

Critical Reviews in Food Science and Nutrition



ISSN: 1040-8398 (Print) 1549-7852 (Online) Journal homepage: http://www.tandfonline.com/loi/bfsn20

Effect of nuts on energy intake, hunger, and fullness, a systematic review and meta-analysis of randomized clinical trials

Masoumeh Akhlaghi, Saeed Ghobadi, Morteza Zare & Sahar Foshati

To cite this article: Masoumeh Akhlaghi, Saeed Ghobadi, Morteza Zare & Sahar Foshati (2018): Effect of nuts on energy intake, hunger, and fullness, a systematic review and meta-analysis of randomized clinical trials, Critical Reviews in Food Science and Nutrition, DOI: 10.1080/10408398.2018.1514486

To link to this article: https://doi.org/10.1080/10408398.2018.1514486

	Published online: 13 Nov 2018.
	Submit your article to this journal 🗷
lılı	Article views: 5
CrossMark	View Crossmark data 🗗



REVIEW



Effect of nuts on energy intake, hunger, and fullness, a systematic review and meta-analysis of randomized clinical trials

Masoumeh Akhlaghi^a, Saeed Ghobadi^b, Morteza Zare^c, and Sahar Foshati^d

^aNutrition Research Center, School of Nutrition and Food Sciences, Shiraz University of Medical Sciences, Shiraz, Iran; ^bStudent Research Committee, School of Nutrition and Food Sciences, Shiraz University of Medical Sciences, Shiraz, Iran; ^cNeuroscience Research Center, Institute of Neuropharmacology, Kerman University of Medical Sciences, Kerman, Iran; ^dDepartment of Clinical Nutrition School of Nutrition and Food Sciences, Shiraz University of Medical Sciences, Shiraz, Iran

ABSTRACT

Despite high energy content, nut consumption has not been associated with weight gain in epidemiological and clinical investigations. Since a satiety effect a decreased sense of hunger in the following meals has been proposed as the mechanism of nuts against obesity, the present metaanalysis was performed to examine how nut consumption affects "energy intake", "hunger", and "fullness". "Weight" was not included among the search terms but weight data were extracted from the collected articles in order to help interpreting the results. A total of 1048 trials were found, of which 31 passed the eligibility process. Daily energy intake was mostly assessed by 3-day food records, and hunger and fullness were estimated by visual analog scale (VAS). Nut consumption was associated with increased energy intake (mean difference (MD) (nuts - control) = 76.3 kcal; 95% Cl: 22.7, 130 kcal; P = 0.005). In contrast, hunger suppressed following nut consumption (MD = $-6.54 \,\mathrm{mm}$ VAS; 95% CI: -12.7, $-0.42 \,\mathrm{mm}$ VAS; P = 0.03). Nuts did not affect sense of fullness and weight in the overall estimate. Subgroup analysis based on participants' BMI revealed that energy intake of overweight/obese individuals was increased following nut consumption while such effect was not observed in normal weight participants. In conclusion, pooled estimates of available clinical trials showed increased energy intake following nut consumption in overweight/obese individuals but not in persons with normal weight. Nut consumption was associated with decreased hunger but no effect was observed on fullness and weight.

KEYWORDS

Nuts; energy intake; hunger; satiety; fullness; weight; obesity

Introduction

Nuts are highly nutritious, containing ample amounts of protein, unsaturated fatty acids, vitamins, minerals, and phytochemicals (Bolling, McKay, and Blumberg 2010). Thanks to this composition, nuts have health benefits for prevention of diseases, especially metabolic syndrome and cardiovascular diseases (Ros 2015). On the other hand, due to high fat content, nuts are energy-dense; and thus, assumed to increase the risk of weight gain and obesity. However, crosssectional studies have shown an inverse association between nut consumption and the prevalence of general and abdominal obesity and metabolic syndrome (Ibarrola-Jurado et al. 2013). Likewise, prospective cohorts have found lower risk of weight gain and obesity in individuals with higher nut consumption (Bes-Rastrollo et al. 2009). A recent meta-analysis of prospective cohorts and clinical trials also showed decreased risk of overweight/obesity and lower measures of weight, body mass index, and waist circumference in higher nut consumptions (Li et al. 2018).

Based on evidence from epidemiological and interventional studies, a number of mechanisms have been suggested

for the inverse association of nut consumption and body weight (Tan, Dhillon, and Mattes 2014). One of these mechanisms is a satiety effect which is proposed to occur following nut consumption and may extend until the next meal or meals (Tan, Dhillon, and Mattes 2014). According to this hypothesis, the satiety effect postpones the sense of hunger and enables the person to compensate for extra energy acquired from nuts by decreasing food ingestion over the next meals, something that is called energy compensation and may help in weight control (Tan and Mattes 2013).

Based on the aforementioned hypothesis, a number of clinical trials have examined the effect of nuts on hunger and satiety but the results are rather complex and contradictory (e.g., Tan and Mattes 2013; Hull et al. 2015; Kirkmeyer and Mattes 2000; Burton-Freeman 2005). Since there is no conclusive evidence on this subject, we performed a meta-analysis on available randomized clinical trials to see how nut consumption affects daily energy intake and hunger. Because previous meta-analyses have investigated weight effects of nuts (Li et al. 2018; Flores-Mateo et al. 2013) we did not include "weight" among our search terms but extracted weight data

from the collected articles in order to help us better interpret the results.

Methods

A systematic review and meta-analysis was conducted in accordance with the guidelines of Preferred Reporting Items for Systematic Reviews and Meta-Analyses at all stages of design, implementation, and reporting (PRISMA-P Group 2015).

Search

PubMed, Scopus, and EMBASE databases were searched from the earliest online available date through August 2018 to find randomized clinical trials examining the effect of nuts on daily energy intake, hunger, satiety, and fullness in adults aged >18 years. Search terms were nuts, walnut, almond, peanut, pecan, hazelnut, filbert, pistachio, macadamia, cashew, Brazil nut, pine nut, chestnut, energy, food intake, appetite, satiety, satiation, hunger, and fullness (search strings are provided as supplementary material). The search was performed by one investigator and screening titles and abstracts was performed by two independent investigators. Final decision was made through discussion with a third investigator. There was no restriction on language. The meta-analysis was not pre-registered in any database.

Eligibility criteria

Randomized clinical trials, in either parallel or crossover design, investigating the effect of nuts on daily energy intake, hunger, and fullness were included. Trials on satiety were not included in this report because a few studies (n = 5) assessed the effect of nuts on satiety. Trials had to be randomized, but the blindness was not a requirement because in majority of the studies nuts were administered in their regular form. Trials were included if they provided similar conditions in the treatment and control groups except for nut consumption, executed control over diet and physical activity of participants, and administered certain amount of nuts to the treatment group. It is worth noting that, because of technical difficulty in preparation of an identical control food for nuts, administration of a control food/load to the control group was not necessary but the presence of the control group itself was. Studies were excluded: 1) if study participants were involved in pathologic conditions affecting either energy intake or appetite, such as malnutrition, type 2 diabetes, cancer, and organ failure; 2) if an isocaloric and nutritionally comparable control was not applied (Food and Drug Administration, XXXX). For instance, in some trials, different doses of nuts were administered in different arms but no control group (i.e. non-effective compound) was used. Also, in some studies regular nuts were compared with high-oleic nuts or in some trials shelled nuts were compared with in-shell nuts. Such studies were excluded; 3) if nuts and control strategies were not isocaloric; 4) if diets of control and nut groups were not controlled for dietary nut consumption; 5) if nuts were not

given in similar quantities to all participants in nut-treated group; 6) if a suitable technique for evaluating hunger and fullness (e.g., measuring the amount of foods consumed or using a subjective tool, such as visual analog scale (VAS)) was not applied; 7) if nut ingredients, such as nut oil, were examined; 8) if specific types of nuts, for instance high oleic peanuts, was used; 9) if nuts were mixed with other foods, such as raisins, and so the observed effects could not be solely attributed to nuts; 10) in crossover designs, if a washout period was not established between the two sequences or if all subjects did not experience both control and nut conditions; 11) if sufficient information (i.e. the mean and standard deviation or error) for parameters of interest (daily energy intake, hunger, fullness, and weight) was not provided; 12) if baseline values were not reported; and 13) in repeated publications, where only one report with the most relevant design to the aims of this meta-analysis was used. Scores of the study quality (Jadad scale) were not reported because of difficulty in blindness of the studies.

Risk of bias assessment

Risk of bias was determined according to a recent systematic review on appetite and energy intake (Halford et al. 2018). Risk of bias was rated as low or high for each of 4 issues: 1) power calculation and meeting the calculated sample size; 2) results reported from 100% sample size or the use of intention-to-treat analysis; 3) the percentage of dropouts; and 4) complete report of outcomes.

Statistical analysis

Pooled effects were calculated by estimating mean difference (MD) and standard error of energy intake, hunger, fullness, and weight between control and nut-treated groups using the inverse-variance random-effect model. Heterogeneity was assessed by I² statistics (Higgins and Green, 2011). Subgroup analysis was performed for each of the investigated variables (energy, hunger, fullness, weight) based on subjects' weight status (normal (BMI < 25 kg/m²) vs. overweight/obese (BMI \geq 25 kg/m²)(. Subgroup analysis was also performed based on nut type (walnut, almond, hazelnut, and peanut) for energy intake, but due to insufficient number of trials this subgroup analysis was not performed for other parameters. STATA software version 12.0 (StataCorp, USA) was used for data analysis.

Results

Search results

After searching PubMed, Scopus, and EMBASE databases, 1048 articles were identified; 305 were duplicates and put aside, 672 were excluded following screening titles and abstracts, 71 were assessed for eligibility, and 31 trials were entered in the meta-analysis (Figure 1). Non-English articles entered the screening process through reading their English

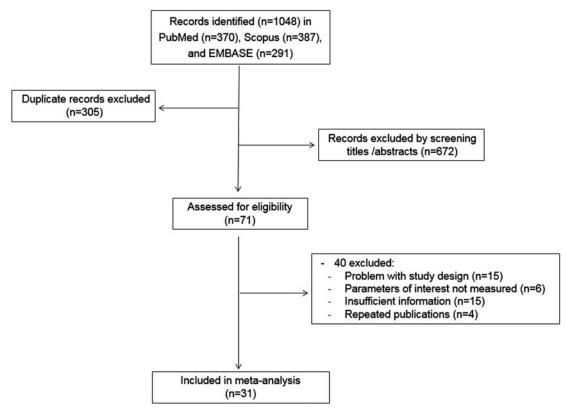


Figure 1. Summary of the screening and selection process of trials included in the meta-analysis of the effect of nuts on energy intake, hunger, and fullness.

abstract. None of the trials written in other languages passed the eligibility process.

Table 1 presents a summary of the included trials. Twenty three, 14, 10, and 15 trials assessed daily energy intake, hunger, fullness, and weight, respectively. Daily energy intake was mostly assessed by 3- to 7-day food records, and hunger and fullness were estimated by visual analog scale (expressed as mm). Three studies were cited twice because they had used different doses (Hull et al. 2015; Tey et al. 2013) or different types of nuts (Kirkmeyer and Mattes, 2000).

A total of 1467 participants were included in the metaanalysis. They were either healthy or with relatively mild medical conditions like overweight, obesity, hyperlipidemia, metabolic syndrome, and prediabetes. The participants had a wide variety of BMI, including normal (< 25 kg/m²), overweight $(25-29.9 \text{ kg/m}^2)$, and obese $(\geq 30 \text{ kg/m}^2)$. Nuts included almond, walnut, peanut, hazelnut, and pecan in doses ranging from 20.0 g/day to 87.5 g/day. There was large variability in the type of control food, varying from no control food (i.e. control group just followed their usual diet instead of getting a control food plus usual diet) to receiving snacks, cereals, or fruits as the control food.

Effect of nuts on daily energy intake

As stated, 23 trials investigated the effect of nuts on daily energy intake. An overall significant increasing effect of nuts on energy intake (MD (nuts - control) = 76.3 kcal; 95% CI: 22.7, 130 kcal; P = 0.005) was observed (Figure 2). Subgroup

analysis based on participants' BMI showed significant increased energy intake following consumption of nuts in overweight/obese individuals (MD = 125 kcal; 95% CI: 70.8, 180 kcal; P < 0.001) but not in individuals with normal weight (MD = -27.5 kcal; 95% CI: -10.5, 50.5 kcal; P = 0.49) (Figure 2-A). Subgroup analysis of energy intake based on nuts type showed that walnut, almond, and hazelnut consumption was associated with higher energy intake although the association was statistically significant only for hazelnut (MD = 227 kcal; 95% CI: 25.6, 427.6 kcal; P = 0.03) (Figure 2-B). There was high level of heterogeneity in the overall effect ($I^2 = 82.1\%$) and walnut ($I^2 = 92.8\%$) and almond $(I^2 = 71.3\%)$ subgroups.

Effect of nuts on hunger

Pooled estimate of 14 trials showed suppressed sense of hunger following nut consumption (MD = -6.54 mm VAS; 95% CI: -12.7, -0.42 mm VAS; P = 0.03) (Figure 3). Subgroup analysis based on participants' BMI did not show a significant association between nut consumption and BMI. There was high heterogeneity in the overall effect and the BMI subgroups ($I^2 = 78.2\% - 97.7\%$).

Effect of nuts on sense of fullness

Ten studies reported the effect of nuts on sense of fullness (MD = 0.03 mm VAS; 95% CI: -12.2, 12.3 mm VAS; P = 1)(Figure 4). Nuts did not seem to affect fullness in the overall effect, but in subgroup analysis normal individuals showed

Table 1. Characteristics of the clinical trials included in the meta-analysis of the effect of nuts on energy intake, hunger, and satiety.

First author, year (ref)	Subjects	Subjects number	Age bolling, McKay, and Blumberg 2010 (y)	McKay, and Blumberg 2010 (kg/m²)	Study design	Intervention	Nut dose	Control	Outcome
Chon at al 2015	010010 120420 120000	AE	20+012	20.7	, , , , , ,	pacaly	≈ 30 21		_
Chell et al. 2015 Dhillon. Tan. and	Overweight/obese	50	34.3 + 13.0	30.5 + 3.5	Parallel	Almond	63 g 15% energy intake	1 1	E H F W*
Mattes 2016		}					(6:		
Hull et al. 2015	Healthy	32	48.4 ± 1.0	22.7 ± 0.26	Crossover	Almond	28 g	ı	*, *H
Hull et al. 2015	Healthy	32	48.4 ± 1.0	22.7 ± 0.26	Crossover	Almond	42 g	ı	** *H
Jenkins et al. 2008	Hyperlipidemic	27	64.0 ± 9	25.5 ± 4	Crossover	Almond	73 g	Muffin	E, W
Jung et al. 2017	Overweight	84	52.4 ± 0.6	25.4 ± 0.22	Crossover	Almond	56 g	Cookie	*ш
Kirkmeyer and	Healthy	24	22 ± 2.5	Normal	Crossover	Almond	80.4	Chestnut	E*, H*
Mattes, 2000									
Liu et al. 2018	Healthy	85	26.55 ± 5.3	22.2 ± 3.12	Parallel	Almond	56 g	ı	E, W*
Sayer et al. 2017	Overweight/obese	22	35.0 ± 3.0	30.0 ± 1.0	Crossover	Almond	28 g	Baked food	ж. *Н
Tan and Matttes, 2013	Overweight/obese	55	30.8 ± 10.5	27.6 ± 4.6	Parallel	Almond	43 g	ı	E*, H*, F*, W*
Wien et al. 2010	Prediabetes	92	53.5 ± 10.0	29.5 ± 5.0	Parallel	Almond	56 g	ı	E, W*
Zaveri and	Overweight/obese	32	37.7 ± 6.6	30.1 ± 3.2	Parallel	Almond	56 g	Cereal bar	E*, H*, W*
Drummond, 2009									
Agebratt et al. 2016	Healthy	30	23.5 ± 3.7	22.3 ± 1.9	Parallel	Walnut	7 kcal/kg	Apple/pear	E*, W*
Burton-Freeman, 2005	Healthy	25	31.5 ± 3	23.0 ± 2	Crossover	Walnut	20 g	Strawberry	E*, H*, F*
								snake	
Casas-Agustench et al. 2009	Healthy	29	22 ± 4.0	24.1 ± 4.5	Crossover	Walnut	54 g	Olive oil	* ±
Holt et al. 2015	Hypercholesterolemic	30	60.0 ± 4.7	24.6 ± 3.4	Parallel	Walnut	35 a	ı	ш
Neale et al. 2017	Overweight/obese	170	43.0 + 8.4	32.6+4.3	Parallel	Walnut	30 g	1	* M*
Rock et al. 2017	Overweight/obese	100	52.8 ± 1.5	32.4 ± 0.4	Parallel	Walnut	15% energy intake	ı	* X
Morgan and	Healthy	19	41.0 ± 11.0	24.0 ± 4.5	Parallel	Pecan	68 g	1	, ш
Clayshulte, 2000							n		
Casas-Agustench	Metabolic syndrome	20	51.8 ± 8.4	30.8 ± 3.1	Parallel	Walnut/	30 g	ı	E*, W*
et al. 2011						almond/ hazelnut			
B250117 of 10 7014	Ozodol+deiomeoro	70	1 + 19	205 + 2	1010101	Doznit	201 cm of 2 62 fram 200	Dotato crico	* *
Claesson et al. 2014	Overweight/obese Healthy	75	01 ± 1	30.5±5 777+17	Parallel	Peanut	84 g males/ 30 g remales 20 kgal/kg	rotato crisp Candy	*M *4
Alves et al. 2014	Overweight/obese	ł 4	27.7 ± 2.2	29.6 ± 2.4	Parallel	Peanut	56 d	-	H, F, W*
Moreira Alves et al. 2014	Overweight/obese	4	27.7 ± 2.2	29.6 ± 2.4	Parallel	Peanut	56 g	ı	. Ш
Duarte Moreira Alves	Overweight/obese	47	27.1 ± 0.7	29.8 ± 2.4	Parallel	Peanut	56 g	Biscuit	*± ′*±
et al. 2014	;;;;;;;;;;;;;;;;;;;;;;;;;;;;;;;;;;;;;;	7	40 5 + 1 6	0.00	امااديدا	+11200	, cc		*//\
JOHNSON Et al. 2013	nealth	‡ ;	0.1 H C.04	9.1.0 1.0 ± 0.9	ratallel	realini	25 g	Cidilis	^ ` →
Kirkmeyer and Mattes, 2000b	Healthy	74	22 ± 2.5	Normal	Crossover	Peanut	8/.5	Chestnut	., H +, H
Reis et al. 2011	Healthy	13	28.5 ± 10.0	22.7 ± 2.5	Crossover	Peanut	63 q	Cheese	*ш
							n	sandwich	
Pearson et al. 2017	Healthy	52	38.6 ± 14.1	23.9 ± 3.1	Parallel	Hazelnut	42 g	Potato crisp	*ш
Tey et al. 2013	Overweight/obese	70	42.5 ± 13.3	30.6 ± 4.6	Parallel	Hazelnut	30 g	ı	E, W*
Tey et al. 2013	Overweight/obese	74	42.0 ± 11.9	30.7 ± 5.3	Parallel	Hazelnut	60 g	1	E, W*
O + 2000 000 2001/01/1									٠

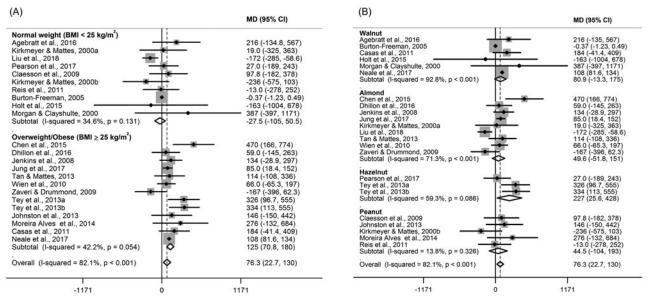


Figure 2. Forest plot of clinical trials examining the effect of nuts on energy intake with subgroup analysis based on: A) participants' BMI; B) nut type. Values are the mean difference (MD) between nuts and control groups with 95% Cls.

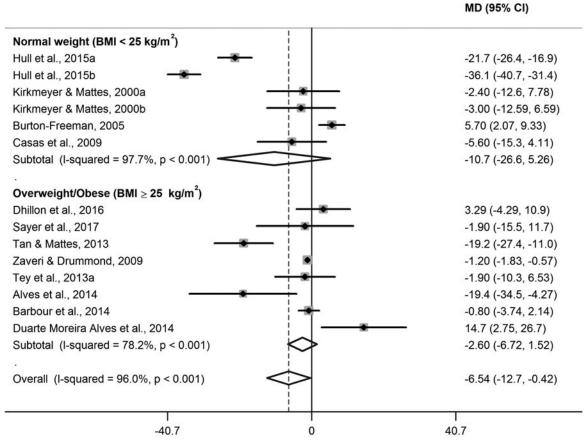


Figure 3. Forest plot of clinical trials examining the effect of nuts on hunger with subgroup analysis based on participants' BMI. Values are the mean difference (MD) between nuts and control groups with 95% Cls.

non-significant increased sense of fullness (MD = 17.9 mm VAS; 95% CI: -2.78, 38.6 mm VAS; P = 0.09) and overweight/ obese individuals indicated non-significant decreased fullness (MD = $-8.43 \,\text{mm}$ VAS; 95% CI: -18.8, 1.91 mm VAS; P = 0.11) following nut consumption. High heterogeneity was observed in the overall effect and the BMI subgroups ($I^2 = 85\% - 99.7\%$).

Effect of nuts on weight

Assessment of weight was not among the primary aims of this work. Therefore, weight was not included in the search terms, but it assessed in studies which were included in the analysis based on the main outcomes. Fifteen out of 31 trials reported weight. These trials did not show any effect from

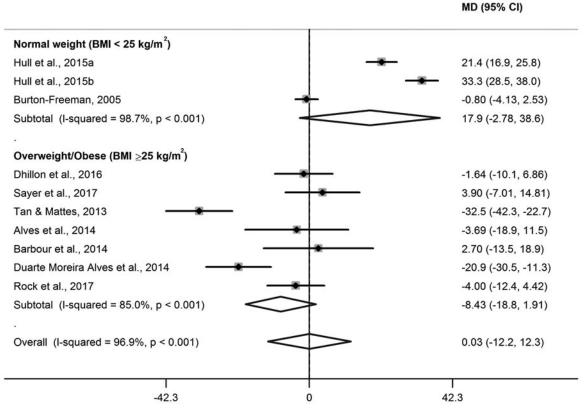


Figure 4. Forest plot of clinical trials examining the effect of nuts on sense of fullness with subgroup analysis based on participants' BMI. Values are the mean difference (MD) between nuts and control groups with 95% Cls.

nuts on weight (MD = -0.09 kg; 95% CI: -0.59, 0.41 kg;P = 0.72) (Figure 5). The overweight/obese subgroup indicated the same result as the overall effect but trials on normal weight individuals revealed a non-significant inverse association between nut consumption and weight -1.19, 0.15 kg; P = 0.13). (MD = -0.52 kg; 95%)CI: Heterogeneity in normal weight subgroup was low $(I^2 = 0)$ but a moderate heterogeneity was observed among the whole trials and in overweight/obese subgroup ($I^2 = 59.7\%-68.0\%$).

Risk of bias assessment

Risk of bias assessment is reported in Supplementary Table 1. The most frequent risks of bias were due to absence/not reporting power calculation and analysis on less than 100% of sample size or absence of intention-to-treat analysis.

Discussion

Concerning the current hypothesis on the beneficial effect of nuts on weight and their satiety effect as one of the mechanisms by which nuts exert such effect, we conducted this meta-analysis to investigate randomized clinical trials examining the effect of nuts on hunger, satiety, and energy intake. According to results of this meta-analysis, consumption of nuts increases daily energy intake and suppresses hunger but it has no effect on sense of fullness or weight. Subgroup analysis based on BMI suggested that weight

status may influence the effect of nuts on energy intake and sense of fullness.

Nuts are classified as high energy density foods, producing >4 kcal energy per gram (Brufau, Boatella, and Rafecas 2006). This high energy density is due to nuts' high fat content (from about half to as much as 75% of weight). Despite high energy content, consumption of nuts has not been accompanied with obesity in epidemiological and clinical investigations (Ibarrola-Jurado et al. 2013; Li et al. 2018). The absence of congruence between energy content and weight effects of nuts has been proposed to be, at least partly, due to a satiety effect that is generated after nut consumption, extends for hours, and influences food consumption over the next meals (Tan, Dhillon, and Mattes 2014). This satiety effect is proposed to either decrease daily energy intake or at least keep it unchanged (Halford et al. 2018).

However, results of this meta-analysis showed a positive association between ingestion of nuts and energy intake; although the magnitude of the increased energy intake was lower than that obtained from the consumed nuts. The average nut consumption in trials of energy was 50.9 g/day. Based on food composition tables, this amount of nuts should provide about 300 kcal/day while the quantity of energy intake estimated from trials was 76.3 kcal/day. There may be several explanations for this inequality between calculated and acquired energy from nuts. One explanation is incomplete digestion of nuts in which a number of cell walls in nuts remain intact during digestion process and pass undigested through gastrointestinal tract (Mandalari et al. 2014). In vitro experiments showed that even prolonged

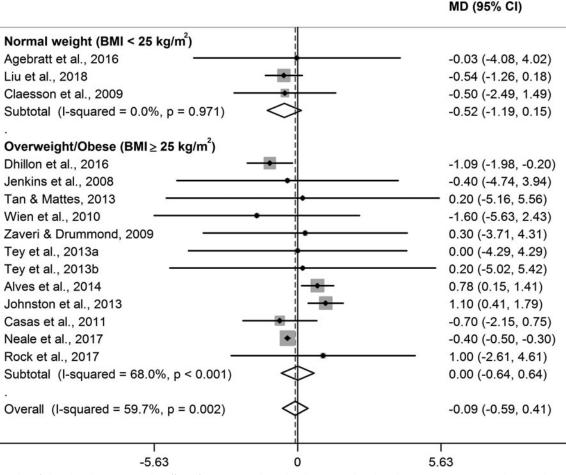


Figure 5. Forest plot of clinical trials examining the effect of nuts on weight with subgroup analysis based on participants' BMI. Values are the mean difference (MD) between nuts and control groups with 95% Cls.

digestion periods are unable to release encapsulated lipids from the intact cells (Grundy et al. 2016). In some cases, slow penetration of lipase into the cell walls may lead to the release of fat in the ileum or colon where sites of fat absorption have already passed (Grundy et al. 2015). Second explanation is that nuts may increase energy expenditure. Nuts, including peanuts (Alper and Mattes, 2002) and walnuts (Tapsell et al. 2009), have shown potential to increase resting energy expenditure (Alper and Mattes, 2002) and body fat oxidation (Tapsell et al. 2009). Mono- (Casas-Agustench et al. 2009) and poly- (Casas-Agustench et al. 2009; Piers et al. 2002) unsaturated fatty acids have shown to augment postprandial oxidation. In this regard, animal studies reported higher sympathetic activity following consumption of safflower oil in comparison with beef tallow diet (Matsuo et al. 1995). Also, safflower-enriched diets increased the expression of peroxisome proliferator-activated receptor-α gene (PPARA) which is involved in fatty acid oxidation (Zhang et al. 2010). In addition, polyunsaturated fatty acids are suggested to upregulate uncoupling protein genes, genes implicated in proton leak across mitochondrial inner membrane and dissipation of energy in the form of heat instead of ATP (Armstrong and Towle, 2001).

Due to paucity of data, we did not report data on satiety, but trials on hunger indicated an inverse association between nut consumption and sense of hunger. This finding

is in line with previously proposed hypothesis on the satiety effect of nuts if we note that hunger is the opposite sense of satiety (de Graaf, 2011). Interestingly, nut consumption appeared to be ineffective on the sense of fullness in the overall effect estimate which could be because nuts are energy-dense, are ingested in small quantities, and thus do not occupy much of the stomach volume. Mechanisms of the satiety-promoting effect of nuts are not completely understood but it is suggested that specific hormones, such as cholecystokinin, peptide YY, and glucagon-like peptide 1, may be involved (Reis et al. 2013; Pasman et al. 2008). Fatty acid composition of nuts, in particular polyunsaturated fatty acids, may be effective in induction of nuts satiety effect through stimulation of satiety-related hormones (Stevenson, Clevenger, and Cooper 2015; Kozimor, Chang, and Cooper 2013; Maljaars et al. 2009).

Subgroup analysis revealed that weight status may influence the effect of nuts on the assessed variables. Compared to normal weight individuals, people with overweight/obese showed trends towards less suppressed hunger (MD = -2.60 vs. -10.66, mm VAS), less sense of fullness (MD = -8.43 vs. 17.9, mm VAS), higher weight (MD = 00 vs. -0.52, kg) and increased energy intake (MD = 125 vs. -27.5, kcal/day) after nut consumption. These weight-dependent effects of nuts may be explained by the fact that obese people have altered appetite sensitivity and regulation. Compared to people with

normal weight, overweight/obese individuals have poorer ability to perceive signals of hunger and satiety (Herbert and Pollatos, 2014). Obese persons are more sensitive to appetitive stimulations of ghrelin (Druce et al. 2005). Additionally, a large cross-sectional study showed that genetic susceptibility to obesity is associated with reduced satiety responsiveness (Llewellyn et al. 2014). Unfortunately, no study has yet compared nut effect on satiety and weight in overweight/obese vs. normal weight individuals. Also, studies on the effect of nuts on hunger and satiety are limited and the findings are still inconclusive. More studies need to focus on these areas.

Strengths and limitations

There were limitations with this work. We did not report data on satiety because few studies assessed it. Furthermore, methods of assessing energy intake, such as food record or recall, are not quite reliable, especially obese individuals may underreport their food intake. In addition, application of different methods of dietary assessment can increase risk of bias. Time of nut consumption might affect hunger, fullness, and energy intake, but subgroup analysis based on time was not possible because in some studies the time of nut administration was not specified and participants were free to eat nuts as a snack or with meals. However, this meta-analysis was the first that investigated the effect of nuts on energy intake, hunger, and fullness.

Conclusions

In conclusion, pooled estimates of available clinical trials showed increased energy intake following nut consumption in overweight/obese individuals but not in normal weight ones. In the overall effect, nut consumption was associated with decreased hunger but no effect was observed on fullness and weight. Person's body weight may influence the effect of nuts on fullness and weight but results were not significant in this regard. More trials are needed to confirm these results.

Acknowledgment

The authors wish to thank Mr. H. Argasi at the Research Consultation Center (RCC) of Shiraz University of Medical Sciences for his invaluable assistance in editing this manuscript. All authors have read and approved the final manuscript.

Disclosure statement

MA, SG, MZ, and SF, no conflicts of interest.

Funding

The authors received no funding for this study.

Abbreviations

BMI body mass index

confidence interval mean difference VAS visual analog scale

References

Agebratt, C., E. Ström, T. Romu, O. Dahlqvist-Leinhard, M. Borga, P. Leandersson, and F. H. Nystrom. 2016. A randomized study of the effects of additional fruit and nuts consumption on hepatic fat content, cardiovascular risk factors and basal metabolic rate. PLoS One 11(1):e0147149.

Alper, C. M., and R. D. Mattes. 2002. Effects of chronic peanut consumption on energy balance and hedonics. International Journal of Obesity and Related Metabolic Disorders: Journal of the International Association for the Study of Obesity 26(8):1129-37.

Alves, R. D., A. P. Moreira, V. S. Macedo, R. de Cássia Gonçalves Alfenas, J. Bressan, R. Mattes, and N. M. Costa. 2014. Regular intake of high-oleic peanuts improves fat oxidation and body composition in overweight/obese men pursuing a energy-restricted diet. Obesity (Silver Spring) 22(6):1422-9.

Armstrong, M. B., and H. C. Towle. 2001. Polyunsaturated fatty acids stimulate hepatic UCP-2 expression via a PPAR alpha-mediated pathway. American Journal of Physiology-Endocrinology and Metabolism 281(6):E1197-204.

Barbour, J. A., P. R. Howe, J. D. Buckley, G. C. Wright, J. Bryan, and A. M. Coates. 2014. Lower energy intake following consumption of hi-oleic and regular peanuts compared with iso-energetic consumption of potato crisps. Appetite 82:124-30.

Bes-Rastrollo, M., N. M. Wedick, M. A. Martinez-Gonzalez, T. Y. Li, L. Sampson, and F. B. Hu. 2009. Prospective study of nut consumption, long-term weight change, and obesity risk in women. The American Journal of Clinical Nutrition 89(6):1913-9.

Bolling, B. W., D. L. McKay, and J. B. Blumberg. 2010. The phytochemical composition and antioxidant actions of tree nuts. Asia Pacific Journal of Clinical Nutrition 19(1):117-23.

Brufau, G., J. Boatella, and M. Rafecas. 2006. Nuts: source of energy and macronutrients. The British Journal of Nutrition 96 Suppl 2:

Burton-Freeman, B. 2005. Sex and cognitive dietary restraint influence cholecystokinin release and satiety in response to preloads varying in fatty acid composition and content. The Journal of Nutrition 135(6):1407-14.

Casas-Agustench, P., P. López-Uriarte, M. Bulló, E. Ros, A. Gómez-Flores, and J. Salas-Salvadó. 2009. Acute effects of three high-fat meals with different fat saturations on energy expenditure, substrate oxidation and satiety. Clinical Nutrition 28(1):39-45.

Casas-Agustench, P., P. López-Uriarte, M. Bulló, E. Ros, J. J. Cabré-Vila, and J. Salas-Salvadó. 2011. Effects of one serving of mixed nuts on serum lipids, insulin resistance and inflammatory markers in patients with the metabolic syndrome. Nutrition, Metabolism and Cardiovascular Diseases 21(2):126-35.

Chen, C. Y., M. Holbrook, M. A. Duess, M. M. Dohadwala, N. M. Hamburg, B. F. Asztalos, P. E. Milbury, J. B. Blumberg, and J. A. Vita. 2015. Effect of almond consumption on vascular function in patients with coronary artery disease: a randomized, controlled, cross-over trial. Nutrition Journal 14(1):61.

Claesson, A. L., G. Holm, A. Ernersson, T. Lindström, and F. H. Nystrom. 2009. Two weeks of overfeeding with candy, but not peanuts, increases insulin levels and body weight. Scandinavian Journal of Clinical and Laboratory Investigation 69(5):598-605.

de Graaf, C. 2011. Trustworthy satiety claims are good for science and society. Comment on 'Satiety. No way to slim'. Appetite 57(3): 778-83. discussion 784-90.

Dhillon, J., S. Y. Tan, and R. D. Mattes. 2016. Almond consumption during energy restriction lowers truncal fat and blood pressure in compliant overweight or obese adults. The Journal of Nutrition 146(12):2513-9.

Druce, M. R., A. M. Wren, A. J. Park, J. E. Milton, M. Patterson, G. Frost, M. A. Ghatei, C. Small, and S. R. Bloom. 2005. Ghrelin

- increases food intake in obese as well as lean subjects. International Journal of Obesity (2005) 29(9):1130-6.
- Duarte Moreira Alves, R., A. P. Boroni Moreira, V. Silva Macedo, and N. M. Brunoro Costa. 2014. Gonçalves alfenas rde C, bressan J. High-oleic peanuts increase diet-induced thermogenesis in overweight and obese men. Nutrición Hospitalaria 29(5):1024-32.
- Flores-Mateo, G., D. Rojas-Rueda, J. Basora, E. Ros, and J. Salas-Salvadó. 2013. Nut intake and adiposity: Meta-analysis of clinical trials. The American Journal of Clinical Nutrition 97(6):1346-55.
- Food and Drug Administration. Guidance for Industry: E 10 Choice of control group and related issues in clinical trials. Published in May 2001. http://www.fda.gov/downloads/drugs/guidancecomplianceregulatoryinformation/guidances/ucm073139.pdf.
- Grundy, M. M., F. Carrière, A. R. Mackie, D. A. Gray, P. J. Butterworth, and P. R. Ellis. 2016. The role of plant cell wall encapsulation and porosity in regulating lipolysis during the digestion of almond seeds. Food & Function 7(1):69-78.
- Grundy, M. M., P. J. Wilde, P. J. Butterworth, R. Gray, and P. R. Ellis. 2015. Impact of cell wall encapsulation of almonds on in vitro duodenal lipolysis. Food Chemistry 185:405-12.
- Halford, J.C.G., U. Masic, C. F. M. Marsaux, A. J. Jones, A. Lluch, L. Marciani, M. Mars, S. Vinoy, M. Westerterp-Plantenga, and D.J. Mela. 2018. Systematic review of the evidence for sustained efficacy of dietary interventions for reducing appetite or energy intake. Obesity Reviews Jun 25.
- Herbert, B. M., and O. Pollatos. 2014. Attenuated interoceptive sensitivity in overweight and obese individuals. Eating Behaviors 15(3): 445 - 8.
- Higgins J.P.T., and S. Green (editors). 2011. Cochrane Handbook for Systematic Reviews of Interventions. Version 5.1.0 [updated March 2011]. The Cochrane Collaboration. http://handbook.cochrane.org/ $index.htm \#part_2_general_methods_for_cochrane_reviews.htm$
- Holt, R. R., S. J. Yim, G. C. Shearer, R. M. Hackman, D. Djurica, J. W. Newman, A. W. Shindel, and C. L. Keen. 2015. Effects of short-term walnut consumption on human microvascular function and its relationship to plasma epoxide content. The Journal of Nutritional Biochemistry 26(12):1458-66.
- Hull, S., R. Re, L. Chambers, A. Echaniz, and M. S. Wickham. 2015. A mid-morning snack of almonds generates satiety and appropriate adjustment of subsequent food intake in healthy women. European Journal of Nutrition 54(5):803–10.
- Ibarrola-Jurado, N., M. Bulló, M. Guasch-Ferré, E. Ros, M. A. Martínez-González, D. Corella, M. Fiol, J. Wärnberg, R. Estruch, P. Román., et al. 2013. Cross-sectional assessment of nut consumption and obesity, metabolic syndrome and other cardiometabolic risk factors: the PREDIMED study. PLoS One 8(2):e57367.
- Jenkins, D. J., C. W. Kendall, A. Marchie, A. R. Josse, T. H. Nguyen, D. A. Faulkner, K. G. Lapsley, and J. Blumberg. 2008. Almonds reduce biomarkers of lipid peroxidation in older hyperlipidemic subjects. The Journal of Nutrition 138(5):908-13.
- Johnson, C. S., C. M. Trier, and K. R. Fleming. 2013. The effect of peanut and grain bar preloads on postmeal satiety, glycemia, and weight loss in healthy individuals: an acute and a chronic randomized intervention trial. Nutrition Journal 12(1):35.
- Jung H., C. O. Chen, J. B. Blumberg, and H. K. Kwak. 2018. The effect of almonds on vitamin E status and cardiovascular risk factors in Korean adults: A randomized clinical trial. European Journal of Nutrition. 57(6):2069-79. doi: 10.1007/s00394-017-1480-5.
- Kirkmeyer, S. V., and R. D. Mattes. 2000. Effects of food attributes on hunger and food intake. International Journal of Obesity and Related Metabolic Disorders: Journal of the International Association for the Study of Obesity 24(9):1167-75.
- Kozimor, A., H. Chang, and J. A. Cooper. 2013. Effects of dietary fatty acid composition from a high fat meal on satiety. Appetite 69:39-45.
- Li, H., X. Li, S. Yuan, Y. Jin, and J. Lu. 2018. Nut consumption and risk of metabolic syndrome and overweight/obesity: a Meta-analysis of prospective cohort studies and randomized trials. Nutrition & Metabolism 15(1):46.
- Liu, Y., H. J. Hwang, H. S. Kim, and H. Park. 2018. Time and intervention effects of daily almond intake on the changes of lipid profile

- and body composition among free-living healthy adults. Journal of Medicinal Food 21(4):340-7.
- Llewellyn, C. H., M. Trzaskowski, C. H. M. van Jaarsveld, R. Plomin, and J. Wardle. 2014. Satiety mechanisms in genetic risk of obesity. JAMA Pediatrics 168(4):338-44.
- Maljaars, J., E. A. Romeyn, E. Haddeman, H. P. Peters, and A. A. Masclee. 2009. Effect of fat saturation on satiety, hormone release, and food intake. The American Journal of Clinical Nutrition 89(4): 1019-24.
- Mandalari, G., M. M. Grundy, T. Grassby, M. L. Parker, K. L. Cross, S. Chessa, C. Bisignano, D. Barreca, E. Bellocco, G. Laganà., et al. 2014. The effects of processing and mastication on almond lipid bioaccessibility using novel methods of in vitro digestion modelling and micro-structural analysis. British Journal of Nutrition 112(09): 1521-9. Nov 14
- Matsuo, T., Y. Shimomura, S. Saitoh, K. Tokuyama, H. Takeuchi, and M. Suzuki. 1995. Sympathetic activity is lower in rats fed a beef tallow diet than in rats fed a safflower oil diet. Metabolism: Clinical and Experimental 44(7):934-9.
- Moher, D., L. Shamseer, M. Clarke, D. Ghersi, A. Liberati, M. Petticrew, P. Shekelle, and L. A. Stewart. PRISMA-P Group 2015. Preferred reporting items for systematic review and Meta-analysis protocols (PRISMA-P) 2015 statement. Systematic Reviews 4(1):1.
- Moreira Alves, R. D., A. P. Boroni Moreira, V. S. Macedo, J. Bressan, R. de Cássia Gonçalves Alfenas, R. Mattes, and N. M. Brunoro Costa. 2014. High-oleic peanuts: new perspective to attenuate glucose homeostasis disruption and inflammation related obesity. Obesity 22(9):1981-8. Silver Spring)
- Morgan, W. A., and B. J. Clayshulte. 2000. Pecans lower low-density lipoprotein cholesterol in people with normal lipid levels. Journal of The American Dietetic Association 100(3):312-8.
- Neale, E. P., L. C. Tapsell, A. Martin, M. J. Batterham, C. Wibisono, and Y. C. Probst. 2017. Impact of providing walnut samples in a lifestyle intervention for weight loss: a secondary analysis of the health track trial. Food & Nutrition Research 61(1):1344522.
- Pasman, W. J., J. Heimerikx, C. M. Rubingh, R. van den Berg, M. O'Shea, L. Gambelli, H. F. Hendriks, A. W. Einerhand, C. Scott, H. G. Keizer, and L. I. Mennen. 2008. The effect of korean pine nut oil on in vitro CCK release, on appetite sensations and on gut hormones in post-menopausal overweight women. Lipids in Health and Disease 7(1):10.
- Pearson, K. R., S. L. Tey, A. R. Gray, A. Chisholm, and R. C. Brown. 2017. Energy compensation and nutrient displacement following regular consumption of hazelnuts and other energy-dense snack foods in non-obese individuals. European Journal of Nutrition 56(3): 1255-67.
- Piers, L. S., K. Z. Walker, R. M. Stoney, M. J. Soares, and K. O'Dea. 2002. The influence of the type of dietary fat on postprandial fat oxidation rates: monounsaturated (olive oil) vs. saturated fat (cream). International Journal of Obesity and Related Metabolic Disorders. Journal of the International Association for the Study of Obesity 26(6):814-21.
- Reis, C. E., D. N. Ribeiro, N. M. Costa, J. Bressan, R. C. Alfenas, and R. D. Mattes. 2013. Acute and second-meal effects of peanuts on glycaemic response and appetite in obese women with high type 2 diabetes risk: A randomised cross-over clinical trial. The British Journal of Nutrition 109(11):2015-23.
- Reis, C. E., L. A. Bordalo, A. L. Rocha, D. M. Freitas, M. V. da Silva, V. C. de Faria, H. S. Martino, N. M. Costa, and R. C. Alfenas. 2011. Ground roasted peanuts leads to a lower post-prandial glycemic response than raw peanuts. Nutricion Hospitalaria 26(4):745-51.
- Rock, C. L., S. W. Flatt, H. S. Barkai, B. Pakiz, and D. D. Heath. 2017. A walnut-containing meal had similar effects on early satiety, CCK, and PYY, but attenuated the postprandial GLP-1 and insulin response compared to a nut-free control meal. Appetite 117:51-7.
- Ros, E. 2015. Nuts and CVD. The British Journal of Nutrition 113(Suppl 2):S111-S20.
- Sayer, R. D., J. Dhillon, G. G. Tamer, M. A. Cornier, N. Chen, A. J. Wright, W. W. Campbell, and R. D. Mattes. 2017. Consuming almonds vs. isoenergetic baked food does not differentially influence



- postprandial appetite or neural reward responses to visual food stimuli. Nutrients 9(8):807.
- Stevenson, J. L., H. C. Clevenger, and J. A. Cooper. 2015. Hunger and satiety responses to high-fat meals of varying fatty acid composition in women with obesity. Obesity (Silver Spring, Md.) 23(10):1980-6.
- Tan, S. Y., and R. D. Mattes. 2013. Appetitive, dietary and health effects of almonds consumed with meals or as snacks: A randomized, controlled trial. European Journal of Clinical Nutrition 67(11):
- Tan, S. Y., J. Dhillon, and R. D. Mattes. 2014. A review of the effects of nuts on appetite, food intake, metabolism, and body weight. The American Journal of Clinical Nutrition 100(Suppl 1):412S-22S.
- Tapsell, L., M. Batterham, S. Y. Tan, and E. Warensjö. 2009. The effect of a calorie controlled diet containing walnuts on substrate oxidation during 8-hours in a room calorimeter. Journal of the American College of Nutrition 28(5):611-7.

- Tey, S. L., A. R. Gray, A. W. Chisholm, C. M. Delahunty, and R. C. Brown. 2013. The dose of hazelnuts influences acceptance and diet quality but not inflammatory markers and body composition in overweight and obese individuals. The Journal of Nutrition 143(8): 1254-62.
- Wien, M., D. Bleich, M. Raghuwanshi, S. Gould-Forgerite, J. Gomes, L. Monahan-Couch, and K. Oda. 2010. Almond consumption and cardiovascular risk factors in adults with prediabetes. Journal of the American College of Nutrition 29(3):189-97.
- Zaveri, S., and S. Drummond. 2009. The effect of including a conventional snack (cereal bar) and a nonconventional snack (almonds) on hunger, eating frequency, dietary intake and body weight. Journal of Human Nutrition and Dietetics 22(5):461-8.
- Zhang, Z., Q. Li, F. Liu, Y. Sun, and J. Zhang. 2010. Prevention of diet-induced obesity by safflower oil: insights at the levels of PPARalpha, orexin, and ghrelin gene expression of adipocytes in mice. Acta Biochimica et Biophysica Sinica 42(3):202-8.