome” OR “gut bacteria” OR “body mass index” OR “bmi” OR “bariatric surgery” OR “obesity” OR “overweight” OR “gastric banding” OR “hypertension” OR “blood pressure increase” OR “atrial fibrillation”.

Study Selection

Results from searches of the databases were combined yielding a total of 3,105 (see [Figure 1](#)) published articles. After removal of duplicates (N=207), the remaining 2,898 articles were screened to determine if they met inclusion criteria, using a systematic inspection of article title, abstract, and if necessary, full text. From this screening, an additional 2,826 articles were excluded based on abstract and title review. After full-text review, an additional 32 articles were excluded. Specifically, 26 studies were excluded for not measuring a target outcome, 1 study was excluded for using children as participants, 1 study was excluded for not being an original study, 2 studies were excluded for having no control group, and 2 were excluded because the sample was comprised of both vegetarians and vegans. A review of the reference sections of screened-in articles and other sources revealed additional 20 articles, yielding a total of 60 studies undergoing quality assessment.

No case-control studies were selected to undergo evaluation. To assess the methodological rigor of screened in studies, all were rated using the Newcastle-Ottawa Quality Assessment Scale (NOS)⁹ for either 1) cohort or 2) cross-sectional designs.¹⁰ Both scales are widely used in systematic reviews and have strong psychometric support. Each study was assessed independently by two raters, and any rating disagreement was settled by discussion and consensus. Ratings of “Fair” or “Good” from both raters were required for inclusion. From those assessments, 16 articles were removed for failing quality assessment (see [Supplementary Table](#)).

A total of 44 studies were screened into the review (See [Table 1–7](#)). As some studies reported multiple outcomes, they may appear in more than one table. Effect sizes were computed for studies that provided standard deviations or standard errors of the mean. To facilitate comparisons between studies in Tables, the following abbreviations were used: VG=vegan; VEG=vegetarian or ovo-lacto vegetarian; PESC=pescatarian, pisco-

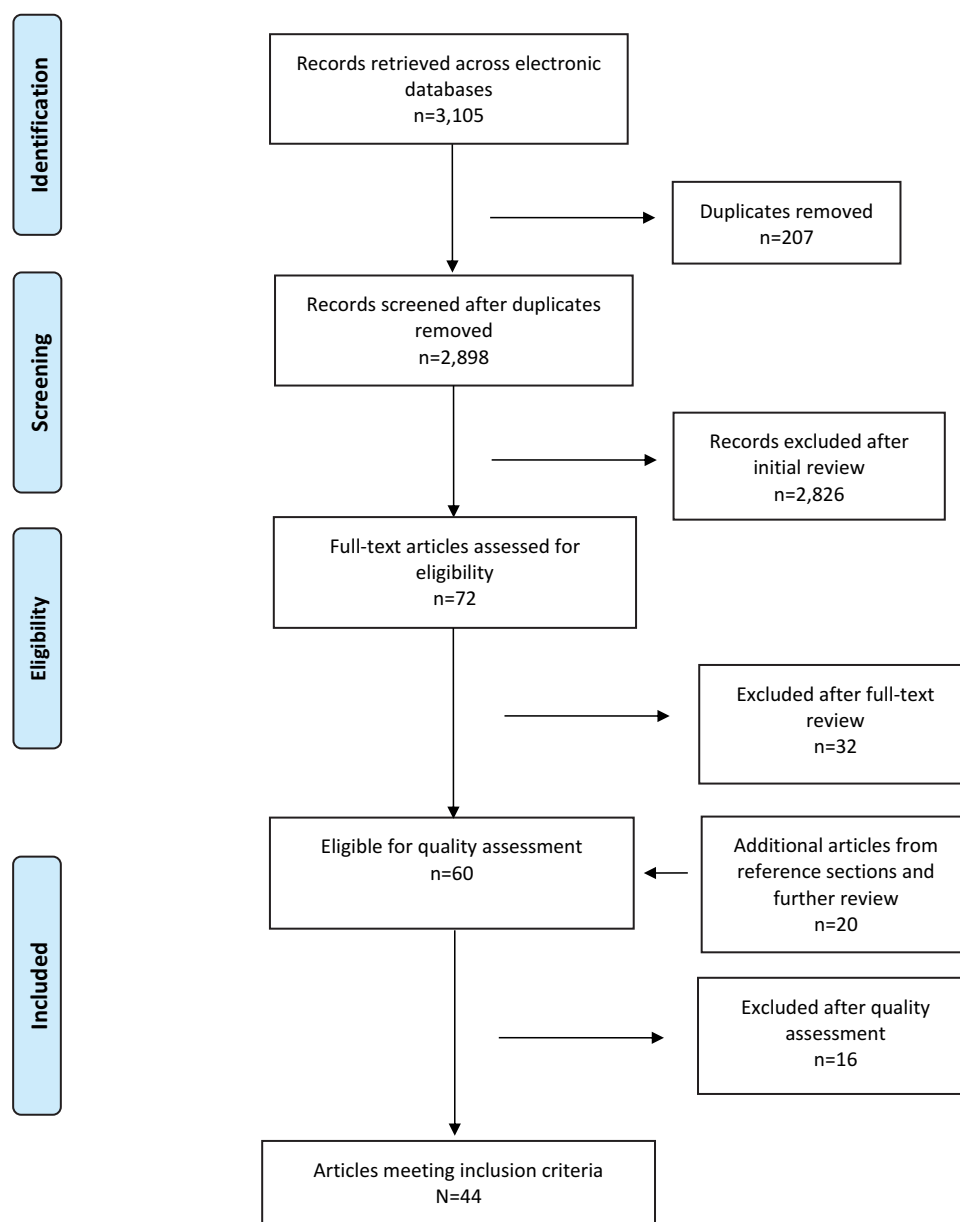


Figure 1 Search outcomes.

vegetarian or “fisheater;” OMN=omnivore, non-vegetarian or “meateater.”

Results

Diabetes. Studies investigating the relationship between diabetes and a vegan diet have focused on NIDDM. Four observational studies^{11–14} were published between 2009 and 2019 (see Table 1). Two US studies reported data from the same cohort of participants,^{11,12} with a total of 60,903 participants in the first study and 49,140 of those individuals also in the second study, conducted two years later. The cases of NIDDM and Insulin-Dependent Diabetes Mellitus (IDDM)

found during the first assessment were removed from the follow-up analyses. In these two studies, vegans were less likely to develop NIDDM than omnivores, but not other comparison groups, when controlling for BMI. In a UK study, compared to omnivores there was no significantly lower risk for incidence of NIDDM when BMI was controlled although there was a significantly lower risk of NIDDM in vegans when BMI was not controlled.¹³ However, an Indian study reported no significant reduction in risk for vegans and other dietary groups.¹⁴

Cancers. Three observational studies of cancer risk in dietary groups were included in the review (see Table 2).^{15–17} They

Table 1 NIDDM

Study (Author, Year)	Location	Cohort	Design	Sample Description	M/F	Age (Mean Years)	Results/Follow-Up	Adjustment	Effect Sizes
Tonstad et al (2009) ¹¹	North America	The Adventist Health Study-2	C	N=60,903 (VG=2,731, VEG=20,408, PESC=5,617, semi-VEG=3,386, OMN=28,761)	M/F	53	VG had a significantly reduced prevalence of NIDDM relative to OMN. There were no significant differences in prevalence of NIDDM between VG and other VEG and PESC groups.	Age, sex, BMI, ethnicity, education, income, physical activity, television watching, sleep habits, and alcohol use.	OR CI=0.40, 0.66 (NIDDM, VG ^a)
Tonstad et al (2013) ¹²	US and Canada	Adventist Health Study-2	C	N=41,387 (VG=3,566, VEG=14,096, PESC=3,635, semi-VEG=2,386, OMN=17,704)	M/F	60.1	In an originally non-diabetic group, after two years, cases of diabetes developed in 0.54% of VG, 1.08% of lacto-ovo VEG, 1.29% of PESC, 0.92% of semi-VEG and 2.12% of OMN. VG had a significantly lower diabetes incidence than OMN, but not other comparison groups.	Age, sex, BMI, education, income, television watching, physical activity, sleep, alcohol use, and smoking.	OR CI=0.24, 0.62 (diabetes, VG ^a)
Agrawal et al (2014) ¹⁴	India	Indian's Third National Family Health Survey	C	N=156,317 (VG=2,560, lacto-VEG=37,797, VEG=5,002, semi-VEG=8,140, PESC=3,446, OMN=99,372)	M/F		There was no significant difference in prevalence of diabetes between VG and other diet groups.	Age, sex, BMI, education, household wealth, rural/urban residence, religion, caste, smoking, alcohol use, television watching	OR CI=0.61, 1.36 (diabetes, VG ^a)
Papier et al (2019) ¹³	UK	EPIC-Oxford	L, 17.6 y F/U	N=45,314 (VG=1,781, VEG=13,645, PESC=7,092, semi-VEG=7,615, OMN=15,181)	M/F	44.5	No significant difference in diabetes incidence between VG and other groups.	Age, Sex, BMI, recruitment method, region of residence, education, Townsend deprivation index, ethnicity, smoking, alcohol intake, physical activity	HR CI=0.66, 1.48 (diabetes, VG ^a)

Notes: ^aWith OMN as reference group

Abbreviations: NIDDM, Non-insulin Dependent Diabetes Mellitus; VG, vegan; VEG, vegetarian or ovo-lacto vegetarian; PESC, pescatarian, pesco-vegetarian or "fish-eater"; OMN, omnivore, non-vegetarian or "meateater"; UK, United Kingdom; US, United States; N, number; C, cross-sectional; L, longitudinal; y F/U, year follow-up; M, males; F, females; BMI, body mass index; CI, confidence interval; OR, odds ratio.

Table 2 Cancer

Study (Author, Year)	Location	Cohort	Design	Sample	M:F	Age (Mean Years)	Results/Follow-Up	Adjustment	Effect Sizes
Tantamango-Bartley et al (2012) ¹⁵	US	Adventist Health Study-2	C	N=69,120 (VG=4,922, VEG=19,735, semi-VEG=3,881, PESC=6,846, OMN=33,736)	M/F		There was no significant difference in cancer incidence between VG and OMN.	Race, family history of cancer, BMI, education, smoking, alcohol, age at menarche, pregnancies, breastfeeding, use of oral contraceptives, hormone replacement therapy, menopause status	HR CI=0.73, 1.00 (overall cancer, VG ^a); HR CI=0.55, 1.17 (GIT, VG ^a); HR CI=0.28, 1.23 (RT, VG ^a); HR IC=1.05, 2.84 (UT, VG ^a); HR CI=0.57, 1.17 (MC, VG ^a); HR CI=0.50, 1.01 (FC, VG ^a)
Key et al (2014) ¹⁶	UK	Oxford Vegetarian Study, EPIC-Oxford cohort	L, 14.9 y F/U	N=61,647 (VG=2,246, VEG=18,298, PESC=8,612, OMN=32,491)	M/F	43.98	After an average follow-up of 14.9 y, there was no significant incidence reduction found for colorectum, female breast, prostate, and all cancer in VG compared to OMN.	Sex, smoking, alcohol consumption, physical activity, parity and oral contraceptive use, method of recruitment, BMI	PR CI=0.82, 2.11 (colorectum, VG ^a); PR CI=0.61, 1.34 (female breast, VG ^a); PR CI=0.31, 1.20 (prostate, VG ^a); PR CI=0.68, 1.00 (all sites, VG ^a)
Penniecook-Sawyers et al (2016) ¹⁷	US	Adventist Health Study-2	L, 7.8 y F/U	N=50,404 (VG=3,748, VEG=14,336, PESC=5,179, semi-VEG=2,930, OMN=24,211)	F only		VG showed consistently lower (but non-significant) breast cancer incidence when compared with OMN.	Age, race, BMI	HR CI=0.62, 1.13 (breast cancer, VG ^a)

Notes: ^aWith OMN as reference group.

Abbreviations: VG, vegan; VEG, vegetarian or ovo-lacto vegetarian; PESC, pescatarian, pesco-vegetarian or "fish-eater"; OMN, omnivore, non-vegetarian or "meateater"; UK, United Kingdom; US, United States; N, number; C, cross-sectional; L, longitudinal; y F/U, year follow-up; M, males; F, females; BMI, body mass index; CI, confidence interval; HR, hazard ratio; GIT, cancer of the gastrointestinal tract; RT, cancer of the respiratory tract and intrathoracic organs; UT, cancer of the urinary tract; MC, all male cancer; FC, all female cancer; PR, prostate cancer.

Table 3 Mortality

Study (Author, Year)	Location	Cohort	Design	Sample Description	M:F	Age (Mean Years)	Results/Follow-Up	Adjustment	Effect Sizes
Key et al (1999) ²⁰	United States, United Kingdom, Germany	Adventist Mortality Study, Health Food Shoppers Study, Heidelberg Study, Oxford VEG Study	L, 10.6 y F/U	N=66,294 (VG=753, VEG=23,265, PESC=2,375, Low meat OMN=8,135, Reg meat OMN=31,766)	M/F		No significant difference in death rate ratios of stomach cancer, colorectal cancer, lung cancer, breast cancer, prostate cancer, ischemic heart disease, cerebrovascular disease, other causes, and all causes between VG and other diet groups.	Age, sex, smoking status, study	DRR CI=0.43, 11.2 (stomach, VG ^a); DRR CI=0.11, 6.17 (colorectal, VG ^a); DRR CI=0.39, 20.0 (lung, VG ^a); DRR CI=0.46, 1.21 (IHD, VG ^a); DRR CI=0.25, 1.98 (CVD, VG ^a); DRR CI=0.92, 1.93 (other causes, VG ^a); DRR CI=0.70, 1.44 (all causes, VG ^a)
Orlich et al (2013) ¹⁹	United States, Canada	Adventist Health Study-2	L, 5.79 y F/U	N=73,309 (VG=5,548, VEG=21,177, PESC=7,194, semi-VEG=4,031, OMN=35,359)	M/F	56.9	For both genders combined, VGs had lower other mortality than OMNs, this difference was not significant for either gender separately. No significant difference in HR of death from all causes, ischemic heart disease, cardiovascular disease, and cancer between VG and other diet groups, for women and all participants. VG males had similar death rates of all causes, ischemic heart disease, and cardiovascular disease compared to VEG males, PESC males, and SEMIVEG males, but lower death rates compared to OMN males. No significant difference in HR of death from cancer and other causes between VG males and males in other diet groups.	Age, sex, race, smoking, exercise, personal income, educational level, marital status, alcohol intake, region, sleep, menopause in women, hormone therapy in postmenopausal women	HR CI=0.73, 1.01 (all-cause, VG ^a); HR CI=0.60, 1.33 (IHD, VG ^a); HR CI=0.71, 1.16 (CVD, VG ^a); HR CI=0.68, 1.24 (cancer, VG ^a); HR CI=0.56, 0.99 (other-cause, VG ^a); HR CI=0.56, 0.92 (all-cause, VG ^a , M); HR CI=0.21, 0.94 (IHD, VG ^a , M); HR CI=0.38, 0.89 (CVD, VG ^a , M); HR CI=0.48, 1.36 (cancer, VG ^a , M); HR CI=0.53, 1.22 (other-cause, VG ^a , M); HR CI=0.78, 1.20 (all-cause, VG ^a , F); HR CI=0.87, 2.24 (IHD, VG ^a , F); HR CI=0.88, 1.60 (CVD, VG ^a , F); HR CI=0.69, 1.44 (cancer, VG ^a , F); HR CI=0.47, 1.05 (other-cause, VG ^a , F);

Appleby et al, (2016) ¹⁸	United Kingdom	Oxford Veg Study Cohort, EPIC-Oxford	L, 5-, 10-, 15-y F/U	N=60,310 (VG=2,228, VEG=18,096, PESC=8,516, Low meat OMN=13,039, Regular meat OMN=18,431)	M/F	43.4	No significant difference in HRs of malignant cancer, ischemic heart disease, cerebrovascular disease, disease of the respiratory, and all causes between VG and other diet groups.	Sex, BMI, smoking, alcohol consumption, physical activity, marriage status, nutritional supplements, study method, parity, oral contraceptive use, hormone therapy use, prior blood pressure, prior high blood pressure, medical treatment, use of separate models	HR CI=0.85, 1.42 (cancer, VG ^a); HR CI=0.88, 1.66 (circulatory disease, VG ^a); HR CI=0.53, 1.55 (IHD, VG ^a); HR CI=0.97, 2.69 (CVD, VG ^a); HR CI=0.86, 2.56 (Respiratory, VG ^a); HR CI=0.94, 1.30 (all-cause, VG ^a)
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Notes: ^aWith OMN as reference group.

Abbreviations: VG, vegan; VEG, vegetarian or ovo-lacto vegetarian; PESC, pescatarian, pesco-vegetarian or "fish-eater"; OMN, omnivore, non-vegetarian or "meateater"; US, United States; N, number; C, cross-sectional; L, longitudinal; y F/U, year follow-up; M, males; F, females; BMI, body mass index; CI, confidence interval; OR, odds ratio; DRR, death rate ratio; IHD, ischemic heart disease; CVD, cardiovascular disease; HR, hazard ratio.

were published between 2012 and 2016 and included 7,198 vegans, 38,028 vegetarians, 15,446 pescatarians, 3,885 semi-vegetarians and 66,204 omnivores for a total of 130,761 participants. Tantamango-Bartley et al¹⁵ and Penniecook-Sawyers et al¹⁷ reported different data from a cohort of participants where there was substantial overlap with the 44,674 female participants in the Tantamango-Bartley et al¹⁵ study included in the Penniecook-Sawyers et al¹⁷ study, although the latter study included data from more US states (48 vs 38 in the Tantamango-Bartley et al study).

Two cohort studies examined whether overall cancer diagnosis rates were related to the type of diet that individuals consume.^{15,16} According to Tantamango-Bartley et al,¹⁵ once several factors were controlled for including but not limited to BMI, education, smoking, and menopause, there was no significant difference in cancer incidence between vegans and omnivores in both genders combined and for female-specific cancers. In a similar vein, Key et al¹⁶ reported a 14% lower risk of any cancer diagnosis in vegans compared to omnivores, but it was not a statistically significant reduction. However, when BMI was not a covariate, there was a specific risk reduction for "any" cancer and for female cancers. There was no significant risk reduction in the incidence of other specific diseases (colorectal, breast, or prostate cancer). Confirming the Key et al¹⁶ finding for breast cancer incidence, Penniecook-Sawyers et al¹⁷ also found a lower risk of breast cancer in vegans that was not statistically significant.

Mortality. The 3 articles reporting mortality rates in dietary groups included several cohort studies (see Table 3).^{18–20} They were published between 1999 and 2016. One report included data from the EPIC-Oxford Study,¹⁸ one included data from the Adventist Health Study-2,¹⁹ and one included data from four cohorts (Adventist Mortality Study, Health Food Shoppers Study, Heidelberg Study Cohort, Oxford Vegetarian Study Cohort).²⁰ The total sample size was 199,913 observations (mean age=50.81), including 8,529 vegans, 53,790 omnivores, 18,085 pescatarians, 25,205 semi-vegetarians (including 21,174 low meat omnivores), and 62,538 vegetarians. Participants from the Oxford Vegetarian Study (11,047) were included in both Key et al²⁰ and Appleby et al.¹⁸

In the majority of comparisons between vegans and other dietary groups, no significant differences in mortality risk were found. In a UK study,¹⁸ and one combining data from five studies,²⁰ the death rate hazard ratio (HR) for vegans compared to omnivores was not significantly different for malignant cancer (stomach cancer, colorectal

Table 4 Gut Microbiome

Study (Author, Year)	Location	Cohort	Design	Sample Description	M: F	Age (Mean Years)	Results/Follow-Up	Adjustment	Effect Sizes
Zimmer et al (2012) ²⁵	Germany		C	N=498 (VG=105, VEG=144, OMN=249)	M/ F	53.04	Total counts of Bacteroides spp., Bifidobacterium spp., Escherichia coli and Enterobacteriaceae spp. were significantly lower in VG samples than in controls, whereas others (E. coli biovars, Klebsiella spp., Enterobacter spp., other Enterobacteriaceae, Enterococcus spp., Lactobacillus spp., Citrobacter spp. and Clostridium spp.) were not. Subjects on a VG or VEG diet showed significantly lower stool pH than did controls, and stool pH and counts of E. coli and Enterobacteriaceae were significantly correlated across all subgroups.	Age, sex	
Ferrocino et al (2015) ²⁴	Italy		C	N=153 (VG=51, VEG=51, OMN= 51)	M/ F	38	The fecal microbiota of the VEG and VG volunteers showed significantly lower microbial counts of the Bacteroides fragilis group. The mesophilic/thermophilic LAB loads (on MRS agar) were also low in the VG and VEG groups.		
Ruengsomwong et al (2016) ²⁷	Thailand		C	N=72 (VG=3 ovo-lacto-VEG=4, lacto-VEG=28 ovo-VEG=1, OMN=36)	M/ F	51.35	No significant differences were found between groups in the genus and species of their microbiotas.		
Wu et al (2016) ²⁶	US	The Controlled Feeding Experiment	C	N=31 (VG=15, OMN=16)	M/ F		The VG group had a significantly healthier metabolome compared to OMN but the gut microbiota were not significantly different. Higher consumption of fermentable substrate in VG was not associated with higher levels of fecal short chain fatty acids.		

Abbreviations: VG, vegan; VEG, vegetarian or ovo-lacto vegetarian; PESCO, pescatarian, pesco-vegetarian or “fish-eater”; OMN, omnivore, non-vegetarian or “meateater”; US, United States; N, number; C, cross-sectional; M, males; F, females.

Table 5 Hypertension

Study (Author, Year)	Location	Cohort	Design	Sample Description	M:F	Age (Mean Years)	Results/Follow-Up	Adjustment	Effect Sizes
Sanders & Key (1987) ²⁹	UK		C	N=44 (VG=22, OMN=22)	M/F		VG males had significant higher DBP than OMN males. No significant difference in SBP between VGs and OMNs, nor in DBP between VG females and OMN females.	Age, height, weight, physical activity, smoking habits, sex	d=1.20 (DBP, VG v. OMN, M); d=0.19 (DBP, VG v. OMN, F); d=0.19 (SBP, VG v. OMN, M); d=0.76 (SBP, VG v. OMN, F)
Famodu et al (1998) ³⁰	Nigeria		C	N=76 (VG=8, VEG=28, OMN=40)	M/F	48.6	No significant difference in SBP and DBP between the three groups.		d=0.10 (SBP, VG v. VEG); d=0.03 (SBP, VG v. OMN); d=0.09 (DBP, VG v. VEG); d=0.06 (DBP, VG v. OMN)
Toohy et al (1998) ³²	US		C	N=188 (VG=45, VEG=143)	M/F	50.8	No significant difference in SBP and DBP between the two groups, regardless of sex.		d=0.26 (SBP, M); d=0.09 (SBP, F); d=0.11 (DBP, M); d=0.34 (DBP, F)
Appleby et al (2002) ³³	UK	EPIC-Oxford cohort	C	N=11,004 (VG=739, VEG=3,800, PESC=1,728, OMN=4,737)	M/F		There were no significant differences in SBP and DBP between VG and other diet groups, regardless of gender.	Age, BMI, non-dietary factors, macronutrients, micronutrients	CI=122.7, 127.6 (SBP, VG, M); CI=117.9, 121.1 (SBP, VG, F); CI=75.8, 78.9 (DBP, VG, M); CI=72.1, 74.2 (DBP, VG, F); CI=123.7, 126.3 (SBP, OMN, M); CI=118.8, 120.0 (SBP, OMN, F); CI=76.1, 77.8 (DBP, OMN, M); CI=73.1, 73.9 (DBP, OMN, F)
Goff et al (2005) ³⁴	UK		C	N=49 (VG=24, OMN=25)	M/F	35.5	VGs had significantly lower SBP than OMNs. No significant difference in DBP between the two groups.	Age, sex, BMI	d=0.70 (SBP); d=0.34 (DBP); CI=-20.6, -1.3 (SBP); CI=-9.0, 2.5 (DBP)
Waldmann et al (2005) ³⁵	Germany	German VG Study	C	N=154 (strict VG=98, moderate VG=56)	M/F	44.2	No significant difference in SBP and DBP between the two groups.		d=0.28 (SBP); d=0.13 (DBP)
Fontana et al (2007) ³⁶	US		C	N=63 (Low calorie, low protein VG=21, OMN (endurance runner and sedentary)=42)	M/F	51.3	VGs had significantly lower SBP and DBP than both OMN groups.		d=1.28 (SBP, VG v. OMN ^a); d=1.93 (SBP, VG v. OMN ^b); d=1.00 (DBP, VG v. OMN ^a); d=1.77 (DBP, VG v. OMN ^b)
Lin et al (2010) ³⁷	Taiwan		C	N=204 (VG=102, OMN=102)	M	46.6	VG nuns had significantly lower SBP and significantly higher DBP than OMNs.	Age, BMI, education	d=0.29 (SBP); d=0.37 (DBP)

(Continued)

Table 5 (Continued).

Study (Author, Year)	Location	Cohort	Design	Sample Description	M:F	Age (Mean Years)	Results/Follow-Up	Adjustment	Effect Sizes
Petterson et al (2012) ³⁸	US	Adventist Health Study-2	C	N = 500 (VG=49, VEG=184, partial VEG=69, OMN=198)	M/F	62.6	VGs had significantly lower SBP and DBP than OMNs. No significant difference in SBP and DBP between VG, partialVEG, and VEG.	Age, sex	β CI=-12.9, -1.4 (SBP, VG ⁵); β CI=-9.0, -3.1 (DBP, VG ⁵)
Chiu et al (2015) ³⁹	Taiwan	MJ Health Screening	L, 2.12 y F/U	N=49,098 (VG=1,913, VEG=4,415 lacto-VEG=1,855, OMN=40,915)	M/F	48.9	VGs had lower risk of high SBP (≥ 130 mmHg) and high DBP (≥ 85 mmHg) than OMNs. Additional year of following VG diet does not lower risk of high SBP and high DBP compared to OMN diet. Comparisons between VGs and other VEG groups were not conducted.	Age, sex, study site	OR CI=0.73, 0.90 (SBP, VG ⁵); OR CI=0.73, 0.91 (DBP, VG ⁵)
Tong et al (2018) ³¹	UK	UK Biobank	C	N=418,749 (VG=378, VEG=6,366, PESC=9,674, poultry-OMN=4,381, low-OMN=199,784, OMN=198,166)	M/F	56.5	VG white women had lower SBP than regular- and low-OMN white women but were not significantly different from those in other dietary groups. VG white women had lower DBP than regular- and low-OMN and VEG white women but were not significantly different from those in other dietary groups. VG white men had lower SBP than regular- and low-OMN and VEG white men but were not significantly different from those in other dietary groups. VG white men had lower DBP than those in all other dietary groups.	Age, sex	CI=129.7, 134.3 (SBP, VG, F); CI=136.1, 136.3 (SBP, OMN, F); CI=134.9, 135.1 (SBP, low-OMN, F); CI=132.2, 133.4 (SBP, poultry-OMN, F); CI=132.3, 133.1 (SBP, PESC, F); CI=132.8, 133.8 (SBP, VEG, F); CI=76.5, 79.1 (DBP, VG, F); CI=81.1, 81.2 (DBP, OMN, F); CI=80.4, 80.5 (DBP, low-OMN, F); CI=78.5, 79.2 (DBP, poultry-OMN, F); CI=78.9, 79.3 (SBP, PESC, F); CI=79.3, 79.9 (SBP, VEG, F); CI=132.9, 138.2 (SBP, VG, M); CI=141.6, 141.8 (SBP, OMN, M); CI=140.7, 141.0 (SBP, low-OMN, M); CI=137.7, 138.9 (SBP, PESC, M); CI=138.3, 139.8 (SBP, VEG, M); CI=78.6, 81.7 (DBP, VG, M); CI=84.6, 84.7 (DBP, OMN, M); CI=83.9, 84.1 (DBP, low-OMN, M); CI=81.9, 83.1 (DBP, poultry-OMN, M); CI=82.0, 82.8 (SBP, PESC, M); CI=82.6, 83.5 (SBP, VEG, M);

Notes: ^aEndurance runner; ^bsedentary; ^cwith OMN as reference group.

Abbreviations: VG, vegan; VEG, vegetarian or ovo-lacto vegetarian; PESC, pescatarian, pesco-vegetarian or "fish-eater"; OMN, omnivore, non-vegetarian or "meateater"; UK, United Kingdom; US, United States; N, number; C, cross-sectional; L, longitudinal; y F/U, year follow-up; M, males; F, females; BMI, body mass index; CI, confidence interval; OR, odds ratio; d, effect size; SBP, systolic blood pressure; DBP, diastolic blood pressure; β , beta coefficient; mmHg, millimeters of mercury.

Table 6 Lipids

Study (Author, Year)	Location	Cohort	Design	Sample Description	M:F	Age (Years)	Results/Follow-Up	Adjustment	Effect Sizes
Sanders et al (1978) ⁴⁸	UK		C	N=54 (VG=26, OMN=28)	M/F	38	VG had significantly lower serum cholesterol level and serum triglyceride concentrations than OMN.	Age, sex, height, ethnic origin, SES	d=2.36 (serum TC); d=0.79 (serum TG)
Sanders & Key (1987) ²⁹	UK		C	N=44 (VG=22, OMN=22)	M/F		Plasma cholesterol concentrations were significantly lower in the male but not female VG compared with their respective controls.	Age, sex, body build	d=1.2 (plasma cholesterol, M); d=0.27 (plasma cholesterol, F)
Thorogood et al (1990) ⁴⁹	UK	Oxford VEG Study	C	N= 208 (VG=52, VEG=52, PESC=52, OMN=52)	M/F	42.41	In VG women the TC and LDL-C levels were significantly lower than in OMN and PESC women. In VG men, TC was significantly lower than in OMN men but not PESC men. In VG men, LDL-C levels did not significantly differ from PESC and OMN men. There were no significant differences in HDL-C among dietary groups.	Age, sex, BMI	d=1.06 (TC, VG v. OMN, M); d=0.74 (LDL-C, VG v. OMN, M); d=0 (HDL-C, VG v. OMN, M); d= 0.39 (TC, VG v. VEG, M); d=0.37 (LDL-C, VG v. VEG, M); d=0.04 (HDL-C, VG v. VEG, M); d= 0.65 (TC, VG v. PESC, M); d=0.58 (LDL-C, VG v. PESC, M); d=0.22 (HDL-C, VG v. PESC, M); d=1.20 (TC, VG v. OMN, F); d=1.34 (LDL-C, VG v. OMN, F); d=0.31 (HDL-C, VG v. OMN, F); d=0.66 (TC, VG v. VEG, F); d=0.66 (LDL-C, VG v. VEG, F); d=0.20 (HDL-C, VG v. VEG, F); d=0.97 (TC, VG v. PESC, F); d=0.94 (LDL-C, VG v. PESC, F); d=0.45 (HDL-C, VG v. PESC, F);

(Continued)

Table 6 (Continued).

Study (Author, Year)	Location	Cohort	Design	Sample Description	M:F	Age (Years)	Results/Follow-Up	Adjustment	Effect Sizes
Famodu et al (1998) ³⁰	Nigeria	Ilisan Cohort	C	N=76 (VG=8, semi-VEG=28, OMN=40)	M/F	48.6	VG had significantly lower serum TC and triglycerides than OMN. TC and triglycerides were not significantly different between VG and semi-VEG.		d=0.85 (serum TC, VG v. OMN); d=0.39 (serum TC, VG v. semi-VEG); d=1.03 (serum TG, VG v. OMN); d=0.71 (serum TG, VG v. semi-VEG)
Toohy et al (1998) ³²	Northeastern US		C	N=188 (VG=45, VEG=143; African American)	M/F	50.8	Serum TC, LDL-C, ratio of HDL-C, and triglycerides were significantly lower in VG compared to VEG.	Age, energy intake, waist to hip ratio	d=0.77 (serum TC, M); d=0.44 (LDL-C, M); d=0.85 (HDL-C, M); d=0.66 (TG, M); d=0.98 (serum TC, F); d=0.76 (LDL-C, F); d=0.37 (HDL-C, F); d=0.30 (TG, F)

Thomas et al (1999) ⁴⁰	UK		C	N=1,097 (VG=143, VEG=578, OMN=376)	F only	47.66	TC and LDL-C concentration were significantly higher in OMN than in VG, regardless of menopausal status. Mean HDL-C concentration was not significantly different between VG and OMN. VG and VEG were not directly compared, regardless of menopausal status.	Age, BMI, number of years past menopause, parity and hour of day of blood sample collection	CI=3.43, 3.67 (TC, VG ^a); CI=4.06, 4.29 (TC, OMN ^a); CI=3.78, 3.92 (TC, VEG ^a); CI=1.94, 2.13 (LDL-C, VG ^a); CI=2.52, 2.73 (LDL-C, OMN ^a); CI=2.23, 2.35 (LDL-C, VEG ^a); CI=1.23, 1.34 (HDL-C, VG ^a); CI=1.26, 1.35 (HDL-C, OMN ^a); CI=1.29, 1.34 (HDL-C, VEG ^a); CI=4.28, 4.81 (TC, VG ^b); CI=5.25, 5.48 (TC, OMN ^b); CI=4.90, 5.13 (TC, VEG ^b); CI=2.57, 3.02 (LDL-C, VG ^b); CI=2.52, 2.73 (LDL-C, OMN ^b); CI=2.23, 2.35 (LDL-C, VEG ^b); CI=1.23, 1.34 (HDL-C, VG ^b); CI=1.26, 1.35 (HDL-C, OMN ^b); CI=1.29, 1.34 (HDL-C, VEG ^b);
Goff et al (2005) ³⁴	UK		C	N=49 (VG=24, OMN=25)	M/F	35.5	VG group had significantly lower fasting plasma triacylglycerol than OMN. No differences in TC, HDL-C, LDL-C were found between VGs and OMNs.	Age, sex, BMI	d=0.54 (TC); OR CI=-0.96, 0.06 (TC); d=0.29 (HDL-C); OR CI=-0.32, 0.09 (HDL-C); d=0.06 (LDL-C); OR CI=-0.53, 0.44 (LDL-C); OR CI=-0.93, -0.37 (TG)
Waldmann et al (2005) ³⁵	Germany	The German Vegan Study	C	N=154 (strict VG=98, moderate VG=56)	M/F	44.2	Only TC differed statistically significantly between the subgroups.		

(Continued)

Table 6 (Continued).

Study (Author, Year)	Location	Cohort	Design	Sample Description	M:F	Age (Years)	Results/Follow-Up	Adjustment	Effect Sizes
Fontana et al (2007) ³⁶	US		C	N=63 (Low calorie, low protein VG=21, OMN (endurance runner and sedentary) =42)	M/F	51.3	Plasma concentrations of TC, LDL-C and triglycerides were significantly lower in the low-calorie low-protein VG diet than in the OMN group (sedentary). There were no significant differences in HDL-C between the VG and sedentary OMN groups.		d=1.55 (TC, VG v. OMN ⁵); d=1.38 (LDL-C, VG v. OMN ⁵); d=0.33 (HDL-C, VG v. OMN ⁵); d=1.30 (TG, VG v. OMN ⁵)
Lin et al (2010) ³⁷	Taiwan		C	N=204 (VG=102, OMN=102)	F only	46.6	VG nuns had significantly lower cholesterol levels than OMN. No significant difference was found in triglycerides.	Age, BMI, education	d=0.55 (cholesterol); d=0.26 (TG)
Shang et al (2011) ⁴²	Taiwan	TMJ Health Screening Center Health Check-Up Database	C	N=93,209 (VG=1,116, lacto VEG=4,313, PESC=2,461, non VEG=85,319)	M/F	37.1	Compared with VG, OMN and PESC had significantly lower risk of developing low HDL-C, respectively. No difference in low HDL-C risk between VG and VEG nor in high TG risk between VG and other groups.	Age, sex, education status, smoking status, drinking status, physical activity at work and leisure	HR CI=0.74, 1.09 (high TG, OMN ⁴); HR CI=0.71, 1.02 (high TG, PESC ⁴); HR CI=0.78, 1.09 (high TG, VEG ⁴); HR CI=0.62, 0.84 (low HDL-C, OMN ⁴); HR CI=0.57, 0.84 (low HDL-C, PESC ⁴); HR CI=0.83, 1.17 (low HDL-C, VEG ⁴)
Vinagre et al (2013) ⁴¹	Brazil		C	N=50 (VG=21, OMN=29)	M/F	36.16	Serum LDL-C and cholesterol were significantly lower in VG than in OMN, but HDL-C and triglycerides were not significantly different.	Sex, gender, BMI, cholesterol, LDL-C, HDL-C, triglycerides, glucose, and intake of energy, carbohydrates, protein, fiber, fat, saturated fat, polyunsaturated fat, monounsaturated fat, and cholesterol.	d=0.87 (TC); d=0.80 (LDL); d=0.39 (HDL); d=0.25 (TG)

Jian et al (2014) ⁴³	Taiwan	2002 Taiwanese Survey on the Prevalence of Hyperglycemia, Hyperlipidemia and Hypertension (TsSHHH)	C	N=6,808 (VG=144, VEG=195, OMN=6,469)	M/F	46.2	VG (men and women) had significantly lower HDL-C than OMN. LDL-C and TC were not significantly different in women but were significantly lower in VG v. OMN men. Triglycerides were significantly higher in VG v. OMN women. VG and VEG women and men did not significantly differ on any of these comparisons.	d=0.33 (HDL-C, VG v. OMN, F); d=0.12 (LDL-C, VG v. OMN, F); d=0.01 (TC, VG v. OMN, F); d=0.35 (TG, VG v. OMN, F); d=0.39 (HDL-C, VG v. OMN, M); d=0.45 (LDL-C, VG v. OMN, M); d=0.49 (TC, VG vs OMN, M); d=0.15 (TG, VG v. OMN, M)
Huang et al (2014) ⁴⁴	Taiwan	TsSHHH	C	N=3,551 (VG=99, VEG=127, OMN=3,325)	F only	43.3	In premenopausal women: HDL-C was significantly lower in VG compared to OMN. Triglycerides were significantly higher in VG compared to OMN. TC and LDL-C were not significant. In postmenopausal women: HDL-C was significantly lower in VG compared to OMN. TC was significantly higher in OMN compared to VG. LDL-C and triglycerides were not significant. There were no significant differences between VG and VEG in either pre- or postmenopausal women.	d=0.48 (HDL-C, VG v. OMN ^a); d=0.20 (HDL-C, VG v. VEG ^a); d=0.14 (LDL-C, VG v. OMN ^a); d=0.38 (LDL-C, VG v. VEG ^a); d=0.45 (TG, VG v. OMN ^a); d=0.40 (TG, VG v. VEG ^a); d=0.05 (TC, VG v. OMN ^a); d=0.24 (TC, VG v. VEG ^a); d=0.32 (HDL-C, VG v. OMN ^b); d=0.05 (HDL-C, VG v. VEG ^b); d=0.28 (LDL-C, VG v. OMN ^b); d=0.22 (LDL-C, VG v. VEG ^b); d=0 (TG, VG v. OMN ^b); d=0.25 (TG, VG v. VEG ^b); d=0.17 (TC, VG v. VEG ^b); d=0.37 (TC, VG v. OMN ^b)

(Continued)

Table 6 (Continued).

Study (Author, Year)	Location	Cohort	Design	Sample Description	M:F	Age (Years)	Results/Follow-Up	Adjustment	Effect Sizes
Bradbury et al (2014) ⁴⁵	UK	EPIC-Oxford Cohort	C	N=1,694 (VG=422, VEG=423, PESC=425, OMN=424)	M/F		VG had significantly lower serum TC than OMN, VEG and PESC. VG males had significantly lower HDL-C than PESC but were not significantly different from OMN and VEG. VG females had significantly lower HDL-C than PESC and OMN but were not significantly different from VEG.	Age, alcohol, physical activity	CI=4.28, 4.55 (TC, VG, M); CI=5.15, 5.41 (TC, OMN, M); CI=4.94, 5.21 (TC, PESC, M); CI=4.73, 5.00 (TC, VEG, M); CI=1.11, 1.19 (HDL-C, VG, M); CI=1.12, 1.21 (HDL-C, OMN, M); CI=1.18, 1.27 (HDL-C, PESC, M); CI=1.11, 1.20 (HDL-C, VEG, M); CI=4.36, 4.55 (TC, VG, F); CI=4.95, 5.15 (TC, OMN, F); CI=4.76, 4.96 (TC, PESC, F); CI=4.66, 4.86 (TC, VEG, F); CI=1.34, 1.42 (HDL-C, VG, F); CI=1.45, 1.52 (HDL-C, OMN, F); CI=1.43, 1.50 (HDL-C, PESC, F); CI=1.38, 1.45 (HDL-C, VEG, F)
Chiu et al (2015) ³⁹	Taiwan (Republic of China)	MJ Health Screening	L, 2.12 y F/U	N=49,098 (VG=1,913, VEG=4,415 lacto-VEG=1,855, OMN=40,915)	M/F	48.9	VG had a significantly higher risk of low HDL-C than OMN, while VG had significantly lower risk of high LDL-C AND TC than OMN. VG had significantly higher risk of high triglycerides than OMN. VG and VEG were not compared directly.	Age, sex, education LTPA, alcohol consumption, study site	OR CI=0.99, 1.24 (TG, VG°, B=baseline measurement); OR CI=0.36, 0.45 (TC, VG°, B); OR CI=0.40, 0.50 (LDL-C, VG°, B); OR CI= 1.39, 1.68 (HDL-C, VG°, B); OR CI=0.90, 1.08 (TG, VG°, T=temporal effect); OR CI=0.89, 1.05 (TC, VG°, T); OR CI=0.81, 1.03 (LDL-C, VG°, T); OR CI= 0.94, 1.12 (HDL-C, VG°, T);

Elorinne et al (2016) ⁴⁶	Finland	C	N=41 (VG=22, OMN=19)	M/F	34	Serum TC, HDL-C, and LDL-C were significantly lower in VG than OMN. Triglycerides were not significantly different.		
Kuchta et al (2016) ⁴⁷	Poland	C	N=42 (VG=21, OMN=21)	M/F	28	There were significantly lower concentrations of TC, LDL-C, and non-HDL-C, and no difference in triglycerides in VG versus OMN.		

Notes: ^aPremenopausal women, ^bpostmenopausal women, ^csedentary OMN, ^dVG as reference group, ^eOMN as reference group.

Abbreviations: VG, vegan; VEG=vegetarian or ovo-lacto vegetarian; PES, pescatarian; pesco-vegetarian or "fish-eater"; OMN, omnivore; non-vegetarian or "meateater"; UK, United Kingdom; US, United States; N, number; C, cross-sectional; y F/U, year follow-up; M, males; F, females; TC, total cholesterol; LDL-C, low-density lipoproteins; HDL-C, high-density lipoproteins; TG, triglycerides; BMI, body mass index; SES, socio-economic status; LTPA, leisure time physical activity; CI, confidence interval; HR, hazard ratio; OR, odds ratio; d, effect size; T, temporal effect; B, baseline data.

cancer, lung cancer, breast cancer, and prostate cancer), ischemic heart disease, cardiovascular disease, cerebrovascular disease, diseases of the respiratory system, other causes and all causes.^{18–20} However, in a US study using an omnivore comparison group, vegans had a significantly lower death rate HR from other causes (excluding heart disease, cardiovascular disease, and cancer).¹⁹ Moreover, for men, vegans' death rate HR (from all causes, ischemic heart disease, and cardiovascular disease) was significantly lower compared to omnivores.

Gut Microbiome. The gut microbiome consists of various commensal microbial species (thought to be >500) in the gastrointestinal tract including bacteria, viruses, archaea, and eukaryotic microbes.^{21,22} It has been increasingly studied due to its link to diseases, such as cardiovascular disease, inflammatory bowel disease, and infections (eg, *Clostridium difficile*). Research has found positive effects of a plant-based diet on the microbiome, and, in turn, on disease.²³

Four cross-sectional studies (see Table 4)^{24–27} assessed the gut microbiota of 754 participants (total). In European studies significantly lower microbial counts of the *Bacteroides fragilis* group and the mesophilic/thermophilic LAB loads (on MRS agar)²⁴ and of the *Bacteroides* spp., *Bifidobacterium* spp., *Escherichia coli* and *Enterobacteriaceae* spp. groups as well as lower stool pH²⁵ were found in vegan fecal microbiota compared to omnivores. Contrary to these findings, a US study found neither a difference in the gut microbiota between vegans and omnivores nor an association between higher consumption of fermentable substrate and higher levels of fecal short-chain fatty acids. The only reported significant finding was a healthier metabolome for vegans compared to omnivores.²⁶ Similarly, Ruengsomwong et al²⁷ compared several Thai vegetarian groups including a small vegan sample and found no significant differences in the genus and species of their microbiotas.

Hypertension. Hypertension is a risk factor for cardiovascular and cerebrovascular diseases.²⁸ The 11 included articles^{29–39} reporting blood pressure data in vegans and other dietary groups were published between 1987 and 2018 (see Table 5). Data from large cohorts were reported in five studies, including the Adventist Health Study-2 (N=1),³⁸ EPIC-Oxford (N=1),³³ UK Biobank (N=1),³¹ German Vegan Study (N=1),³⁵ and MJ Health Screening (N=1).³⁹ The total sample size was 478,218 (mean age=55.5 years), including 3,399 vegans, 448,412 omnivores

Table 7 Overweight

Study (Author, Year)	Location	Cohort	Design	Sample Description	M:F	Age (Mean Years)	Results/Follow-Up	Adjustment	Effect Sizes
Sanders et al (1978) ⁴⁸	UK		C	N=54 (VG=26, OMN=28)	M/F	38	VG had significantly lower weight than OMN.	Age, sex, height, ethnic origin, SES	d=0.78 (weight)
Barr et al (1998) ⁵⁹	Canada		L, 13 mo F/U	N=45 (VG=8, VEG=15, OMN=22)	F only	27.2	VGs had similar BMI with VEGs, but lower BMI than OMNs.		d=1.08 (BMI, VG v. OMN); d=0.23 (BMI, VG v. VEG)
Famodu et al (1998) ³⁰	Nigeria		C	N=76 (8=VG, 28=VEG, 40=OMN)	M/F	48.6	No significant group difference in BMI.		d=0.73 (BMI, VG v. OMN); d=0.67 (BMI, VG v. VEG)
Spencer et al (2003) ⁶⁰	UK	EPIC-Oxford	C	N=37,875 (VG=1,553, VEG=12,307, PESC=6,191, OMN=17,824)	M/F	40.5	VGs had significantly lower BMI than VEGs and OMNs, for both genders. BMI of VGs and PESC were not significantly different from each other, for both genders.	Age, sex, lifestyle factors, dietary factors	CI=22.83, 23.43 (BMI, VG, M); CI=23.54, 23.80 (BMI, VEG, M); CI=23.97, 24.20 (BMI, OMN, M); CI=23.27, 23.64 (BMI, PESC, M); CI=22.32, 22.79 (BMI, VG, F); CI=22.88–23.04 (BMI, VEG, F); CI=23.17–23.31 (BMI, OMN, F); CI=22.73, 22.92 (BMI, PESC, F)
Fontana et al (2005) ⁵¹	US		C	N=36 (Raw Food VG=18, OMN=18)	M/F	54.2	Raw Food VGs had significantly lower BMI compared to OMNs.	Age, sex, SES	d=1.72 (BMI)
Newby et al (2005) ⁵²	Sweden	Swedish Mammography Cohort	C	N=55,459 (VG=83, lacto-VEG=159, semi-VEG=960, OMN=54,257)	F only	51.39	BMI of semi-VEG, lacto-VEG, and VG were significantly lower than that of OMN (p<0.01). BMI of the vegetarian diets were not significantly different from each other. The prevalence of overweight was 30% for OMNs, 24% for semi-VEGs, 21% for lacto-VEGs, and 23% for VGs. The prevalence of obesity was 10% for OMNs, 5% for semi-VEGs, 4% for lacto-VEGs, and 6% for VGs. VGs had significant lower overweight/obesity (BMI>25) OR than OMN.		d=0.36 (BMI, VG v. OMN); d=0.08 (BMI, VG v. semi-VEG); d=0.03 (BMI, VG v. lacto-VEG)

Rosell et al (2005) ⁵³	UK	EPIC-Oxford	C	N=659 (VG=232, VEG=231, OMN=196)	M only	46.8	VG had significantly lower BMI than OMN. No significance test was conducted to compare VGs and VEGs.	d=1.03 (BMI, VG v. OMN); d=0.03 (BMI, VG v. VEG)
Waldmann et al (2005) ³⁵	Germany	German VG Study	C	N=154 (strict VG=98, moderate VG=56)	M/F	44.2	No difference in BMI between groups.	d=0 (BMI)
Rosell et al (2006) ⁵⁸	UK	EPIC-Oxford	L, 5 y F/U	N=21,966 (VG=609, VEG=5,277, PESC=2,504, OMN=10,784, reverted=1,826, converted=966)	M/F	43.8	No difference in weight gain (g/year) between vegan and other groups for men. VG women had significantly lower weight gain than OMN women over the 5 years.	CI=373, 439 (WG, OMN, M); CI=298, 456 (WG, PESC, M); CI=339, 433 (WG, VEG, M); CI=178, 390 (WG, VG, M); CI=374, 563 (WG, reverted, M); CI=133, 351 (WG, converted, M); CI=403, 443 (WG, OMN, F); CI=300, 376 (WG, PESC, F); CI=364, 420 (WG, VEG, F); CI=211, 396 (WG, VG, F); CI=388, 479 (WG, reverted, F); CI=238, 365 (WG, converted, F);
Fontana et al (2007) ³⁶	US		C	N=63 (Low calorie, low protein VG=21, OMN (endurance runner and sedentary)=42)	M/F	51.3	Low-calorie low-protein VGs had significant lower BMI than sedentary OMNs (p<0.01). No difference between low-calorie low protein VGs and endurance OMN runners was found in their BMI.	d=1.79 (BMI, VG v. OMN ^a); d=0.08 (BMI, VG v. OMN ^b)
Ho-Pham et al (2009) ⁵⁴	Vietnam		C	N=210 (VG=105, OMN=105)	F only	62	No significant group difference in BMI.	d=0 (BMI)
Tonstad et al (2009) ¹¹	US, Canada	The Adventist Health Study-2	C	N=60,903 (VG=2,731, VEG=20,408, PESC=5,617, semi-VEG=3,386, OMN=28,761)	M/F	53	BMI of VGs was significantly lower than that of OMN. BMI was lowest in VG and incrementally higher in lacto-ovo VEG, PESC, semi-VEG, and OMN.	d=0.96 (BMI, VG v. OMN); d=0.44 (BMI, VG v. VEG); d=0.56 (BMI, VG v. PESC); d=0.73 (BMI, VG v. semi-VEG)

(Continued)

Table 7 (Continued).

Study (Author, Year)	Location	Cohort	Design	Sample Description	M:F	Age (Mean Years)	Results/Follow-Up	Adjustment	Effect Sizes
Rizzo et al (2013) ⁵⁵	US, Canada	Adventist Health Study-2	C	N=71,752 (VG=5,694, VEG=21,799, Semi-VEG=4,042, PESC=6,583, OMN=33,634)	M/F	59	BMI of VGs were significantly lower than that of OMNs, VEGs, semi-VEGs, and PESCs. The prevalence of overweight (BMI=25–29.99) was 23.7% for VGs, 33.0% for VEGs, 35.5% for PESCs, 37.0% for semi-VEGs, and 37.3% for OMNs. The prevalence of obesity (BMI≥30) was 9.4% for VGs, 16.7% for VEGs, 17.9% for PESCs, 24.2% for semi-VEGs, and 33.3% for OMNs.		CI=24.0, 24.2 (BMI, VG), CI=26.0, 26.2 (BMI, VEG), CI=26.0, 26.2 (BMI, PESC), CI=27.3, 27.6 (BMI, semi-VEG), CI=28.6, 28.7 (BMI, OMN)
Schmidt et al (2013) ⁵⁶	UK	EPIC-Oxford	C	N=1,629 (VG=412, VEG=404, PESC=405, OMN=408)	M/F	42	VGs had significant lower BMI than OMNs, for both genders. Significance tests comparing VGs, PESCs, and VEGs were not conducted.	Age, sex	d= 0.79 (BMI, VG v. OMN, M); d=0.26 (BMI, VG v. PESC, M); d=0.31 (BMI, VG v. VEG, M); d=0.57 (BMI, VG v. OMN, F); d=0.18 (BMI, VG v. PESC, F); d=0.29 (BMI, VG v. VEG, F)
Agrawal et al (2014) ¹⁴	India	Indian's Third National Family Health Survey	C	N=156,317 (VG=2,560, lacto-VEG=37,797, VEG=5,002, semi-VEG=8,140, PESC=3,446, OMN=99,372)	M/F		BMI was not significantly different among the dietary groups. The percentage of overweight (BMI≥25) in VGs did not significantly differ from that of OMNs, PESCs and semi-VEGs, but was significantly lower than that of lacto-ovo-VEGs and lacto-VEGs. The percentage of obesity (BMI≥30) in VGs did not significantly differ from that of lacto-ovo VEGs, PESCs, semi-VEGs, and OMNs, but was significantly lower than that of lacto-VEGs.	Age, sex, education, household wealth, rural/urban residence, religion, caste, smoking, alcohol use, television watching	d=0.05 (BMI, VG v. OMN); d=0.16 (BMI, VG v. lacto-VEG); d=0.12 (BMI, VG v. lacto-ovo VEG); d=0.05 (BMI, VG v. PESC); d=0.02 (BMI, VG v. semi-VEG)

Chiu et al (2015) ³⁹	Taiwan	MJ Health Screening	F, 2.12 y F/U	N=49,098 (VG=1,913, VEG=4,415 lacto- VEG=1,855, OMN=40,915)	M/F	48.9	At baseline, VGs had significantly lower OR of overweight (BMI≥27) than OMNs. Compared to OMN, each additional year of VG diet reduce the risk of obesity by 7%. VG were not compared directly to VEG.	Age, sex, education LTPA, alcohol consumption, study site	OR CI=0.59, 0.78 (obesity, VG v. OMN, B); OR CI=0.88, 0.99 (obesity, VG v. OMN, T)
Tonstad et al (2015) ⁵⁷	US	Adventist Health Study-2	C	N= 65,981 (VG=5,389, VEG=18,390, semi-VEG=3,681, PESC=6,420, OMN=32,101)	M/F	56.7	VGs had significantly lower BMI than OMNs. No significance tests were conducted to compare VGs, VEGs, semi-VEGs, and PESCs.		d=0.87 (BMI, VG v. OMN); d=0.39 (BMI, VG v. lacto-ovo VEG); d=0.64 (BMI, VG v. semi-VEG); d=0.47 (BMI, VG v. PESC)
Tong et al (2018) ³¹	UK	UK Biobank	C	N=418,749 (VG=378, VEG=6,366, PESC=9,674, poultry- OMN=4,381, low- OMN=199,784, OMN=198,166)	M/F	56.5	VG white women had significantly lower BMI than regular- and low-OMN white women but were not significantly different from other dietary groups. VG white men had significantly lower BMI than white men in all other dietary groups.	Age, sex	CI=24.2, 25.5 (BMI, VG, F); CI=27.4, 27.5 (BMI, OMN, F); CI=26.6, 26.6 (BMI, low-OMN, F); CI=24.9, 25.2 (BMI, poultry-OMN, F); CI=24.9, 25.2 (BMI, PESC, F); CI=25.1, 25.4 (BMI, VEG, F); CI=24.2, 25.5 (BMI, VG, M); CI=28.0, 28.0 (BMI, OMN, M); CI=27.3, 27.4 (BMI, low-OMN, M); CI=25.7, 26.2 (BMI, poultry-OMN, M); CI=25.6, 25.9 (BMI, PESC, M); CI=25.6, 26.0 (BMI, VEG, M)

Notes: ^aSedentary OMN, ^bendurance OMN.

Abbreviations: VG=vegetarian; VEG=vegetarian or ovo-lacto vegetarian; PESC=pescatarian, pesco-vegetarian or “fish eater”; OMN=omnivore, non-vegetarian or “meateater.” UK= United Kingdom, US= United States, N=number, C=cross-sectional, L=longitudinal, y F/U=year follow-up, mo F/U= month follow-up, M=males, F=females, BMI=body mass index, SES=socio-economic status, LTPA=leisure time physical activity, CI=confidence interval, OR=odds ratio, d=effect size, WG=weight gain, B=baseline data, T=temporal effect.

14,936 vegetarians and 11,402 pescatarians, and 69 partial (semi) vegetarians.

In US studies, vegans had lower systolic blood pressure (SBP) and diastolic blood pressure (DBP) compared to omnivores,^{36,38} although no significant differences in blood pressure were found between vegans, vegetarians, and partial vegetarians.^{32,38} The majority of the remaining comparisons (across several studies)^{29,30,32–35} found no significant differences in blood pressure between vegans and other dietary groups with the following exceptions: 1) 2 studies reported significantly lower SBP in vegans compared to omnivores,^{34,37} 2) 1 study found higher DBP in vegans than omnivores³⁷ and another reported higher DBP in vegan men than omnivore men,²⁹ and 3) among white participants, one study found in women significantly lower SBP (vegans versus omnivore) and DBP (vegans versus omnivores and vegetarians) and in men significantly lower SBP (vegans versus omnivores and vegetarians) and DBP (vegans versus all other dietary groups).³¹

Lipids. High serum low-density lipoprotein (LDL-C) cholesterol is a leading risk factor for several diseases, including stroke, coronary heart disease, and myocardial infarction.²⁸ Observational studies (N=18; see Table 6),^{29,30,32,34–37,39–49} comparing cholesterol levels among dietary groups were published between 1978 and 2016 included a total of 149,838 participants, comprised of vegans (n=4,299), vegetarians (n=5,933), pescatarians (n=2,398), and omnivores (n=137,208). Most studies (N=11) found that total cholesterol (TC) was significantly lower in vegans compared to omnivores,^{30,36,37,41,44,45,47–49} vegetarians,³² and moderate vegans (animal products < 5% diet).³⁵ Only one study reported no difference in TC in vegans compared to omnivores.³⁴ In most studies, LDL-C was found to be significantly lower, in vegans compared to omnivores,^{36,40,41,43,46,47,49} with one showing lower levels in pescatarians⁴⁹ (women) and another in vegetarians.³²

High-density lipoprotein (HDL-C) levels were lower in vegan groups across studies,^{32,43–45} with subgroup analyses showing significantly lower HDL-C in vegan women.⁴⁴ Shang et al⁴² found that compared to vegans, omnivores and pescatarians were at reduced risk of developing low HDL-C. At the same time, other research reported no significant differences in HDL-C levels between vegans and other diet groups.^{34,36,40,41}

Several studies reported that vegans had significantly lower triglycerides compared to omnivores^{30,34,36,48} and vegetarians.³² Other research found no difference in

triglycerides in vegans compared to omnivores,^{37,41,44,46,47} or higher triglycerides in vegans compared to omnivores.^{39,43,44}

Overweight. In studies, overweight is typically assessed using BMI, which is calculated by dividing weight in kilograms by height in meters squared. Individuals with a BMI between 25 and 29.9 are considered overweight, while those with a BMI equal to or greater than 30 are considered obese.⁵⁰

The 18 observational studies^{11,14,30,31,35,36,39,48,51–60} were published between 1978 and 2018 (see Table 7). In 12 articles, data from larger studies were reported, including EPIC-Oxford (N=4),^{53,56,58,60} Adventist Health Study-2 (N=3),^{11,55,57} Indian's Third National Family Health Survey (N=1),¹⁴ German Vegan Study (N=1),³⁵ Swedish Mammography Cohort (N=1),⁵² MJ Health Screening (N=1)³⁹ and UK Biobank (N=1).³¹ Including only the most recent reports, the total sample size for these studies was 808,079, including 12,853 vegans, 92,381 vegetarians, 12,781 semi-vegetarians, 28,152 pescatarians, and 661,912 omnivores.

In US studies, vegans had significantly lower BMI than omnivores, vegetarians, pescatarians, and semi-vegetarians.^{11,36,51,55,57} The BMI of Canadian vegans was not significantly different from that of vegetarians but was significantly lower than that of Canadian omnivores.⁵⁹

In the UK, vegans had significantly lower BMI than omnivores, regardless of sex.^{31,48,53,56,58} One study⁶⁰ found the BMI of vegans was significantly lower than that of vegetarians and was not significantly different from that of pescatarians. Further, weight gain over 5 years was significantly less in vegan compared to omnivore women, but this difference was not observed in men.⁵⁸ However, when white men were selected for analysis, BMIs for vegans were significantly lower than those of all other diet groups.³¹

In India, diet groups had similar BMI profiles.¹⁴ The percentage of vegans that were overweight did not significantly differ from that of pescatarians, semi-vegetarians, and omnivores, but was significantly lower compared to vegetarians and lacto-vegetarians. The same pattern of results was found in comparisons of obesity rates with the exception that the percentage of vegans who were obese did not significantly differ from the percentage of vegetarians who were obese.

The mean BMI of Swedish vegans did not differ from those of vegetarians and semi-vegetarians, but was significantly lower than that of omnivores.⁵² Taiwanese vegans

had significantly lower BMIs than omnivores, with each additional year of following a vegan diet lowering the risk of obesity.³⁹ No significant differences were found in the BMIs of Vietnamese⁵⁴ and Nigerian³⁰ vegans versus omnivores.

Discussion

Based on a review of this literature, several health benefits of a vegan diet were identified. Compared to omnivores, vegans were less likely to be diagnosed with NIDDM in the US^{11,12} but not India.¹⁴ There was reduced mortality risk for all vegans (other causes) and male vegans (all causes, ischemic heart disease, cardiovascular disease) in the US¹⁹ but not UK¹⁸ or in a larger study combining data from multiple cohorts.²⁰ Vegans were at reduced risk for incidence of “any” cancer^{15,16} (US and UK) and of female cancers¹⁵ (US), but only when analyses were not adjusted for BMI.

Mean blood pressure was lower in vegans compared to omnivores from the US,^{36,38} whereas in other locales there were mixed findings for analyses of SBP and DBP. Similarly, compared to omnivores, vegans’ BMIs were significantly lower in some countries (eg, US,^{11,36,51,55,57} Sweden,⁵² Canada,⁵⁹ Taiwan³⁹) but not others (eg, India,¹⁴ Nigeria,³⁰ Vietnam⁵⁴). The majority of observational studies found superior lipid profiles (except HDL-C and triglycerides) in vegans compared to other diet groups.^{30,31,37,46-49} However, observational studies of the vegan microbiome reported mixed results.²⁴⁻²⁷

Significantly lower rates of NIDDM were found for vegans versus omnivores in two large-scale North American studies.^{11,12} Although not reported in Table 1, when BMI was not included as a covariate, in the US, vegans were less likely to be diagnosed with NIDDM compared to all dietary groups, suggesting that BMI is an important mediator for the analyses of vegans versus vegetarians and pescatarians. Similarly, in a UK study, vegans’ risk of NIDDM was significantly less than omnivores’, but only when there was no BMI adjustment.¹³ These results are consistent with other research reporting an average diabetes risk reduction of 47–78% in vegans.⁶¹ However, adopting a vegan diet did not result in a significant difference in diabetes rates in a large Indian study.¹⁴ Agrawal et al¹⁴ surmise that the lack of reduced relative risk for their vegan sample compared to the advantage reported for US vegan diabetics was due to 1) the belief among Indians that consuming ghee was permitted

on a vegan diet, 2) the consumption of more refined rice by Indian vegans, or 3) reverse causality with some diabetics adopting a vegan diet to better manage their disease. It is unclear whether cultural and socio-economic factors affect diet content and whether other intersecting factors (eg, diabetics choosing a vegan diet to manage symptoms), in turn, affect diet-related prevalence of NIDDM.

Two seminal studies of vegan and omnivore diets provided evidence that a vegan diet decreases “any” cancer risk, but only when the effects of BMI are not partialled out, with the implication being that it is an important mediator.^{15,16} However, in a recent meta-analysis,⁶² when the results of these studies were aggregated, there was a significant risk reduction for vegans even when BMI was covaried, because of the larger sample size of the combined studies. However, there was no support that a vegan diet offered protection against diagnosis of most specific cancers with the exception of female-specific cancers, but only when BMI was not a covariate.¹⁵ Relatively few studies have investigated the link between veganism and cancer probably due to the low prevalence of vegan cancers in the population when this research was conducted.⁶³

For both sexes together, vegans’ mortality rate for causes other than heart disease, cardiovascular disease and cancer was significantly lower than that of omnivores in a US study¹⁹ but not significantly different in a UK study,¹⁸ and combined analysis of five data sets.²⁰ In the Orlich et al study, men’s death rates were significantly lower in vegans from all causes, ischemic heart disease and cardiovascular disease.¹⁹ Dinu et al⁶² found a benefit in mortality rates for vegans considered together with vegetarians, but not for vegans alone. However, they did not break down their analysis by sex. It is unclear why, despite the many health benefits of a vegan diet (eg, reduced risk of NIDDM and cancer (“any”), lower TC and LDL cholesterol, and lower BMI), there was not more evidence for a mortality advantage. One possibility is that the relatively advanced health care systems in the countries where this research was conducted were able to effectively treat diseases, therefore delaying mortality in sick people.¹⁸ Insufficient sample size representing the vegan population is also partly responsible for the sparseness of this evidence.²⁰

Investigations of the microbiome of vegans and omnivores have reported mixed results. Two cross-sectional studies of vegans and omnivores reported a comparatively favorable profile for the gut microbiota in the vegan

sample, suggesting potential positive effects on fecal flora of a vegan diet.^{24,25} At the same time, a fecal analysis examining the gut make-up of individuals following different vegetarian diets did not reveal significant differences.²⁷ However, the Ruengsomwong et al sample of vegans was small ($n=3$). Similarly, in US urban vegans, the microbiome was not significantly different from that of omnivores, but there were significant differences in their plasma metabolome.²⁶ Specifically, in the plasma metabolome of vegans, compared to omnivores there were more numerous bacterial metabolites and fewer amino acid metabolites, differences associated with improved health outcomes such as increases in beneficial bacterially generated metabolites (eg, equol) and reductions in harmful metabolites like trimethylamine. The authors note that gut microbiota affected the make-up of the plasma metabolome to a greater extent in vegans than in omnivores and that environmental differences can affect gut microbiota, an observation that is in line with the geographic differences in findings described in these articles.

Compared to omnivores, the lower mean SBPs and DBPs in vegans were found in US studies.^{36,38} Moreover, vegans' SBP and DBP levels did not significantly differ from those of vegetarians, semi-vegetarians, and pescatarians. In the remainder of the research, there were no significant blood pressure differences in comparisons of vegans to other dietary groups^{29,30,32–35} with only a few exceptions.^{29,31,34,37} Other considerations may have impacted these results, for example that persons with hypertension routinely take medications to lower their blood pressures. Indeed, examination of means for dietary groups revealed that most were well within normotensive ranges.

The large majority of observational studies found more favorable lipid profiles in vegans than in omnivores and other comparison groups.^{30,36,37,41,45–49} These findings are consistent with results from a recent meta-analysis assessing the relationship between lipids and vegetarian diets.⁴ There is a strong relationship between high levels of TC and LDL-C and intake of saturated fat, which in the human omnivore diet is derived predominantly from animal sources.⁶⁴

The varied results for triglycerides may be explained by geographic differences. Vegans had higher triglyceride levels in studies conducted in Taiwan whereas other research showed they had either lower levels or levels that were not significantly different. In trying to explain this discrepancy, Shang et al⁴² note that the omnivore diet

in Taiwan differs from the typical Western diet in that there are more plant foods, no dairy and lesser quantities of other animal-derived foods. This finding is consistent with a meta-analysis of vegan diets and cardio-metabolic risk factors that also found a different pattern of results in some Taiwan studies.⁶⁵

The results for HDL-C levels were also mixed, with studies reporting either less favorable levels for vegans or non-significant differences. Low HDL-C and high triglycerides have been considered components of metabolic syndrome and independent risk factors for cardiovascular disease.⁴³ However, other research has cast doubt on this relationship with respect to HDL-C⁴⁴ as interventions⁶⁶ and genetic variants⁶⁷ that increase HDL-C do not reduce coronary heart disease risk. As later follow-up analyses showed no associations, Chiu et al³⁹ proposed that poorer triglycerides and HDL-C in vegetarians and vegans may be due to “reverse causation”, in which individuals with medical conditions switch to a vegetarian diet in order to improve their health.

Vegans had lower BMI or overweight/obesity rates than omnivores in some countries (eg, US,^{11,36,51,55,57} Sweden,⁵² Canada,⁵⁹ Taiwan³⁹), but these differences were not found in other countries (eg, India,¹⁴ Nigeria,³⁰ Vietnam⁵⁴). In the Nigerian study,³⁰ there were only 8 vegans, so that comparison was likely insufficiently powered. The Vietnamese study⁵⁴ assessed vegan Buddhist nuns and matched controls. In both groups, the mean BMI was 24, well within the normal range. The larger US studies in this review did not report average BMI of their female participants. However, in the US, the BMI of the average adult woman is 26.5,⁶⁸ which is in the overweight range. The data for the Indian sample¹⁴ were divided according to weight status category (eg, underweight, overweight, obese) and the proportions in each following different diets were compared. Across diet categories, the percentage of individuals who were overweight or obese was small by Western standards, ranging from 10% in pescatarians (11.5% in vegans) to 16.2% in lacto-vegetarians. Considering these results and those of the other observational studies that were reviewed, compared to an omnivore diet consuming a vegan diet appears to provide some protection against overweight in non-Asian countries, possibly because obesity is more prevalent in these locations. In terms of the differences between vegans versus vegetarians, semi-vegetarians, and pescatarians, the results were inconsistent.

An important finding from this review was that demographic variables moderated the relationship between consuming a vegan diet and some outcomes. For example, in a US

study, there were sex differences in mortality (from all causes, ischemic heart disease and cardiovascular disease) with male vegans reporting significantly lower rates than their omnivore counterparts while there was not a significant difference in female mortality rates.¹⁹ In the UK, vegan white women had lower SBP and DBP than omnivore white women and lower DBP than vegetarian white women. Vegan white men had lower SBP and DBP than their vegetarian and omnivore counterparts, with DBP also lower than all other white male dietary groups. Geographic location exerted a moderating effect, with the most wide-ranging evidence for the health benefits of a vegan diet coming from the US. In American studies there were lower rates of NIDDM compared to omnivores;^{11,12} reduced incidence of “any” and female cancer relative to omnivores when data were aggregated with UK data;¹⁵ decreased “other cause” mortality risk in men, as well as reduced mortality risk due to cardiovascular disease, coronary heart disease, and all cause compared to omnivores;¹⁹ significantly lower SBP and DBP compared to omnivores,³⁶ but not vegetarians;^{32,38} superior lipid profiles to both vegetarians³² and omnivores;^{36,38} and significantly lower BMI compared to omnivores.^{11,36,51,55,57} In the UK, the evidence was still strong but not as extensive as in the US. In UK studies, compared to omnivores, vegans had reduced risk of “any” cancer when data were aggregated with US data;¹⁶ decreased TC and LDL-C;^{29,40,45,48,49} and significantly lower BMI.^{48,53,56,60} In the UK, there was evidence for BMI mediating the relationship between vegan diet choice and reduced incidence of NIDDM.¹³ The evidence for lower blood pressure in vegans versus omnivores was mixed.^{29,33,34} However, in Asian countries, there was less evidence for the health advantages of a vegan diet.^{43,65} In India, the proportion of overweight and obese vegans did not significantly differ from the proportion of overweight and obese omnivores and pescatarians, but were significantly lower than the corresponding percentages of lacto-vegetarians.¹⁴ Of interest, compared to vegans, Indian lacto-vegetarians also had a lower risk of NIDDM, despite having a higher risk of overweight and obesity. In Taiwan, there was a lower risk of high SBP and high DBP in vegans compared to omnivores,³⁹ although in a small study vegan nuns had higher DBP than omnivore nuns.³⁷ However, lipid profiles for vegans in Taiwan did not compare as favorably to their omnivore counterparts with most studies showing significantly worse HDL-C^{39,42–44} and triglycerides^{43,44} for Taiwanese vegans with only a few comparisons showing an advantage for TC^{37,43,44} and LDL-C^{39,43} while others reported non-significant differences.^{43,44} In one Taiwanese study, vegans had a significantly reduced risk of overweight.³⁹

However, in one study from Vietnam,⁶⁹ there was no significant difference in BMIs between vegan and omnivore nuns although the sample size was relatively small.

As to why these discrepancies in results exist, there are several possible explanations. We screened in more studies from the US and UK, so the relative dearth of evidence in Asian countries may be due, in part, to less research that was screened into our review or that may not have been performed at all. Other possibilities include regional differences in cuisine (the typical foods consumed on both vegan and omnivore diets), and international disparities in obesity rates, socio-economic statuses or other religious and/or cultural factors. Regardless of the reason, the evidence for the benefits of a vegan diet was strongest in the US, less strong in the UK, and weakest in Asian countries.

Other important causal factors related to how the reviewed studies were conducted may have affected results. The healthfulness of the vegan diet can make a difference in outcome for both cardiovascular disease⁷⁰ and NIDDM.⁷¹ For free-living vegans, the reason for choosing the diet (eg, health, concern for animals) may be associated with diet quality.⁷² Individuals citing health as their primary reason for adopting a vegan diet reported consuming fewer sweets and more fruit whereas those citing concern for animals as the primary reason reported greater intake of soy, foods containing vitamin D and vitamin supplements (D and B₁₂). Geographic location is also an important consideration as vegans in the US (Adventist 2) reported consuming more fiber and vitamin C than their British counterparts (EPIC-Oxford).¹⁹ These results suggest that evaluating how the content of vegan diets affects outcomes is an important area for further research.

An important mediator that affected outcomes was BMI. In several studies, results were reported both with and without BMI as a covariate. In our Tables, we only listed results that were adjusted for BMI. However, in studies of NIDDM and cancer, inclusion of BMI as a covariate not only made a difference in the results of individual studies, but also affected the aggregate research that was reviewed. Adjustment for BMI in studies of NIDDM affected the overall pattern of results in that US vegans were no longer at reduced risk of NIDDM compared to vegetarians and pescatarians.^{11,12} However, even with adjustment for BMI, vegans’ risk of NIDDM was less than that of omnivores, suggesting that following a vegan diet did affect diabetes risk independent of BMI in the US. In the UK, adopting a vegan versus omnivore diet significantly reduced the risk of NIDDM only when the effects

of BMI were not covaried.¹³ For cancers, in each of the two studies that were reviewed, adjustment for BMI in analyses shifted the pattern of the results such that reduced risk of any cancer for vegans compared to omnivores was no longer significant.^{15,16} However, a meta-analysis that statistically aggregated that data showed, that even with the inclusion of BMI, there was a reduced risk of any cancer in vegans.⁶² Aggregating the data functioned to increase the sample size and consequently shrink the confidence interval so that the analysis was significant.

Dietary adherence is not always measured in observational studies, so the extent to which vegans adhere to their diets is largely unknown. Self-identified vegans (127 for ethical reasons and 80 for health reasons) reported mean numbers of diet violations since beginning their diets, 3.22 ± 6 for ethical vegans and 16 ± 63.8 for health vegans.⁷³ As statistics for duration on their respective diets were not reported, adherence was difficult to assess. Nonetheless, low violation estimates suggest an overall high adherence rate. Regardless, the extent to which adherence is important to health outcomes is unclear. One study compared highly restrictive to more moderate vegans and with one exception (TC) reported non-significant differences (in blood pressure, BMI and some lipid measures³⁵). However, other research suggests that level of adherence to a plant-based diet can make a difference in lowering insulin resistance and in risk of both diagnosis of pre-diabetes and NIDDM.⁷⁴

One final consideration that can affect the outcome is the dietary group that was selected for comparison to vegans. In a tally of results listed in our tables, the majority of the significant findings were obtained in comparisons of vegans to omnivores (47.2% of total vegan-omnivore comparisons). There were fewer significant findings when comparing vegans to vegetarians (20% of total vegan-vegetarian comparisons), pescatarians (12.5% of total vegan-pescatarian comparisons) and semi-vegetarians (2.5% of total vegan-semi-vegetarian comparisons). This is not surprising considering that an omnivore diet is more discrepant from a vegan diet than either vegetarian or pescatarian diets. The relatively small number of favorable comparisons for vegan versus semi-vegetarian diets was largely driven by the large number of null findings comparing these two groups on measures of mortality. Regardless, this survey of findings is consistent with the idea that there is more research support for the benefits of a vegan diet when it is compared to an omnivore diet (as opposed to other vegetarian/pescatarian diets).

Several limitations to this review should be acknowledged. First, although adjustment for potential confounders

was one criterion for judging a study's inclusion, there may have been other confounders that were not addressed (eg, medication usage). Second, several of our included studies reported multiple outcomes. Consequently, the conclusions for some of the outcomes were based, at least in part, on data from the same participants. Third, to evaluate research in the aggregate and facilitate comparisons, dietary groups with different names but similar definitions were given uniform names (eg, meat-eaters were called omnivores). There may have been small differences across studies in how these groups were defined, although there is inevitably some diversity in food choices among people following common diets. Fourth, because these studies investigated free-living vegans, the amount of time that individuals practiced the diet was not uniform and varied both within and across studies. Fifth, although an attempt was made to cover diverse outcomes in this review; there were some areas that were omitted. For example, we did not review studies of mental health outcomes and we excluded studies of children on vegan diets. Sixth, because the health benefits of a vegan diet is a broad topic, we were limited in the extent to which we could cover each medical condition or nutritional status. Seventh, because we used scores on the NOS scales to evaluate study quality, there may have been some studies that were excluded, not because they were poorly executed but because they did not report sufficient detail to score high enough to be screened into the review.

Conclusion

To summarize, there was strong evidence that free-living vegans were less likely than omnivores to develop NIDDM (in the US). Vegans had reduced TC and LDL-C levels, and lower BMIs (in non-Asian locales) compared to omnivores, and in some cases, other dietary groups.

There was evidence that when the effects of BMI were not covaried or when results were aggregated across studies, vegans were less likely than omnivores to develop cancer (all cause). Similarly, there was evidence that BMI mediated the relationship between vegan vs omnivore diet status and development of NIDDM in the UK and between vegan vs vegetarian and pescatarian diet status and development of NIDDM in the US.

There was inconsistent evidence that a vegan diet reduced risk of diagnosis of female cancers and improved mortality rates. There was also mixed evidence that vegans had lower triglycerides and blood pressure, and healthier microbiota.

Finally, we found no evidence that vegans were at reduced risk of developing most specific cancers. Further, in Indian vegans, there was not a reduced probability of overweight,

obesity, or NIDDM. Moreover, the preponderance of evidence showed lower HDL-C in vegans compared to omnivores or no significant differences.

Evidence supporting the health benefits of a vegan diet was strongest in the US and weaker in Asian countries. As the number of vegans increases worldwide, there is a need for additional studies of the diet's protective and therapeutic effects, the pathways through which it exerts them, and the role of moderators and other causal factors in how they are manifested.

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