

Chapter

Scientific Insights on the Dietary Management of Obesity Through the Administration of Green Tea Catechins

Vadivel V*, Rajalakshmi P, Ravichandran N and Brindha P

Centre for Advanced Research in Indian System of Medicine (CARISM),
SASTRA Deemed University, India

***Corresponding Author:** vadivel@carism.sastra.edu

First Published **April 22, 2019**

Copyright: © 2019 Vadivel V, Rajalakshmi P, Ravichandran N and Brindha P.

This article is distributed under the terms of the Creative Commons Attribution 4.0 International License (<http://creativecommons.org/licenses/by/4.0/>), which permits unrestricted use, distribution, and reproduction in any medium, provided you give appropriate credit to the original author(s) and the source.

Acknowledgement: Authors are thankful to the Hon'ble Vice-Chancellor of SASTRA Deemed University, Thanjavur, India for his encouragement and support.

Abstract

Obesity is considered as a primary fitness problem that threatens hundreds of thousands of humans at some point of the sector. Occurrence of weight problems ends in incidence of various continual diseases and therefore, remedy/prevention of weight problems is an important target of the current research trends. Catechins is likewise recognized as one of the feasible element, which is accountable for the anti-weight problems attributes of suitable for eating nuts. Hence, the prevailing evaluation accumulating the summation of scientific literature related to anti-obesity assets of nut consumption with regards to their catechin content material and also explains various molecular mechanisms through which, catechins attain prevention of body weight advantage.

Keywords

Obesity; Edible Nuts; Catechins; Dietary Supplements; Body Weight Loss

Introduction

Obesity is increasingly identified as a critical public fitness burden, due to the fact it's far associated with an improved threat for plenty persistent illnesses, such as metabolic syndrome, i.e., high blood pressure, diabetes, arteriosclerosis and coronary heart disease (CHD) [1]. Being obese or overweight, described as a frame mass index between 25 and 30 and a frame mass index (BMI) > 30, respectively [2], is related to a higher risk of growing cardiovascular ailment (CVD) and kind-2 diabetes mellitus [3] and increased mortality [4,5]. For instance, inside the San Antonio coronary heart study, 80% of overweight topics had been hypertensive and diabetic, 85% of diabetics were hypertensive and obese, and 67% of hypertensive have been diabetic and overweight [6]. In 1999 – 2000, the age-adjusted prevalence of obese and obese inside the US became anticipated at 64.5%

and 30.5%, respectively [7]. Internationally obesity is now taken into consideration as a chief fitness trouble of epidemic percentage that threatens thousands and thousands of lives [8]. Almost two thirds of the United States person populace is overweight [7] and 26% are obese, with some states reporting obesity prices as high as 33% [9].

According to the scientific suggestions concerning overweight and weight problems, weight loss and subsequent BMI discount is usually recommended to decrease blood stress, serum lipids and glucose variables to ultimately decrease the risk of CVD and diabetes [2]. Consequently, frame weight reduction remains an essential goal for the prevention of many chronic diseases. Due to the complicated nature of weight problems (genetic elements make contributions about 66% and environmental factors 33%), treatment of weight problems is also a complex problem. A greatest healing method ought to cope with each factor by changing power balance in a greater poor path by increasing electricity expenditure and/or reducing strength consumption and also by improving insulin signaling and metabolism [5].

During the last a long time, several techniques consisting of non-pharmacological and pharmacological were advanced with a purpose to obtain lengthy-time period body weight loss. Non-pharmacological techniques intention to converting eating behavior to: 1. reduce the caloric consumption, fat and simple sugars and 2. growth energy expenditure (EE), particularly by means of bodily hobby. Pharmacological techniques aim to: 1. change eating dependency, i.e., reduce starvation or appetite and boom satiety, 2. boom EE, in particular by means of thermogenesis, 3. given that, conventional weight control programmes have best restrained fulfillment, particularly in lengthy-term efficacy; there's a growing hobby in alternative strategies for weight control, which includes pharmacological interventions [7].

Studies on weight problems inside the area of meals science have centered on the look for meals ingredient that suppresses the accumulation of body fats. In this connection, in current years, the focus on bioactive meals substances and their potential position in stopping weight gain has increased at the side of the worldwide increas-

ing obesity epidemic. Weight problem develops because of every day fine power stability, which in itself can be very small (200 kJ/day), but over the direction of months and years can amount to numerous kilograms of fats [10]. One implication of this is that a growth in EE through similar quantities (200 kJ/day) may want to help to save you this kind of weight gain, furnished that those small consequences can be sustained. Numerous bioactive food ingredients which includes capsaicin, caffeine and catechins had been counseled to be able to eliciting an boom in weight loss program-triggered thermogenesis and thereby each day EE; however, the size consistency of the effects found with those distinct elements vary [11,12].

There may be a growing body of research showing that intake of common nut reduces BMI and body weight. Even though many factors are explaining the mechanism through which the nuts controlling the frame weight, till now there may be no concrete end is derived to denote the thing accountable for anti-weight problems residences of nuts. However, the recent studies efforts found out the presence of catechins - a polyphenolic constituent verified to possess sturdy anti-obesity belongings - within the testa of fit for human consumption nuts. For this reason, the prevailing overview became specifically emphasizing the relationship between nut intake and body weight gain almost about the catechins content material of nuts, and also ambitions to discover the strategies for nutritional administration of edible nuts with appropriate catechins stage for the lengthy-term management of body weight and controlling the obesity.

In latest years, the catechins received extra attention because of their capacity health blessings, such as anti-weight problems impact. Consequently, the catechins content, particularly placed inside the testa of edible nuts, may be one of the possible mechanisms for the control of frame weight benefit of the consumers, in addition to different nutrients and non-nutrients. As a result, in the following phase, the anti-weight problems traits of catechins were described by using thinking about the results of previous investigations on green tea catechins.

Catechins

Catechins belong to the flavan-3-ol elegance of flavonoids and are ubiquitous in flowers and broadly found in lots of ingredients. The maximum prominent resources of catechins are green and black tea, grape/wine, sure culmination and coca [13]. In current years, there has been tremendous hobby in the ability health benefits of catechin-rich foods and drinks. Green tea catechins have numerous organic sports that could possibly offer diverse health blessings [14–18]. Beneficial results of inexperienced tea catechins on infection [19]; angiogenesis [20,21] and oxidation [22] are rising areas of studies hobby. An extended body of evidences counseled that several pathways inside the improvement of metabolic syndromes may be undoubtedly tormented by inexperienced tea catechins.

Green tea is derived from drying and steaming of fresh tea leaves and accordingly no oxidation takes place, resulting in high stages of catechins. Green tea catechins are mainly composed of (-)epigallocatechin-three-gallate (EGCG); (-)epigallocatechin (EGC); (-)epicatechin-three-gallate (ECG) and epicatechin (EC). Epigallocatechin gallate (EGCG) is the most abundant catechin in experienced tea, representing approximately 35 – 40% of total catechins and additionally received maximum interest as an anti-obesigenic agent (Figure 1) [23]. The various useful effects of inexperienced tea on obesity, kind-2 diabetes and CVD are associated with its EGCG content material (Table 1) [24–27].

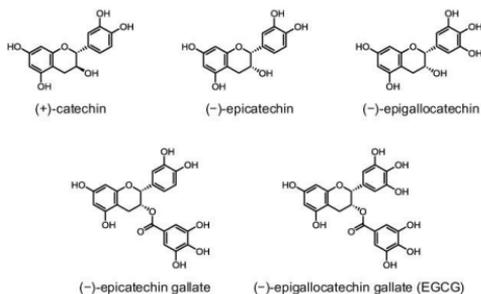


Figure 1: Structure of catechins present in green tea.

Table 1: Human studies assessing the effect of green tea and green tea extract rich in EGCG on weight management.

Citations	Type of study	Popula- tion	Test components (daily dosage)	Durati- on of Intake	Main outcomes		
					Weight (kg)	Fat mass (kg)	BMI
Chantre and Lairon [30]	Multi-center, open, uncontrolled	7 M, 63 F, B M I : 28.9	GTE (375 mg cate- chins, of which 270 mg was EGCG)	12 weeks	- 3.5	Not Repor- ted	Not repor- ted
Hase et al. [33]	Case-control	23 M, BMI: 24 -25	GTE (483.0 mg cate- chins, of which 300 mg was EGCG and 75.5 mg caffeine)	12 weeks	- 0.5	- 1.7	- 0.6
Kajimoto et al. [37]	Double blind, three parallel arm, controlled	98 M, 97 F, B M I : 25.7	Green tea bever- age, low (444 mg catechins, of which 152 mg was EGCG and 50 mg caffeine)	12 weeks	- 1.1	- 3.9%	- 0.4
Kovacs et al. [39]	Randomized par- allel, placebo-controlled	26 M, 78 F, BMI: 25 -35	GTE (573 mg cate- chins, of which 323 mg was EGCG, and 104 mg caffeine)	13 weeks	0.6	0.5	0.2
Nagao et al. [32]	Multi-center, Randomized, dou- bleblind, controlled	140 M, 100 F, BMI: 26.8	Green tea beverage (583 mg catechins, of which 100 mg was EGCG and 72 mg caffeine)	12 weeks	- 1.6	- 1.8	- 0.6
Nagao et al. [31]	Double blind, controlled	35 M, BMI: 24.9 -25.0	GTE (690 mg cate- chins, of which 136 mg was EGCG, and 75 mg caffeine)	12 weeks	- 1.1	- 0.7	- 0.4

Tsuchida et al. [34]	Randomized, double-blind, controlled	43 M, 37 F, BMI: 25.9 – 26.5	GTE (588 mg catechins, of which 115 mg was EGCG and 83 mg caffeine)	12 weeks	- 1.3	- 1.4	- 0.5
Diepvens et al. [46]	Double-blind, placebo-controlled, parallel design	46 F, BMI: 27.7	GTE + low energy diet (1206.9 mg catechins, of which 595.8 mg was EGCG and 236.7 mg caffeine)	12 weeks	0	0	0.1
Chan et al. [45]	Randomized, parallel, placebo-controlled	34 obese F, BMI: 30.9	Capsulated green tea powder (659 mg catechins, of which was 538 mg EGCG and 150 mg caffeine)	12 weeks	- 1.8	- 0.2%	- 0.3
Westert-Plantenga et al. [51]	Randomized, parallel, placebo-controlled	23 M, 53 F, BMI: 25 –35	Low habitual caffeine GTE (<300 mg caffeine) (375 mg catechins, of which 270 mg was EGCG and 150 mg caffeine)	13 weeks	- 2.8	- 2.1	- 0.9
Auvichayapat et al. [36]	Randomized, controlled	18 M, 42 F, BMI: 27 –28	GTE (140.8 mg catechins, of which 100 mg was EGCG and 87 mg caffeine)	12 weeks	- 0.7	- 0.86	- 1.09
Hsu et al. [35]	Randomized, parallel, double-blind, placebocontrolled	78 obese F, BMI: 30.8	GTE (491 mg catechins, of which 302 mg was EGCG and 27 mg caffeine)	12 weeks	- 0.12	- 0.05	Not reported

In most of the instances, gallated catechins, specifically EGCG are extra energetic than different catechins [28]. The found catechin-particular effects of inexperienced tea recommend that EGCG can also act in another way from EC, EGC and ECG in regulating obesity. According to the nature of the specific structures of catechins [29], EGCG contained the biggest range of hydroxyl organizations on its three aromatic rings most of the different catechins and those hydrox-

yl groups are important for hydrogen bonding. Additionally EGCG has both gallyl and galloyl corporations, which have some conformational flexibilities, which could also be essential for interaction with other molecules.

Catechins and Bodyweight

In humans, the outcomes on body weight and body fats in reaction to supplementation with green tea catechins wealthy in EGCG had been explored in numerous intervention studies (Table 2). Chantre and Laeron [30] investigated the outcomes of encapsulated green tea extract in fairly overweight topics (BMI 28.9) and found a 4.6% decrease in body weight and a 4.5% discount in waist-to-hip ratio vs. baseline. In another take a look at, green tea catechins had been imbedded right into a weight reduction verified that after a 12 week supplementation of overweight, body weight and body fats had been reduced considerably while compared to the manipulate organization [31]. This approach presents direct evidence that inexperienced tea catechins can make contributions way of life modifications associated with weight control. Similarly, Nagao et al. [32] confirmed sizable decreases on frame weight and body fat in subjects, who were following their ordinary way of life, at the same time as taking inexperienced tea catechins. This look at is of special interest, as it correlates anti-obesity outcomes of green tea catechins with upgrades in CVD chance factors like systolic blood strain and LDL-C in a fantastically massive population ($n = 240$).

Table 2: Effects of (-)-EGCG on body weight, food intake, adipose tissues, adipogenic hormones, and serum nutrients [54,55,86,61,62].

Parameters	Dose/route/duration	Models	Effects
Body weight	70–92 mg/kg bw, ip, 4–8 days	Rat	Decreasing
	0.5–1% w/w diet, p.o., 4 wk	Rat	No change
	300 mg/kg bw, p.o., 7 days	Mice	No change
	1% w/w diet, p.o., 5 months	Mice	Decreasing
	0.5–1% w/w diet, p.o., 29 days	Mice	Decreasing
Food intake	70–92 mg/kg bw, ip, 4–8 days	Rat	Decreasing
	0.5–1% w/w diet, p.o., 4 wk	Rat	No change
	1%w/w diet, p.o., 5 months	Mice	No change
	0.5–1% w/w diet, p.o., 29 days	Mice	No change
Subcutaneous adipose tissue	70–92 mg/kg bw, ip, 7–8 days	Rat	Decreasing
	1% w/w diet, p.o., 5 months	Mice	Decreasing
Abdominal adipose tissue	70–92 mg/kg bw, ip, 4–8 days	Rat	Decreasing
Epididymal adipose tissue	70–92 mg/kg bw, ip, 4–8 days	Rat	No change
	1% w/w diet, p.o., 5 months	Mice	Decreasing
	0.5–1% w/w diet, p.o., 29 days	Mice	Decreasing
Body fat	70–92 mg/kg bw, ip, 1 wk	Rat	Decreasing
	0.5–1% w/w diet, p.o., 29 days	Mice	Decreasing
Hormones, Leptin	70–92 mg/kg bw, ip, 4–8 days 81 mg/kg bw, p.o., 1 wk	Rat	Decreasing
	1% w/w diet, p.o., 5 months	Mice	Decreasing
	0.5–1% w/w diet, p.o., 29 days	Mice	Decreasing
Cholesterol	70–92 mg/kg bw, ip, 4–8 days	Rat	Decreasing
	0.5–1% w/w diet, p.o., 4 wk	Rat	Decreasing
	500 mg/kg bw, p.o., 7 h	Mice	Decreasing
Glucose	70–92 mg/kg bw, ip, 4–8 days	Rat	Decreasing
	1% w/w diet, p.o., 5 months	Mice	Decreasing
	0.5–1% w/w diet, p.o., 4 wk	Rat	No change
	0.5–1% w/w diet, p.o., 29 days	Mice	No change
Fatty acid	70–92 mg/kg bw, ip, 7 days	Rat	No change
	0.5–1% w/w diet, p.o., 29 days	Mice	No change
Lipid	70–92 mg/kg bw, ip, 4–8 days	Rat	Decreasing
Protein	70–92 mg/kg bw, ip, 4–8 days	Rat	No change
Triglyceride	70–92 mg/kg bw, ip, 4–8 days	Rat	Decreasing
	1% w/w diet, p.o., 5 months	Mice	Decreasing
	0.5–1% w/w diet, p.o., 29 days	Mice	Decreasing
	0.5–1% w/w diet, p.o., 4 wk	Rat	No change

However, greater tightly controlled studies additionally supported the concept that inexperienced tea catechins are powerful in weight and fat weight loss. Two studies on overweight women and men suggested slight discounts in bodyweight and extra stated lower in frame fats [33,34]. Hsu et al. [35] observed slight reductions of 0.12 kg frame weight and 0.05 kg body fats after 12 weeks of supplementation of 78 overweight females (BMI 30.8) with best 27 mg caffeine and 491 mg catechins. Some other current examine mentioned a substantial reduction of frame weight of 0.7 kg after supplementation of catechins (a hundred and 40.8 mg of total catechins containing a 100 mg of EGCG and 27 mg of caffeine) over 12 weeks [36]. In every other take a look at, with a specific approach, every day intake of a manage beverage with low dose of green tea catechins (41 mg) became compared with two high dose degrees (444 and 665 mg) of green tea catechins [37]. In the two higher dose degree corporations, body weight and BMI were substantially reduced.

In overweight girl patients with polycystic ovarian syndrome, a frame weight reduction weight loss of 2.4% turned into suggested following a 12 week supplementation with encapsulated green tea catechins [38]. Unfortunately, this promising locating lacks statistical importance for the among-institution distinction, likely because of the reality that woman patients might have replied differently to the green tea catechins, because of the high degree of weight problems as well as various metabolic modifications due to polycystic syndrome while in comparison to studies of overweight. Kovacs et al. [39] examined how green tea catechins might influence the frame weight after a weight-loss programme. Neither the anticipated frame weight and body fats weight-loss nor a difference in frame weight regain in reaction to the inexperienced tea catechins could be proven. It is not always surprising that impartial elements like green tea catechins did now not similarly increase the already full-size weight-loss on account of a completely low caloric weight loss.

Phung et al. [40] conducted a systematic literature seek and the randomized managed trials that evaluated green tea catechins with-

out or with caffeine and that mentioned BMI, body weight, waist circumference or waist-to-hip ratio had been blanketed. The outcomes imply that, fifteen studies (n = 1243) met with the inclusion standards. On meta-analysis, green tea catechins with caffeine reduced BMI, frame weight and waist circumference, however no longer waist-to-hip ratio compared with caffeine on my own. Green tea catechins ingestion with caffeine in addition extensively reduced frame weight as compared to caffeine. For this reason, the administration of catechins with caffeine is related to statistically considerable reductions in BMI, frame weight and waist circumference; however, the clinical importance of those discounts is discreet the pleasant.

Statistical pooling of facts from 7 trials at the evaluation of catechins with caffeine as compared with a caffeine matched manipulate showed that consuming catechins at a dose ranging from 583 to 714 mg/day over 12 weeks had a significant gain on BMI, frame weight and waist circumference, without a effect on waist-to-hip ratio [41-44]. On pooling, the six trials within the analysis of catechins with caffeine as compared with caffeine-free control. Catechin ingestion significantly decreased frame weight with no impact on BMI, waist circumference or waist-to-hip ratio [45-48]. Of the two caffeine-unfastened trials, pooling the results showed no statistically enormous effect [49]. Curiously, the trial that evaluated EGCG alone [50] confirmed non-large boom in BMI and body weight as compared with placebo. This counseled that the effect of catechins might be because of the mixture, in place of any unmarried catechin.

Few authors speculated that the significance of ordinary caffeine consumption may have masked the outcomes of inexperienced tea catechins. This speculation become recently showed by Westerterp-Plantenga et al. [51], who showed in low level caffeine clients inexperienced tea supplementation similarly notably reduces the body weight and frame fat for the duration of weight maintenance, while in high caffeine purchasers this effect of inexperienced tea catechins couldn't be discovered. However, in contrast, Diepvens et al. [52] found no impact of 206 mg catechins and 236 mg caffeine on body weight and body

fat in obese women over 12 weeks. In all likelihood, the consequences of the catechins are not extra to the weight decreasing effect of the low caloric weight reduction, because the latter already represents a sturdy stimulus to weight reduction frame weight and body fats.

For pharmacologic weight-loss products on marketplace, sufferers are considered to have failed treatment in the event that they have now not done a weight loss 2 kg after 4 weeks of remedy [2]. Green tea catechins with caffeine simplest supplied a mean weight-loss of > 1 kg as compared with a caffeine-matched control, and < 0.5 kg in comparison with a caffeine-free manage taken over an average of 12 weeks. Case reports of catechins consumption have introduced up concerns of hepatotoxicity, and the US nutritional supplements records specialists committee has proposed that the green tea extract products must undergo a label that imply intake together with food, due to the possibility of extreme liver troubles [53] of the rigors that evaluated liver transaminases [50], only one record suggested elevations in the inexperienced tea catechins institution [43]. But, transaminase concentrations have been expanded at baseline, which shows capacity bias in institution allocations. To evaluate issues of liver damage, a randomized trial the usage of high-dose catechins (714 mg/day) became undertaken in healthful men [41]. This trial determined that over three weeks of catechins consumption, there have been no elevations in liver transaminases or reviews of liver disorder.

On the basis of presently to be had literature, ingestion of inexperienced tea catechins with caffeine may additionally positively affect BMI, frame weight and waist circumference. But, the magnitude of impact over 12 weeks is small and no longer likely clinically applicable. In addition, the anti-obesity impact of catechins is depending on the path of management additionally. Even as the results of IP injection of EGCG on frame weight reduction weight loss and meals intake were found, those results aren't observed after oral administration of EGCG inside 7 – 14 days [54] or even 4 weeks [55]. In mice, oral management of ECG or EGCG (three hundred mg/kg BW) within 7 days did no longer adjust body weight [56]. This could be because

of inefficient absorption of EGCG [57] or its speedy metabolism to inactivate molecules within the digestive tract [17] and propose that the results of IP injection of EGCG aren't resulting from interaction of EGCG with meals or via EGCG's motion within gastrointestinal tract. An opportunity cause of the distinction between IP and oral administration may provide distinctive EGCG concentrations in the blood. Although oral administration of EGCG isn't always effective inside 14 days, lengthy-time period oral consumption might also mimic a number of the intense EGCG results caused by IP injection of EGCG and can be beneficial to fitness. This rivalry is evidenced through the truth that oral intake of EGCG can reduce rat, mice and human frame weight [58 - 62] lower serum ldl cholesterol degree in rat, mice and humans [63, 64], boom the rat HDL-C [55], decrease rat and human LDL-C [59, 63] and decrease rat mice and hamster blood triglycerides [65 - 69]. Primarily based on oral and IP results of catechins on serum hormones and nutrients, lengthy-time period intake of EGCG appears to influence the incidence of obesity as reported from scientific studies [30,31].

Animal Studies with Catechins

Numerous intervention studies using animal models have found that the nutritional supplementation of green tea catechins modulates serum lipid profiles [70], and decreases frame weight in addition to adipose tissue mass in rodents (desk 3) [71-73] and these are confirmed via consequences shown in people [74]. EGCG given to rats via an intraperitoneal injection at a dosage of 70 - 92 mg/kg frame weight/day reduced the body weight approximately 20 - 30% within 2 - 7 days [54]. Proximate analysis of animals showed that Sprague-Dowley rats treated with EGCG for 7 days had no change in percent water or protein content, however a 50% decrease in carbohydrate content and 65% reduction in fat content material [54]. EGCG remedy reduced subcutaneous fat by way of forty - 70% and abdominal fat by using 20 - 35%, but now not epididymal fat. A 20% loss of abdominal fats turned into visible in obese male Zucker rats inside four days of EGCG remedy.

The effective dose of EGCG on frame weight is 30 – 50 mg/kg BW. However, rats step by step adapt and within 1 week higher doses of EGCG (100 mg/kg BW) are needed to reduce or prevent body weight will increase [14]. The body weight loss is reversible whilst EGCG management is stopped, animals regain frame weight lost. In aid of anti-obesity impact of EGCG [26], other *in vivo* statistics proven that EGCG reduces the meals uptake, lipid absorption and serum lipids, triglycerides, cholesterol and leptin levels in addition to stimulating EE, fat oxidation, HDL tiers and faecal lipid excretion [75]. Research on anti-obesity action of catechins targeted at the truth that it decreases the fats absorption. Klaus et al. [62] said that faeces power content changed into appreciably and dose-dependently improved through EGCG supplementation. Numerous other researchers reported that inexperienced tea EGCG inhibits the intestinal absorption of nutritional lipids by means of interfering with the emulsification and micellar solubilization of lipids, vital steps concerned in the intestinal absorption of dietary fats, cholesterol and different lipids [76,77].

EGCG additionally said to noticeably lessen epididymal, subcutaneous, mesenteric and retroperitoneal fat weights in high-fat fed mice [73]. These findings suggest that EGCG may contribute to the anti-weight problems effect of green tea catechins. Reduced adipose fat in the course of obese rats handled with EGCG for four days might also give an explanation for the reduced adipose tissue mass and subsequent hypolipidemia of animals treated with catechins [75]. Lee et al. [78] examined the consequences of EGCG at the expression of genes concerned in adipogenesis, lipolysis, beta-oxidation and thermogenesis within the epididymal white adipose tissue. Administration of EGCG (2-five mg/kg/day) in mice displaying no big differences inside the plasma activities of AST and ALT and relative liver weights were now not affected. In addition, no extensive differences have been located within the strength consumption between the control weight-reduction plan-fed mice and EGCG-fed mice. So, EGCG did no longer cause an anorectic effect responsible for prevention and discount of the high-fats eating regimen-precipitated will increase in

body weight and adipose tissue mass. Similarly, the results suggest that, dietary EGCG efficiently decreased frame weight, mass of diverse adipose tissues and plasma concentrations of triglycerides, LDL cholesterol and leptin. This confirms the inverse association between dietary catechin and attention of plasma cholesterol and triglycerides.

Human Trials with Catechins

Long-term treatment (12 weeks) with green tea extract containing 115 mg EGCG consistent with day significantly decreased the body fats (7%), frame weight (2%) and body mass index (BMI) (2%) in both women and men [34]. These findings are supported by different research, wherein wholesome volunteers received a green tea extract containing 270 – 300 mg EGCG reduced the frame weight by means of 1.2% [31] to 1.5% [32]. In a randomized cross-over trial of Oolong tea, which contained 244 mg of EGCG and 270 mg of caffeine, fat oxidation additionally expanded by means of 12% above the manipulate amongst 12 wholesome volunteers, who consumed this tea day by day [79].

Komatsu et al. [80] mentioned that single administration of both Oolong tea (77 mg caffeine and eighty one mg EGCG) or green tea (161 mg caffeine and 156 mg EGCG) increases the electricity expenditure about 111 and 50 kJ, respectively. Similarly, Berube-parent et al. [81] reported a boom of 8% of power expenditure in 24 h after administration of green tea and Guarana extracts containing 600 mg caffeine and 270 mg EGCG. Curiously, higher doses of EGCG (600, 900 and 1200 mg) at a fixed caffeine dose (600 mg) did no longer resulted in addition increase of power expenditure in 24 h [81]. In people also, inexperienced tea extract promoted reduction of frame weight and waist circumference in moderately obese sufferers after 3 months of remedy [30].

In a clinical take a look at France, inexperienced tea extracts containing 25% EGCG exerted its discounts on body weight (4.6%) and waist circumference (4.5%) in fairly overweight patients 3 months af-

ter remedy [30]. In a Japanese look at the subjects ingesting one bottle Oolong tea containing 690 mg catechins per day for 12 weeks had a decrease body weight, body weight index, waist circumference, body fat mass and subcutaneous fats location [31]. From a previous observe in Netherland, Princen et al. [82], concluded that there has been no impact of intake of inexperienced tea catechins (6 cups/day) on body weight index and plasma lipid and antioxidant stages in ordinary weight smokers in the course of a four week length, even as plasma cholesterol and LDL-C tended to decrease after consumption of 3.6 g of green tea catechins/day. Wholesome Japanese men had been given a inexperienced tea extract containing 254 mg catechins [83] and after one hour of management, their plasma degree of EGCG reached 0.27 nM, even as plasma phospholipids, TC and triglycerides did not trade. But, the plasma phosphatidylcholine hydroperoxide degree decreased from 74 pM in controls to forty five pM in EGCG-dealt with topics suggesting that tea catechins are effective antioxidants.

Similarly to research assessing the general anti-weight problems effect of inexperienced tea catechins, several mechanistic research have been done. Specifically, the effect of green tea EGCG on EE and fat oxidation in human beings has obtained a lot of interest. Even as a few have located will increase in EE and fats oxidation by means of 4 and 35%, respectively, after supplementation with inexperienced tea extract containing 270 mg EGCG and 150 mg caffeine [74], others pronounced 2.nine% and 12% for the same parameters after ingesting Oolong tea containing 244 mg EGCG and 270 mg caffeine [79]. The overall fashion of elevated fat oxidation (3.3%) and thermogenesis (4.6%) in response to a beverage containing inexperienced tea EGCG (282 mg), calcium (633 mg) and caffeine (300 mg) is supported by Rudelle et al. [84].

Boschmann and Thielecke [85] pronounced anti-obesigenic consequences of pure EGCG by means of examining the thermogenic and fat oxidation ability of EGCG in overweight guys. EGCG alone has the capacity to increase the fat oxidation, at least inside the post-prandial segment. Fats oxidation extended extensively with the green

tea extract when compared to the control institution, while caffeine by myself resulted in a drastically better growth of fats oxidation [74]. However, no increase on EE was found with EGCG alone [86]. The potential of EGCG to growth the fat oxidation without significantly affecting EE has currently been reported by Klaus et al. [62], who advised the modifications in EE might result from the caffeine present in the tea extracts. There seems to be a most efficient dose of EGCG and caffeine that has the ability to boom fats oxidation. Berube-parent et al. [81] also investigated the results of a steady stage of caffeine (600 mg) combined with various quantities of green tea extracts (270, six hundred, 900, and 1200 mg EGCG). The management of six hundred mg caffeine plus 270 mg EGCG elevated the EE by 750 kJ, although fats oxidation turned into no longer altered.

EGCG and caffeine had been located to modulate EE and fat oxidation fees via one of kind targets. EGCG can inhibit catechol-O-methyltransferase, an enzyme involved inside the degradation of norepinephrine [82]. As a result, as soon as launched the norepinephrine remains in the synaptic cleft longer and presents a extended stimulation of adrenergic receptors. Caffeine additionally inhibits the phosphodiesterase-caused degradation of intracellular cyclic AMP [74]. Each a extended stimulation of adrenergic receptors, especially beta-adrenergic receptors, and an multiplied intracellular cyclic AMP awareness outcomes in an accelerated EE and fats oxidation. The mechanisms, through which green tea catechins lower the power intake and oxidative stress [87].

This assumption is supported by way of the reality that catechins inhibits the pastime of catechol-O-methyltransferase [88], the metabolizing enzyme of norepinephrine and epinephrine, the reality that catechins will increase the EE in obese rats through beta-adrenoceptor activation of thermogenesis in brown adipose tissues [57] and the truth that green tea catechins inward Ca²⁺ currents and modulate stimulus-secretion in bovine adrenal chromaffin cells [89]. Because the sympathoadrenal machine is thought to affect mind features [90], it's far viable that EGCG may additionally reduce meals intake

through stimulating adrenaline pastime [91]. Some of studies have pronounced an increase of EE after the ingestion of caffeine [92–97]. Interestingly, Dulloo et al. [74] located no sizable increase on EE when a hundred and fifty mg of caffeine became administered in tablet form. However, in contrast, Berube-parent et al. [81] showed that, pill preparation containing 600 mg caffeine and ranging quantities of EGCG significantly extended EE.

A latest pass-over, placebo managed have a look at mentioned a boom of fats oxidation of 17% after a supplement containing various inexperienced tea polyphenols and 366 mg EGCG in comparison to manipulate [95]. This look at became finished in everyday weight men for the duration of exercise and virtually supports the findings in animals that green tea catechins exchange the metabolism in favour of improved fat usage. This assumption for this impact is basically as a result of EGCG is supported by way of a have a look at in overweight guys [96]. In this study, the authors pronounced a sizeable reduction inside the postprandial respiration quotient after 3 days supplementation of three hundred mg EGCG per day, suggesting an elevated fat oxidation. It's miles really worth to mention that this examine presents the first proof that a unmarried catechin, particularly EGCG has the capability to reasonably have an effect on fat oxidation.

To date, the maximum extraordinary findings have been pronounced from a protracted-term research that intake of beverage containing 570 mg green tea catechins (218 mg EGCG) for eight weeks increase fats oxidation of 37% and 32% at relaxation and during workout, respectively in 14 healthy topics [97,98]. Interestingly, the beverage contained < 40 mg caffeine, suggesting that EGCG and other catechins are specially liable for this commentary. In any other examine, the results of an encapsulated inexperienced tea extract (250 mg) with low doses of catechin and caffeine, on resting EE and respiratory quotient were tested vs. placebo [36]. The entire day, dose becomes 140 mg catechins with 100 mg EGCG and 87 mg caffeine. The check topics have been obese (BMI 27 – 28) and had a sedentary way of life. Curiously, the outcomes of EGCG were massive for the above parameters after 8 weeks. After 12 weeks, significance becomes

observed most effective for resting EE. Although, the decreased respiration quotient on this research may advocate an accelerated fats oxidation, which complements the long-term study by way of Ota et al. [99] in exercising subjects.

In a double blind, parallel, placebo managed look at related to 38 untrained girl volunteers with a median BMI of 31.2 kg undergone a bodily workout programme for 12 weeks. Half of the volunteers took three hundred mg EGCG similarly to the training programme. In this institution, fats oxidation changed into expanded with the aid of approximately 36% at relative workload whilst in comparison to the control organization. The recurring caffeine intake becomes < 300 mg in keeping with day. It's miles consistent with different proof for effects of EGCG on fat oxidation. In comparison to caffeine, these lengthy-term studies indicate that there is no multiplied metabolic resistance to the consequences of EGCG. The gold standard EGCG boom fats oxidation and guide a weight control has no longer but been installed. The dosage of EGCG utilized in various research ranged from a hundred mg/day [31] to 540 mg/day [38], whilst the duration of the research various from someday to thirteen weeks [81]. The test objects had been administered both within the shape of tablets containing inexperienced tea extract up to 6tablets/day [74] or beverages at as much as 1500 ml/day [79]. the general fine results suggests that the choicest dose and period time for green tea catechins or EGCG administration lies with an affordable variety that may be easily integrated into a weight control programme. Each catechins and caffeine are believed to be liable for the intense results on EE and thermogenesis and additionally fat oxidation located in preceding studies [84].

Currently, the separate and mixed consequences of green tea catechins vs. caffeine on electricity metabolism and fat oxidation over a single day become examined with the aid of Gregersen et al. [100]. Fifteen healthy everyday weight adult males received tablets containing placebo, caffeine on my own (one hundred fifty mg) or caffeine + catechin mixture (600 mg) enriched in EGCG, EGC or combination of catechins in a randomized pass-over double blinded design.

EGCG with caffeine insignificantly raised EE and fats oxidation vs. caffeine only and placebo. The maximum determined effect on EE of about 2% should nonetheless be significant for strength balance over plenty longer length of publicity. The consequences advise that the examined catechin/caffeine combos in sedentary situations exert best small acute effects on EE and fats oxidation, which were now not statistically considerable. Tremendous tendencies discovered suggest that the impact size of catechins/caffeine combinations on EE could at most be within the order of about 2%. Moreover, there does no longer appear to be an effect of the examined catechins/caffeine combos on subjective appetite measures.

Molecular Mechanisms of Catechins to Control Body Weight

Green tea extract containing 25% catechins and rich in EGCG inhibits gastric lipase and pancreatic lipase, the enzymes worried in lipid digestion, *in vitro* at 40 and 80 mg extract in line with g of substrate, respectively [101]. This inhibition is seemingly because of a catechin-brought on lipid emulsification process since the addition of EGCG (55 – 1300 μM) not most effective dose-dependently reduces LDL cholesterol solubility in ciliary micelles however also alters the scale of the combined lecithin/taurocholate/LDL cholesterol micelles [76]. This indicates that the decreased lipid emulsification and digestibility may be chargeable for lowering the intestinal LDL cholesterol absorption, overall fats absorption and serum triglyceride and cholesterol levels.

An *in vitro* study take a look [102] suggest that catechins appears to stimulate the interest of hormone sensitive lipase, that is inside adipocytes and is chargeable for lipid mobilization from adipose tissue to other peripheral tissues. This became evident by way of the fact that, EGCG (10 – 20 $\mu\text{g/ml}$) stimulated a boom in glycerol release by adipocytes into the cytosol after 4 h after incubation with catechin. In contrast, EGCG (one hundred $\mu\text{g/ml}$) inhibited adrenaline and

adrenocorticotrophic hormone-induced lipolysis within the number one fats cells of rats, as indicated by means of decreased launch of fatty acids [103, 104]. The results of EGCG on a selection of lipases are related to the reduced fatty liver and adipose tissue of animals or people treated with catechins.

Green tea catechins are observed to possess anti-lipogenic activity. They inhibit the activity of acetyl CoA carboxylase (ACC), fatty acid synthase (FAS), malic enzyme (ME), glucose-6-phosphate dehydrogenase (G6PDH), glycerol-3-phosphate dehydrogenase (G3PDH) and stearoyl-CoA-desaturase-I (desk 4) [87]. Decrease at the expression of hepatic ME and G6PDH have also been found in obese mice treated with catechins [61]. Catechins have been stated to inhibit the rat liver ACC pastime [105]. Other document suggests that EGCG supplementation down-regulates ACC mRNA expression in overweight mice [61]. EGCG became pronounced to inhibit the activity of chicken FAS [106], so, concurrently decreased lipids, inclusive of triglycerides, phospholipids & cholesterol have been observed. In addition, the interest of G3PDH, the rate limiting step in triglyceride biosynthesis, becomes decreased by way of catechins treatment [107]. Taken collectively, catechins appears to reduce fatty acids and triglycerides synthesis via inhibiting lipogenic enzymes and this can explain the hypolipidic liver, fats cells and blood in catechin administered topics.

Latest research pronounced that nutritional catechins attenuated food plan-caused obesity with the aid of inhibiting mRNA expression of lipogenic enzymes which include FAS and ACC in white adipose tissue [71]. Lee et al. [78] revealed that the, RNA degrees of adipogenic genes which includes Peroxisome proliferator-activated receptor- γ (PPAR- γ), CCAAT-enhancer binding protein- α (CEBP- α), regulatory element-binding protein-1c (SREBP-1c), adipocyte fatty acid-binding protein (aP2), lipoprotein lipase (LPL) and FAS have been notably suppressed dose-dependently by means of EGCG supplementation

in white adipose tissues. In parallel to the suppression of adipogenic genes, mRNA levels of leptin, resistin and adiponin, which are all hormonal elements derived from adipose tissues have been also reduced from 50 – ninety% by EGCG supplementation. These results suggested that EGCG modulates lipid accumulation by using suppressing gene expression of the transcription factors and enzymes associated with adipogenesis and adipocyte-derived hormonal factors in white adipose tissues. Further, Lee et al. [78] additionally found that EGCG improved the mRNA stages of Hormone sensitive lipase (HSL), adipose triglyceride lipase (ATGL), carnitine palmitoyl transferase-1 (CPT-1) and uncoupling protein-2 (UCP-2) genes, contributing to the stimulation of lipid catabolism in adipose tissue. Those results may additionally relate to the mechanisms by way of which the catechins modulate lipolysis in adipocytes.

Numerous LDL cholesterol-related enzymes may be regulated with the aid of inexperienced tea catechins, supporting the viable hypocholesterolaemic impact of catechins. Squalene epoxidase, the rate limiting step in LDL cholesterol biosynthesis, is inhibited with the aid of inexperienced tea catechins [108]. EGCG inhibited the interest of different cholesterol biosynthetic enzymes, lanosterol-14- α -dimethylase and oxido-squalene-lanosterol-cyclase [108]. Together with the inhibition of micelle formation [101] and stimulation of faecal LDL cholesterol excretion by catechins, inhibition of LDL cholesterol-biosynthesis enzymes can be related to the low plasma cholesterol levels found in rats [54].

It must be referred to that, gallated catechins have additionally been inhibited other steroid-related enzymes consisting of eleven-beta-hydroxysteroid dehydrogenase [109], 5-alpha-reductase [110] and aromatase [111]. Reduced five-alpha-reductase activity may be related to weight loss of androgen-dependent organs including prostates and seminal vesicles, found in catechins treated male rat [54]. Decreased aromatase activity can be associated with low blood estrogen degree and the following weight loss of estrogen-structured organs together with uterus, located in EGCG-treated with girl rats [75]. Lately, Lee et

al. [112] mentioned the EGCG stimulate cholesterol α -hydrolase at both mRNA stage and promoter pastime in HepG2 cells, dealt with cholesterol or bile acids. Those results are indicative of an instantaneous modulation of cholesterol α -hydrolase gene expression by using EGCG at transcriptional level.

Green tea catechins can dose-dependently inhibit LDL oxidation in endothelial cells brought about by means of reactive oxygen species [113]. An *in vitro* assay showed that galled epicatechins are more effective in preventing LDL oxidation [114]. Lipid peroxidation enzymes inclusive of lipoxygenase were additionally inhibited via catechins [113]. EGCG also can save you the *in vitro* phosphorylation of glycyrrhizin-binding lipoxygenase by using casein kinase II [55]. These observations advise that the reduction of lipid peroxidation by way of catechins can be related to the mechanism by means of which they modulate lipid shipping in lipoproteins of the lymphatic blood device [114]. It is also cited that, catechins inhibit the radical reaction of apolipoprotein B-100 [115] and will increase the LDL receptor expression in HepG2 and Hela cells [116] and this will relate to the mechanism by way of which catechins reduce blood cholesterol in animals and people [117–119].

A clean assessment of the antimitotic impact of catechins on 3T3-LI preadipocytes started out lately. Preadipocyte proliferation as indicated via improved range of cells [75] and extra incorporation of bromodeoxyuridine [120] become inhibited through EGCG. Weight problem is characterized by way of expanded range and length of fat cells. Subsequently, a thorough research via which, catechins execute modulation of preadipocytic mitogenesis will help in the prevention and control of obesity. The preceding findings established that a suppressive effect of EGCG on preadipocytic proliferation is in all likelihood mediated through Eric mitogen-activated protein kinase (MAPK)-structured and p38 MAPK and JNK MAPK-structured pathways [121,122].

Rats handled with EGCG exhibited tremendous modifications in numerous endocrine parameters [75]. After 7 days of IP remedy with EGCG at a dosage of 85 mg/kg BW, circulating stages of testosterone have been decreased via about 75% in male rats and 17-beta-estradiol stages via 34% in lady rats. Dose-established effects on EGCG at the levels of serum leptin, IFG-I, insulin, boom hormone and luteinizing hormone have been also discovered. EGCG did no longer regulate serum tiers of corticosterone, suggesting the selective consequences of EGCG on endocrine systems. due to the fact, intercourse steroids, insulin and IFG-I are recognized to be anabolic hormones and due to the fact the later enzymes are accountable for stimulating fat cell increase and differentiation [90], the lower in plasma insulin and IFG-I degrees by EGCG can be associated with reduced adipose fats and the subsequent decreases in adipose tissue mass and serum lipid levels in rats [54].

Oral administration of catechins inside the diet (5%) of male rats for 8 weeks showed endocrinological results which includes inducement of goiters, expanded plasma thyroid-stimulating hormone (TSH) stages, decreased frame weights and blood tri-iodothyronine (T3) and Thyroxine (T4) levels, while no extensive trade in follicle-stimulating hormone degrees become located [111]. Due to the fact thyroid hormones are recognized to stimulate the basal metabolic price and carefully concerned with weight loss [90]. The discovered will increase in plasma T3 and T4 stages with ingestion of catechins [111] advocate that other hormones, which include norepinephrine, may be responsible for the increasing thermogenesis and subsequent weight loss in humans/animals treated with green tea catechins. The loss of urge for food might involve neuropeptides other than leptin, due to the fact EGCG is powerful in reducing the frame weight of lean and obese lady and male rats [55]. However, EGCG did not trade the plasma ranges of neurohormones including ACTH, neuropeptide Y, CRF, urocortin and galanin. Plasma tiers of cholecystokinin, a satiety hormone secreted from the gastrointestinal systems and mind are increased in rats given an eating regimen supplemented with catechins [122].

Prolonged postprandial hyperglycemia is a unfavourable aspect for weight problems in addition to diabetes. Park et al. [123] reported the outcomes of circulating green tea catechins on blood glucose and insulin degrees. Oral glucose loading 1 h after green tea extract ingestion in people brought about better blood glucose and insulin levels than on top of things subjects. Gallated catechins had been required for these outcomes, despite the fact that in the intestinal lumen they have been regarded to lower glucose and LDL cholesterol absorption. Treatment with epigallocatechin-3-gallate hindered 2-deoxyglucose uptake into liver, fats, pancreatic beta-cellular, and skeletal muscle cellular lines. The glucose intolerance turned into ameliorated by way of gallated catechin-deficient inexperienced tea extract or inexperienced tea extract mixed with polyethylene glycol, which changed into used as an inhibitor of intestinal absorption of gallated catechins. Those findings may advise that the gallated catechin elevates blood glucose stage by blocking ordinary glucose uptake into the tissues while it's miles in the movement, ensuing in secondary hyperinsulinemia, while it decreases glucose entry into the stream while they are inside the intestinal lumen. Those findings inspire the improvement of non-absorbable derivatives of gallated catechins for preventative remedy of kind 2 diabetes and weight problems, which could especially result in best the tremendous luminal effect [123].

The mechanistic movements of catechins had been found out by diverse *in vitro* and *in vivo* research. Molecular mechanisms doubtlessly contributing to the anti-weight problems outcomes of green tea had been appreciably reviewed [124]. In brief, *in vitro* information proposed that green tea catechins exert their anti-weight problems outcomes through numerous mechanisms consisting of 1. Inhibition of adipocyte differentiation and proliferation, 2. decrease of fats absorption and 3. Inhibition of catechol-O-methyltransferase in brown adipose tissue. Suggested mechanisms by using which EGCG may additionally decrease power consumption, increase EE, and decrease adipose tissue mass and prevent/treat obesity.

In vivo animal research reporting anti-obesity effects of inexperienced tea catechins, especially EGCG are several. The findings comprise: 1. reduction of fats mass, 2. reduction in triglycerides in hyperlipidaemia models as well as discount of unfastened fatty acids and general LDL cholesterol and 3. In combination with staying power workout promoted beta-oxidation and improved exercising ability in mice [125].

Exceedingly little is known about the underlying mechanisms of catechins action within the law of frame weight. Thankfully, positive bases, along with 1. decreasing digestive enzyme activity, 2. increasing lipolytic inhibition, 3. lowering lipogenic activity, 4. increasing fats oxidation and thermogenesis, 5. modulating the expression of lipoproteins, 6. lowering the cell numbers in pre-adipocyte and adipocytes and 7. reducing hormone-inspired proliferation of pre-adipocytes and their differentiation to adipocytes, have these days started out to be examined and should help the *in vivo* anti-obesity consequences of catechins on animals and humans.

Conclusion

From the results of previous dietary intervention studies, it may be concluded that the nut consumption as part of healthy food regimen should save you/control frame weight advantage. Despite the fact that, nuts are electricity-dense and fat-rich food-stuffs, their incorporation has now not been associated with weight benefit, and it is able to lowers the risk of weight problems, which is evidenced with the aid of via inverse courting between nut consumption and BMI in many big cohort research. Epidemiological and medical studies cause the recommendation of frequent nut intake, which in turn, could outcomes in decreased danger of obesity and related diseases. However, further studies are essential to evaluate the possible lengthy-term outcomes of nut intake on weight alternate. It's might critical to emphasize that the recommendation of nuts rather for other power-dense snacks that lack dietary price to facilitate beneficial modifications in nutritional behavior.

The various favorable nutrient profiles of nuts, along with excessive ranges of proteins, fiber, fat with unsaturated fatty acids, the presence of bioactive compounds, specially the catechins seems to play a key function inside the prevention of body weight advantage. Despite the fact that, occurrence of catechins was observed only for few nut sorts that also simplest inside the testa, that is commonly eliminated at some stage in processing techniques. As a result, the processing of nut samples must be re-taken into consideration to keep away from the loss of health beneficial polyphenolic compounds positioned in the testa. Similarly, even though the anti-obesity attributes of catechins of green tea has been established in many clinical trials, till now, the medical evidences are lacking for the catechins contained in safe to eat nuts. subsequently, the anti-obesity traits of catechins present in edible nuts have to be evaluated thru appropriate *in vivo* fashions and also the required degree of catechins to control the obesity must be addressed through human clinical trials, so that the nut consumption alongside appropriate amounts of catechins could be advocated for the manipulate of growing figures of weight problems.

Reference

1. S Yusuf, S Hawken, S Ounpuu, T Dans, A Avezum, F Lanas, et al. INTERHEART Study investigators: Effect of potentially modifiable risk factors associated with myocardial infarction in 52 countries: Case-control study. *Lancet*. 2004; 364: 937 – 952.
2. X Pi-Sunyer, DM Becker, C Bouchard. Clinical guidelines on the identification, evaluation, and treatment of overweight and obesity in adults, National Institutes of Health. 1998. Available Online At: http://www.nhlbi.nih.Gov/guidelines/obesity/ob_gdlns.pdf, 1998.
3. DP Guh, W Zhang, N Bansback, C Amarsi, CL Birmingham, et al. The incidence of co-morbidities related to obe-

- sity and overweight: a systematic review and meta-analysis. *BMC Public Health*. 2009; 9: 88 – 94.
4. GL Burke, A Bertoni, S Shea. The impact of obesity on cardiovascular disease risk factors and subclinical vascular disease. *Archives in Internal Medicine*. 2008; 168: 928 – 935.
 5. KF Adams, A Schatzkin, TB Harris. Overweight, obesity, and mortality in a large prospective cohort of persons 50 to 71 years old. *New England Journal of Medicine*. 2006; 355: 763 – 778.
 6. M Wei, BD Mitchell, SM Haffner, MP Stern. Effects of cigarette smoking, diabetes, high cholesterol, and hypertension on all-cause mortality and cardiovascular disease mortality in Mexican Americans - The San Antonio Heart Study. *American Journal of Epidemiology*. 1996; 144: 1058 – 1065.
 7. KM Flegal, MD Carroll, CL Ogden, CL Johnson. Prevalence and trends in obesity among US adults, 1999–2000. *Journal of the American Medical Association*. 2002; 288: 1723 – 1727.
 8. J Sabate, Y Ang. Nuts and health outcomes: New epidemiologic evidence. *American Journal of Clinical Nutrition*. 2009; 89: 1643S – 1648S.
 9. D Lloyd-Jones, R Adams, M Carnethon. Heart disease and stroke statistics 2009 update: A report from the American Heart Association Statistics Committee and Stroke Statistics Subcommittee. *Circulation*. 2009; 119: 480 – 486.
 10. JO Hill, HR Wyatt, GW Reed. Obesity and the environment: where do we go from here? *Science and Technology Froid*. 2003; 299: 853 – 855.
 11. EMR Kovacs, DJ Mela. Metabolically active functional food ingredients for weight control. *Obesity Reviews*. 2006; 7: 59 – 78.

12. MS Westerterp-Plantenga, K Diepvens, AMCP Joosen. Metabolic effects of spices, teas, and caffeine, *Physiology and Behavior*, 89, 85 – 91, 2006.
13. T Raab, D Barron, FA Vera, V Crespy, M Oliveira, et al. Catechin glucosides: Occurrence, synthesis, and stability. *Journal of the Agricultural and Food Chemistry*. 2010; 58: 2138 – 2149.
14. S Liao, YH Kao, RA Hiipakka. Green tea: Biochemical and biological basis for health benefits. *Vitamins and Hormones*. 2001; 62: 1 – 94.
15. CS Yang, ZY Wang. Tea and Cancer. *Journal of National Cancer Institute*. 1993; 85: 1038 – 1049.
16. LA Mitscher, M Jung, D Shankel, JH Dou. Chemoprotection: A review of the potential therapeutic antioxidant properties of green tea (*Camellia sinensis*) and certain of its constituents. *Medical Research Reviews*. 1997; 17: 327 – 365.
17. JK Lin, YC Liang, SY Lin-Shiau. Cancer chemoprevention by tea polyphenols through mitotic signal transduction blockade. *Biochemical Pharmacology*. 1999; 58: 911 – 915.
18. N Ahmad, H Mukhtar. Green tea polyphenols and cancer: Biologic mechanisms and practical implications. *Nutrition Reviews*. 1999; 57: 78 – 83.
19. V Crespy, G Williamson. A review of the health effects of green tea catechins in in vivo animal models. *Journal of Nutrition*. 2004; 134: 3431S – 3440S.
20. M Dona, I Dell'Aica, F Calabrese, R Benelli, M Morini, et al. Neutrophil restraint by green tea: inhibition of inflammation, associated angiogenesis, and pulmonary fibrosis. *Journal of Immunology*. 2003; 170: 4335 – 4341.
21. SK Rodriguez, W Guo, L Liu, MA Band, EK Paulson, et al. Green tea catechin, epigallocatechin-3-gallate, inhibits vascular endothelial growth factor angiogenic signaling by dis-

- rupting the formation of a receptor complex. *International Journal of Cancer*. 2006; 118: 1635 – 1644.
22. MR Sartippour, ZM Shao, D Heber, P Beatty, L Zhang, et al. Green tea inhibits vascular endothelial growth factor (VEGF) induction in human breast cancer cells. *Journal of Nutrition*. 2002; 132: 2307 – 2311.
 23. K Osada, M Takahashi, S Hoshina, M Nakamura, S Nakamura, et al. Tea catechins inhibit cholesterol oxidation accompanying oxidation of low density lipoprotein *in vitro*. *Comparative Biochemistry and Physiology Part-C: Toxicology and Pharmacology*. 2001; 128: 153 – 164.
 24. CS Yang, JM Landau. Effects of tea consumption on nutrition and health. *Journal of Nutrition*. 2000; 130: 2409 – 2412.
 25. JD Lambert, MJ Lee, H Lu, X Meng, JJ Hong, et al. Epigallocatechin-3-gallate is absorbed but extensively glucuronidated following oral administration to mice. *Journal of Nutrition*. 2003; 133: 4172 – 4177.
 26. S Mandel, O Weinreb, T Amit, MB Youdim. Cell signaling pathways in the neuroprotective actions of the green tea polyphenol (+)-epigallocatechin-3-gallate: implications for neurodegenerative diseases. *Journal of Neurochemistry*. 2004; 88: 1555 – 1569.
 27. SB Moyers, NB Kumar. Green tea polyphenols and cancer chemoprevention: multiple mechanisms and endpoints for phase II trials. *Nutrition Reviews*. 2004; 62: 204 – 211.
 28. OJ Park, YJ Surh. Chemopreventive potential of epigallocatechin gallate and genistein: evidence from epidemiological and laboratory studies. *Toxicology Letters*. 2004; 150: 43 – 56.
 29. T Goto, Y Yoshida, M Kiso, H Nagashima. Simultaneous analysis of individual catechins and caffeine in green tea. *Journal of Chromatography: A*. 1996; 749: 295 – 299.

30. P Chantre, D Lairon. Recent findings of green tea extract AR25 (Exolise) and its activity for the treatment of obesity. *Phytomedicine*. 2002; 9: 3 – 8.
31. T Nagao, Y Komine, S Soga. Ingestion of a tea rich in catechins leads to a reduction in body fat and malondialdehyde-modified LDL in men. *The American Journal of Clinical Nutrition*. 2005; 81: 122 – 129.
32. T Nagao, T Hase, I Tokimitsu. A green tea extract high in catechins reduces body fat and cardiovascular risks in humans. *Obesity*. 2007; 15: 1473 – 1483.
33. T Hase, Y Komine, S Meguro. Anti-obesity effects of tea catechins in humans. *Journal of Oleo Science*. 2001; 50: 599 – 605.
34. T Tsuchida, I Hiroshige, N Haruo. Reduction of body fat in humans by long-term ingestion of catechins. *Progress Medicine*. 2002; 22: 2189 – 2203.
35. CH Hsu, TH Tsai, YH Kao, KC Hwang, TY Tseng, et al. Effect of green tea extract on obese women: a randomized, double-blind, placebo-controlled clinical trial. *Clinical Nutrition*. 2008; 27: 363 – 370.
36. P Auvichayapat, M PrapoChanung, O Tunkamnerdthai, BO Sripanidkulchai, N Auvichayapat, et al. Effectiveness of green tea on weight reduction in obese thais: A randomized, controlled trial. *Physiology and Behavior*. 2008; 93: 486 – 491.
37. O Kajimoto, Y Kajimoto, M Yabune, T Nakamura, K Kotani. Tea catechins with a galloyl moiety reduce body weight and fat. *Journal of Health Sciences*. 2006; 1: 161 – 171.
38. Z Chen, G Yang, M Zhou, M Smith, A Offer, et al. Body mass index and mortality from ischaemic heart disease in a lean

- population: 10 year prospective study of 220,000 adult men. *International Journal of Epidemiology*. 2006; 35: 141 – 150.
39. EMR Kovacs, MPMG Lejeune, I Nijs. Effects of green tea on weight maintenance after body-weight loss. *British Journal of Nutrition*. 2004; 91: 431 – 437.
 40. OJ Phung, WL Baker, LJ Matthews, M Lanosa, A Thorne, et al. Effect of green tea catechins with or without caffeine on anthropometric measures: a systematic review and meta-analysis. *The American Journal of Clinical Nutrition*. 2010; 91: 73 – 81.
 41. J Frank, TW George, JK Lodge. Daily consumption of an aqueous green tea extract supplement does not impair liver function or alter cardiovascular disease risk biomarkers in healthy men. *Journal of Nutrition*. 2009; 139: 58 – 62.
 42. KC Maki, MS Reeves, M Farmer. Green tea catechin consumption enhances exercise-induced abdominal fat loss in overweight and obese adults. *Journal of Nutrition*. 2009; 139: 264 – 270.
 43. T Nagao, S Meguro, T Hase. A catechin-rich beverage improves obesity and blood glucose control in patients with type 2 diabetes. *Obesity*. 2009; 17: 310 – 317.
 44. T Matsuyama, Y Tanaka, I Kamimaki, T Nagao, I Tokimitsu. Catechin safely improved higher levels of fatness, blood pressure, and cholesterol in children. *Obesity*. 2008; 16: 1338 – 1348.
 45. CC Chan, MW Koo, EH Ng, OS Tang, WS Yeung, et al. Effects of Chinese green tea on weight, and hormonal and biochemical profiles in obese patients with polycystic ovary syndrome – A randomized placebo-controlled trial. *Journal of the Society for Gynecologic Investigations*. 2006; 13: 63 – 68.

46. K Diepvens, EMR Kovacs, IMT Nijs. Effect of green tea on resting energy expenditure and substrate oxidation during weight loss in overweight females. *British Journal of Nutrition*. 2005; 94: 1026 – 1034.
47. Y Fukino, M Shimbo, N Aoki, T Okubo, H Iso. Randomized controlled trial for an effect of green tea consumption on insulin resistance and inflammation markers. *Journal of Nutritional Sciences and Vitaminology*. 2005; 51: 335 – 342.
48. DJ Maron, GP Lu, NS Cai. Cholesterol-lowering effect of a theaflavin-enriched green tea extract. *Archives of Internal Medicine*. 2003; 163: 1448 – 1453.
49. M Takeshita, S Takashima, U Harada. Effects of long-term consumption of tea catechins-enriched beverage with no caffeine on body composition in humans. *The Japanese Journal of Pharmacologic Therapy*. 2008; 36: 767 – 776.
50. M Hill, AM Coates, JD Buckley. Can EGCG reduce abdominal fat in obese subjects? *Journal of the American College Nutrition*. 2007; 26: 396S – 402S.
51. MS Westerterp-Plantenga, MP Lejeune, EM Kovacs. Body weight loss and weight maintenance in relation to habitual caffeine intake and green tea supplementation. *Obesity Research*. 2005; 13: 1195 – 1204.
52. K Diepvens, EM Kovacs, N Vogels, MS Westerterp-Plantenga. Metabolic effects of green tea and of phases of weight loss. *Physiology and Behavior*. 2006; 87: 185 – 191.
53. DN Sarma, ML Barrett, ML Chavez. Safety of green tea extracts: a systematic review by the US Pharmacopeia. *Drug Safety*. 2008; 31: 469 – 484.
54. YH Kao, RA Hiipakka, S Liao. Modulation of endocrine systems and food intake by green tea epigallocatechin-gallate. *Endocrinology*. 2000; 141: 980 – 987.

55. M Fukuyo, Y Hara, K Muramatsu. Effect of tea leaf catechin, (-)-pigallocatechin gallate on plasma cholesterol level in rats. *Journal of the Japanese Society for Nutrition and Food Sciences*. 1986; 39: 495 – 500.
56. H Matsuda, T Chisaka, Y Kubomura, J Yamahara. Effects of crude drugs on experimental hypercholesterolemia. I. Tea and its active principles. *Journal of Ethnopharmacology*. 1986; 17: 213 – 224.
57. JJ Choo. Green tea reduces body fat accretion caused by high-fat diet in rats through beta-adrenoceptor activation of thermogenesis in brown adipose tissue. *Journal of Nutritional Biochemistry*. 2003; 14: 671 – 676.
58. N Hasegawa, N Yamda, M Mori. Powdered green tea has antilipogenic effect on Zucker rats fed a high-fat diet. *Phytotherapy Research*. 2003; 17: 477 – 480.
59. G Zheng, K Sayama, T Okubo, LR Juneja, I Oguni. Anti-obesity effects of three major components of green tea, catechins, caffeine and theanine, in mice. *In Vivo*. 2004; 18: 55 – 62.
60. KL Kuo, MS Weng, CT Chiang, YJ Tsai. Comparative studies on the hypolipidemic and growth suppressive effects of oolong, black, pu-erh, and green tea leaves in rats. *Journal of Agricultural Food Chemistry*. 2005; 53: 480 – 489.
61. S Wolfram, D Raederstorff, Y Wang, SR Teixeira. TEAVIG-OTM (Epigallocatechin Gallate) supplementation prevents obesity in rodents by reducing adipose tissue mass. *Annals of Nutrition and Metabolism*. 2005; 49: 54 – 63.
62. S Klaus, S Pultz, C Thone-Reineke, S Wolfram. Epigallocatechin gallate attenuates diet-induced obesity in mice by decreasing energy absorption and increasing fat oxidation. *International Journal of Obesity and Related Metabolic Disorders*. 2005; 29: 615 – 623.

63. K Muramatsu, M Fukuya, Y Hara. Effect of green tea catechins on plasma cholesterol level in cholesterol-fed rats. *Journal of Nutritional Sciences and Vitaminology*. 1986; 32: 613 – 622.
64. S Kono, K Shinchu, K Wakabayashi, S Honjo. Relation of green tea consumption to serum lipids and lipoproteins in Japanese men. *Journal of Epidemiology*. 1996; 6: 128 – 133.
65. K Sayama, S Lin, G Zheng, I Oguni. Anti-obesity effects of three major components of green tea, catechins, caffeine and theanine, in mice. *In Vivo*. 2000; 14: 481 – 484.
66. H Ashida, T Furuyashiki, H Nagayasu, H Bessho, H Sakakibara, T Hashimoto, et al. Anti-obesity actions of green tea: possible involvements in modulation of the glucose uptake system and suppression of the adipogenesis related transcription factors. *Biofactors*. 2004; 22: 135 – 140.
67. TTC Yang, MWL Koo. Hypocholesterolemic effects of Chinese tea. *Pharmacological Research*. 1997; 35: 505 – 512.
68. PT Chan, WP Fong, YL Cheung, Y Huang. Jasmine green tea epicatechins are hypolipidemic in Hamsters (*Mesocricetus auratus*) fed a high fat diet. *Journal of Nutrition*. 1999; 129: 1094 – 1101.
69. M Yang, C Wang, H Chen. Green, oolong and black tea extracts modulate lipid metabolism in hyperlipidemia rats fed high-sucrose diet. *Journal of Nutritional Biochemistry*. 2001; 12: 14 – 20.
70. M Kobayashi, T Unno, Y Suzuki, A Nozawa, Y Sagesaka, et al. Heat-epimerized tea catechins have the same cholesterol-lowering activity as green tea catechins in cholesterol-fed rats. *Biosciences Biotechnology and Biochemistry*. 2005; 69: 2455 – 2458.

71. T Murase, A Nagasawa, J Suzuki, T Hase, I Tokimitsu. Beneficial effects of tea catechins on diet-induced obesity: stimulation of lipid catabolism in the liver. *International Journal of Obesity and Related Metabolic Disorders*. 2002; 26: 1459 – 1464.
72. S Wie, D Raederstorff, Y Wang, SR Teixeira, V Elste, et al. TEAVIGO (epigallocatechin gallate) supplementation prevents obesity in rodents by reducing adipose tissue mass. *Annals of Nutrition and Metabolism*. 2005; 49: 54–63.
73. M Bose, JD Lambert, J Ju, KR Reuhl, SA Shapses, CS Yang. The major green tea polyphenol, (–)-epigallocatechin-3-gallate, inhibits obesity, metabolic syndrome, and fatty liver disease in high-fat-fed mice. *Journal of Nutrition*. 2008; 138: 1677 – 1683.
74. G Dulloo, C Duret, D Rohrer. Efficacy of a green tea extract rich in catechin polyphenols and caffeine in increasing 24-h energy expenditure and fat oxidation in humans. *The American Journal of Clinical Nutrition*. 1999; 70: 1040 – 1045.
75. YH Kao, RA Hippakka, S Liao. Modulation of obesity by a green tea catechin. *The American Journal of Clinical Nutrition*. 2000; 72: 1232 – 1241.
76. DG Raederstorff, MF Schlachter, V Elste, P Weber., Effect of EGCG on lipid absorption and plasma lipid levels in rats. *Journal of Nutritional Biochemistry*. 2003; 14: 326 – 332.
77. S Koo, S Noh. Green tea as inhibitor of the intestinal absorption of lipids: potential mechanism for its lipid-lowering effect. *Journal of Nutritional Biochemistry*. 2007; 18: 179 – 183.
78. MS Lee, CT Kim, Y Kim. Green tea (–)-epigallocatechin-3-gallate reduces body weight with regulation of multiple genes expression in adipose tissue of diet-induced obese

- mice. *Annals of Nutrition and Metabolism*. 2009; 54: 151 – 157.
79. W Rumpler, J Seale, B Clevidence, J Judd, E Wiley, et al. Oolong tea increases metabolic rate and fat oxidation in men. *Journal of Nutrition*. 2001; 131: 2848 – 2852.
80. T Komatsu, M Nakamori, K Komatsu, K Hosoda, M Okamura, et al. Oolong tea increases energy metabolism in Japanese females. *Journal of Medical Investigations*. 2003; 50: 170 – 175.
81. S Berube-Parent, C Pelletier, J Dore. Effects of encapsulated green tea and guarana extracts containing a mixture of epigallocatechin-3-gallate and caffeine on 24 h energy expenditure and fat oxidation in men. *British Journal of Nutrition*. 2005; 94: 432 – 436.
82. HM Princen, D Van, R Buytenhek, C Blonk, LB Tijburg, et al. No effect of consumption of green and black tea on plasma lipid and antioxidant levels and on LDL oxidation in smokers. *Arteriosclerosis, Thrombosis and Vascular Biology*. 1998; 18: 833 – 841.
83. K Nakagawa, M Ninomiya, T Okubo, N Aoi. Tea catechin supplementation increases antioxidant capacity and prevents phospholipid hydroperoxidation in plasma of humans. *Journal of Agricultural and Food Chemistry*. 1999; 47: 3967 – 3973.
84. S Rudelle, MG Ferruzzi, I Cristiani. Effect of a thermogenic beverage on 24-hour energy metabolism in humans. *Obesity*. 2007; 15: 349 – 355.
85. M Boschmann, F Thielecke. The effects of epigallocatechin-3-gallate on thermogenesis and fat oxidation in obese men: A pilot study. *Journal of the American College Nutrition*. 2007; 26: 389S – 395S.

86. RT Borchardt, JA Huber. Catechol O-methyltransferase. 5. Structure–activity relationships for inhibition by flavonoids. *Journal of Medicinal Chemistry*. 1975; 18: 120 – 122.
87. YH Kao, HH Chang, MJ Lee, CL Chen. Tea, obesity, and diabetes. *Molecular Nutrition and Food Research*. 2006; 50: 188 – 210.
88. H Lu, X Meng, CS Yang. Enzymology of methylation of tea catechins and inhibition of catechol-O-methyltransferase by (-)-Epigallocatechin gallate. *Drug Metabolism and Disposition*. 2003; 31: 572 – 579.
89. CY Pan, YH Kao, AP Fox. Enhancement of inward Ca²⁺ currents in bovine chromaffin cells by green tea polyphenol extracts. *Neurochemistry International*. 2002; 40: 131 – 137.
90. GM Besser, MO Thorner. *Comprehensive clinical endocrinology*, 3rd edn. London: MOSBY, an affiliate of Elsevier Sci. 2002; 256 – 278.
91. G Dulloo, J Seydoux, L Girardier. Green tea and thermogenesis: interactions between catechin-polyphenols, caffeine and sympathetic activity. *International Journal of Obesity*. 2000; 24: 252 – 258.
92. KJ Acheson, B Zahorska-Markiewicz, P Pittet, K Anantharaman, E Jequier. Caffeine and coffee: their influence on metabolic rate and substrate utilization in normal weight and obese individuals. *The American Journal of Clinical Nutrition*. 1980; 33: 989 – 397.
93. Astrup, S Toubro, S Cannon, P Hein, L Breum, et al. Caffeine: a double-blind, placebo-controlled study of its thermogenic, metabolic, and cardiovascular effects in healthy volunteers. *The American Journal of Clinical Nutrition*. 1990; 51: 759 – 767.

94. TJ Horton, CA Geissler. Post-prandial thermogenesis with ephedrine caffeine and aspirin in lean, pre-disposed obese and obese women. *International Journal of Obesity and Related Metabolic Disorders*. 1996; 20: 91 – 97.
95. PJ Arciero, CL Bougopoulos, BC Nindl, NL Benowitz. Influence of age on the thermic response to caffeine in women. *Metabolism*. 2000; 49: 101 – 107.
96. G Dulloo, CA Geissler, T Horton, A Collins, DS Miller. Normal caffeine consumption: influence on thermogenesis and daily energy expenditure in lean and post-obese human volunteers. *The American Journal of Clinical Nutrition*. 1989; 49: 44 – 50.
97. D Bracco, JM Ferrarra, MJ Arnaud, E Jequier, Y Schutz. Effects of caffeine on energy metabolism, heart rate, and methylxanthine metabolism in lean and obese women. *The American Journal of Physiology*. 1995; 269: 671 – 678.
98. MC Venables, CJ Hulston, HR Cox. Green tea extract ingestion, fat oxidation, and glucose tolerance in healthy humans. *The American Journal of Clinical Nutrition*. 2008; 87: 778 – 784.
99. N Ota, S Soga, A Shimotoyodome, M Inaba, T Murase, et al. Effects of combination of regular exercise and tea catechins intake on energy expenditure in humans. *Journal of Health Sciences*. 2005; 51: 233 – 236.
100. NT Gregersen, C Bitz, I Krog-Mikkelsen, O Hels, EMR Kovacs, et al. Effect of moderate intakes of different tea catechins and caffeine on acute measures of energy metabolism under sedentary conditions. *British Journal of Nutrition*. 2009; 102: 1187 – 1194.
101. C Juhel, M Armand, Y Pafumi, C Rosier, J Vandermander, et al. Green tea extract [AR25 (R)] inhibits lipolysis of triglyc-

- erides in gastric and duodenal medium *in vitro*. Journal of Nutritional Biochemistry. 2000; 11: 45 – 51.
102. M Mochizuki, N Hasegawa. Effects of green tea catechin-induced lipolysis on cytosol glycerol content in differentiated 3T3-L1 cells. Phytotherapy Research. 2004; 18; 945 – 946.
103. Y Kimura, H Okuda, T Okuda, T Yoshida. Studies on the activities of tannins and related compounds of medicinal plants and drugs. II. Effects of various tannins and related compounds on adrenaline-induced lipolysis in fat cells. Chemical and Pharmaceutical Bulletin. 1983; 31: 2497 – 2500.
104. Y Kimura, H Okuda, T Okuda, T Yoshida. Studies on the activities of tannins and related compounds of medicinal plants and drugs. III. Effects of various tannins and related compounds on adrenocorticotrophic hormon-induced lipolysis and insulin-induced lipogenesis from glucose in fat cells. Chemical and Pharmaceutical Bulletin. 1983; 31: 2501 – 2506.
105. J Watanabe, J Kawabata, R Niki. Isolation and identification of acetyl-CoA carboxylase inhibitors from green tea (*Camellia sinensis*). Biosciences Biotechnology and Biochemistry. 1998; 62: 532 – 534.
106. X Wang, W Tian. Green tea epigallocatechin gallate: A natural inhibitor of fatty-acid synthase. Biochemistry and Biophysics Research Communications. 2001; 288: 1200 – 1206.
107. M Mochizuki, N Hasegawa. Stereospecific effects of catechin isomers on insulin induced lipogenesis in 3T3-L1 cells. Phytotherapy Research. 2004; 18: 449 – 450.
108. Abe, T Seki, K Umehara, T Miyase. Green tea polyphenols: Novel and potent inhibitors of squalene epoxidase. Biochemistry and Biophysics Research Communications. 2000; 268: 767 – 771.

109. Guo, MM Reidenberg. Inhibition of 11β -hydroxysteroid dehydrogenase by bioflavonoids and their interaction with furosemide and gossypol. *Journal of Laboratory and Clinical Medicine*. 1998; 132: 32 – 38.
110. RA Hiipakka, HZ Zhang, W Dai, Q Dai, S Liao. Structure–activity relationships for inhibition of human 5α -reductases by polyphenols. *Biochemistry and Pharmacology*. 2002; 63: 1165 – 1176.
111. Satoh, Y Sakamoto, A Ogata, F Nagai. Inhibition of aromatase activity by green tea extract catechins and their endocrinological effects of oral administration in rats. *Food Chemistry and Toxicology*. 2002; 40: 925 – 933.
112. S Lee, JY Park, H Freake, IS Kwun, Y Kim. Green tea enhances cholesterol 7-hydroxylase gene expression in HepG2 cells. *British Journal of Nutrition*. 2008; 99: 1182 – 1185.
113. CT Ho, Q Chen, H Shi, KQ Zhang, RT Rosen. Antioxidative effect of polyphenol extract prepared from various Chinese teas. *Preventive Medicine*. 1992; 21: 520 – 525.
114. Ikeda, K Tsuda, Y Zuzuki, M Kobayashi. Tea catechins with a galloyl moiety suppress postprandial hypertriacylglycerolemia by delaying lymphatic transport of dietary fat in rats. *Journal of Nutrition*. 2005; 135: 155 – 159.
115. R Hashimoto, M Yaita, K Tanaka, Y Hara, S Kojo. Inhibition of radical reaction of apolipoprotein B-100 and α -tocopherol in human plasma by green tea catechins. *Journal of Agricultural and Food Chemistry*. 2000; 48: 6380 – 6383.
116. DJ Kuhn, AC Burns, A Kazi, QP Dou. Direct inhibition of the ubiquitin–proteasome pathway by ester bond-containing green tea polyphenols is associated with increased expression of sterol regulatory element-binding protein 2 and LDL receptor. *Biochimica Biophysica Acta*. 2004; 1682: 1 – 10.

117. S Tokunaga, IR White, C Frost, K Tanaka. Green tea consumption and serum lipids and lipoproteins in a population of healthy workers in Japan. *Annals of Epidemiology*. 2002; 12: 157 – 165.
118. Imai, K Nakachi. Cross sectional study of effects of drinking green tea on cardiovascular and liver diseases. *British Medical Journal*. 1995; 310: 693 – 696.
119. CH Wu, FH Lu, CS Chang, TC Chang. Relationship among habitual tea consumption, percent body fat, and body fat distribution. *Obesity Research*. 2003; 11: 1088 – 1095.
120. PF Hung, BT Wu, HC Chen, YH Chen, CL Chen, et al. Antimitogenic effect of green tea (+)-epigallocatechin gallate on 3T3-L1 preadipocytes depends on the ERK and Cdk2 pathways. *American Journal of Physiology: Cell Physiology*. 2005; 288: C1094 – C1108.
121. CM Boney, RM Smith, PA Gruppuso. Modulation of insulin-like growth factor I mitogenic signaling in 3T3-L1 preadipocyte differentiation. *Endocrinology*. 1998; 139: 1638 –1644.
122. T Yamamoto, LR Juneja, DC Chu, M Kim. *Chemistry and applications of green tea*. Boca Raton: CRC Press LLC. 1997; 165 – 214.
123. H Park, JY Jin, WK Baek, SH Park, HY Sung, et al. Ambivalent role of gallated catechins in glucose tolerance in humans: a novel insight into non-absorbable gallated catechin-derived inhibitors of glucose absorption. *Journal of Physiology and Pharmacology*. 2009; 60: 101 – 109.
124. S Wolfram, Y Wang, F Thielecke. Anti-obesity effects of green tea: From bedside to bench. *Molecular Nutrition and Food Research*. 2006; 50: 176 – 187.

125. T Murase, S Haramizu, A Shimotoyodome, I Tokimitsu, T Hase. Green tea extract improves running endurance in mice by stimulating lipid utilization during exercise. *American Journal of Physiology: Regulatory, Integrative and Comparative Physiology*. 2006; 290: R1550 – R1556.
126. Shimotoyodome, S Haramizu, M Inaba, T Murase, I Tokimitsu. Exercise and green tea extract stimulate fat oxidation and prevent obesity in mice. *Medicine and Science in Sports and Exercise*. 2005; 37: 1884 – 1892.