

## REVIEW

# Dietary supplements for obesity

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## Keywords

Dietary supplements • Obesity • Weight loss • Nutritional disease

## Summary

*Obesity and associated complications including diabetes, cardiometabolic dysfunction, disability, malignancy and premature mortality are considered epidemic. Research on obesity is therefore of worldwide importance. The development of obesity is a multifactorial phenomenon with contributions from biological, behavioral, genetic and environmental factors. Obesity and its associated issues require various lifestyle modifications and treatment options such medication, exercise, diet, surgery, pharmacological therapy and dietary supplements. Dietary supplements are considered an attractive alternative to traditional therapy due to their low toxicity profile and their accessibility to the general population. Dietary supplements may include one or more dietary ingredients. In this narrative review, we analyze the effects on obesity and obesity-re-*

*lated issues of various natural components. For example, there are a myriad of supplements that have been used as dietary supplements for weight loss such as minerals, vitamins, amino acids, metabolites, herbs, and plant extracts. This narrative review aims to present the benefits and side-effects of several ingredients of dietary supplements for weight loss and treatment of obesity. In particular, the mechanism of action, results of clinical trials, and possible side effects will be presented for the following ingredients:  $\beta$ -Glucans, bitter orange, calcium, vitamin D, chitosan, chromium, cocoa, coleus forskohlii, conjugate linoleic acid, ephedra sinica, fucoxanthin, garcinia cambogia, glucomannan, green coffee, green tea, guar gum, raspberry, hoodia gordonii, irvingia gabonensis, phenylpropylamine, pyruvate, white kidney bean.*

## Introduction

Obesity is one of the most prevalent nutritional diseases in industrialized countries and is recognized as a significant public health issue. By definition, obesity is the excessive accumulation of adipose tissue. Excess body weight is classically determined on the basis of a person's BMI (body mass index):  $BMI = \text{weight (kg)} / \text{height (m)}^2$ . A BMI of  $30 \text{ kg/m}^2$  is recognized as the obesity threshold, while a BMI comprised between 25 and  $29.9 \text{ kg/m}^2$  is used to identify overweight people; BMIs over the obesity threshold are associated with higher morbidity and mortality rates and a wide range of symptoms. Excessive weight gain is considered a multifactorial phenomenon that involves variably interacting biological, behavioral, genetic, and environmental factors [1-4].

Obesity and associated complications like cardiometabolic dysfunction, diabetes, disability, malignancy, and premature death are considered epidemic in the western world [5-7]. Treatment of obesity is therefore of major significance for researchers around the world. In the United States, the obesity rate continues to increase, and worsened during the COVID-19 pandemic [8]. Obesity, overweight and associated issues call for preventive and treatments approaches. Indeed, several solutions have been proposed, such as following a healthy lifestyle, following eating plans for weight loss, and exercising. Moreover, interventional approaches such as surgery,

pharmacological therapy and dietary supplements have also been developed [7, 9]. The U.S. Food and Drug Administration (FDA) has approved various traditional approaches including pharmacological, surgical, and endoscopic bariatric therapies that may promote weight loss of 5-35% [10]. Dietary supplements are considered an attractive alternative to traditional therapy due to their low toxicity and accessibility to the general population. Almost 33.9% of adults attempting weight loss, largely young adults, women, and lower socioeconomic groups, use dietary supplements [7, 10].

Dietary supplements are products intended to aid or supplement the diet and enhance nutritional status. They may include one or more ingredients such as minerals, vitamins, amino acids, metabolites, herbs, or extracts. Since they are generally taken orally, they are mostly available in the form of tablets, capsules, powders, or liquids. Dietary supplements are not suggested alone for treatment of a disease, but they should act synergistically with other treatments to facilitate healing or recovery. Though dietary ingredients may show certain effects in preclinical and clinical settings, the evidence may not be clinically significant in clinical trials [10-12]. Thus, clinical studies and meta-analyses should be performed to prove the effectiveness of a dietary supplement. Moreover, different factors must be considered in selecting dietary supplements, among which purity of the supplement, the patient's overall lifestyle (such as dietary habits and exercise), other health-associated conditions of

the patient (such as concomitant diseases and nutritional status), accurate dosage, food–drug interactions, absorption profiles, and potential side effects are the most relevant [7].

The U.S. Dietary Supplement Health and Education Act of 1994 deregulated the dietary supplement industry. The Office of Dietary Supplements (ODS) at the National Institutes of Health (NIH) has strengthened and enhanced understanding and knowledge of dietary supplements through the evaluation of scientific information, supporting scientific research and public education [13]. A product integrity profile is required for research projects funded by the ODS, including a collaborative approach to documentation of the commercial product. The Obesity Society believes that it is useful to conduct a qualitative analysis of non-FDA approved therapies to provide valid scientific evidence for the guidance of members [10].

Researchers have shown keen interest in the molecular and theoretical mechanisms of bioactive ingredients and the weight-loss effects of commonly used dietary supplements [7]. Table I and following paragraphs report a non-exhaustive list of dietary supplements with evidence supporting their use in weight loss.

## β-Glucans

Glucans are soluble polysaccharide fibers derived from D-glucose and classified by their α or β interchain linkage. β-Glucans consist of D-glucose monomers linked together by beta-glycosidic bonds and are arranged in linear β1-3,1-4-D-glucan structure. Glucans are mostly

found in cell walls of cereal grain endosperm and are also a major structural component of mushroom cell walls. As β-glucans are indigestible and readily fermented by gut microbiota in the colon and small intestine, it has been suggested that they could have a prebiotic role; they do not have significant side effect [14, 15, 52]. Their weight-loss effect is attributed to their being soluble fibers that may increase satiety due to the total time they take to move through the gastrointestinal track, and to their ability to reduce absorption of glucose. Their weight-loss effects were discovered as secondary outcomes of clinical trials designed to evaluate the effects of β-glucans on health conditions like insulin resistance, high blood pressure and dyslipidemia [14, 15].

## Bitter orange

Bitter orange, *Citrus aurantium* or Seville orange, has been used in various traditional South American and Chinese folk medicines for different health conditions [53]. Bitter orange extract has been used as a supplement to treat obesity and to enhance exercise performance. It contains phytochemicals such as octopamine, alkaloids and particularly synephrine. Several studies suggest there is a positive effect of bitter orange extract on weight loss [16, 53-55]. In a clinical trial by Kaats et al., a single dose of synephrine alone or synephrine and flavonoids combined both increased the basal metabolic rate in humans. Synephrine derived from *C. aurantium* is safe and no side effects were observed when up to 98 mg/day was taken for 60 days [16].

Tab. I. Non-exhaustive list of dietary supplements with evidence supporting their use in weight loss.

| Ingredient                 | Mechanism of action  | References |
|----------------------------|--|------------|
| β-Glucans                  | Reduce appetite and glucose absorption   | [14, 15]   |
| Bitter orange              | Increase metabolic rate  | [16]       |
| Calcium-vitamin D          | Improve metabolism, regulate triglyceride storage and adipocyte lipid metabolism | [17]       |
| Chitosan                   | Reduce fat absorption  | [18]       |
| Chromium                   | Increase insulin sensitivity   | [19, 20]   |
| Cocoa                      | Stimulate thermogenesis and lipid catabolism                                     | [21]       |
| <i>Coleus forskohlii</i>   | Increase lipolysis   | [22, 23]   |
| Conjugated linoleic acid   | Increase lipolysis   | [24, 25]   |
| <i>Ephedra sinica</i>      | Increase metabolic rate  | [26]       |
| Fucoxanthin                | Reduce lipogenesis and increases thermogenesis                                   | [27-31]    |
| <i>Garcinia cambogia</i>   | Reduce lipogenesis   | [32, 33]   |
| Glucomannan                | Reduce appetite and fat absorption   | [34-36]    |
| Green coffee               | Increase lipolysis   | [37-40]    |
| Green tea                  | Reduce appetite and increases thermogenesis                                      | [26, 41]   |
| Guar gum                   | Reduce appetite  | [42]       |
| <i>Hoodia gordonii</i>     | Reduce appetite.   | [43-45]    |
| <i>Irvingia gabonensis</i> | Reduce lipogenesis   | [46, 47]   |
| Raspberry                  | Reduce lipogenesis and increases lipolysis                                       | [48]       |
| Phenylpropylamine          | Reduce appetite  | [49]       |
| Pyruvate                   | Reduce appetite and fatigue, increases glucose uptake my skeletal muscles        | [50]       |
| White kidney bean          | Reduce glucose absorption  | [51]       |

## Calcium-vitamin D supplementation

Calcium is an essential nutrient/mineral often associated with healthy teeth and bones. It is also required for muscle, heart, and nerve function and for blood clotting. Vitamin D helps the body absorb calcium after conversion to calcitriol in the kidneys. Combined supplementation of calcium and vitamin D may improve metabolic health, regulate triglyceride storage and adipocyte lipid metabolism, and reduce body weight. Three different studies have reported significant weight loss over time in adults treated with calcium, without any stated side effect [17].

## Chitosan

Chitosan is a natural marine polysaccharide fiber derived from insect exoskeletons and shells of crustaceans like lobster, shellfish etc. [10]. Although it does not occur naturally in human tissues, it appears to be biocompatible, nontoxic, biodegradable and nonimmunogenic [7]. It is used in wound dressings to stop or reduce bleeding, and orally to decrease absorption of lipids in the gastrointestinal tract and reduce body weight [18]. Its mechanism of action for weight loss may involve binding fat molecules in the intestine, preventing their absorption. Chitosan is an insoluble animal fiber that exerts a bile acid sequestration or resin effect, decreasing absorption of cholesterol. It is currently available over the counter for treatment of conditions like obesity, hypertension and hypercholesterolemia [56]. A meta-analysis by Ernst and Pittler showed a statistically significant loss of weight (2.38 kg) after chitosan supplementation for 28 days [57]. Five additional studies on the effects of chitosan supplementation on weight loss reported statistically significant changes between groups, while others reported a weight reduction (2.3 kg) after 6 weeks of supplementation [10, 58].

A Cochrane meta-analysis of 13 clinical trials on chitosan reported a statistically significant weighted mean difference in body weight (1.7 kg) between the chitosan supplement and placebo groups. However, when only higher quality trials were analyzed, the average weight loss fell to 0.6 kg, which was, however, still statistically significant [59].

## Chromium

Chromium is considered an essential trace mineral, present in small amounts in different foods. It is also taken as a supplement. Chromium plays a significant role in the metabolism of amino acids, glucose, and lipids through its effects on insulin. It may directly increase the activity of serotonin and regulate its downstream impact on dopaminergic signaling on central insulin receptors. Chromium is considered to affect numerous pathways involving the central control of satiety, energy homeostasis and food intake by modulating these neurotransmitters [19,

20]. Chromium has been shown to reduce body weight while maintaining lean mass, thus it is favored by manufacturers of dietary supplements for weight loss [60].

## Cocoa

The cocoa bean or cocoa seed is the dried and fermented seed from the fruit of *Theobroma cacao*. These seeds are roasted and ground to obtain cocoa. Cocoa beans contain substantial amounts of various bioactive compounds, such as methylxanthines and antioxidant polyphenols (theobromine and caffeine) that may help weight loss by converting white adipocytes into brown adipocytes and enhancing lipid catabolism and endothelial function, while reducing insulin resistance and oxidative stress [21].

A randomized controlled clinical trial proved that a combination of catechins and cocoa reduces food intake without any side effect in young adults [61], while a second blinded placebo-controlled study proved that cocoa by-products reduced body weight in adults [62].

## *Coleus forskohlii*

Forskolin is a bioactive compound extracted from the root of *Coleus forskohlii*, a relative of the mint family. *Coleus forskohlii* is native to India and has been used in Ayurvedic medicine for centuries for the treatment of various health conditions, like respiratory disorders, heart disease and abdominal colic. Forskolin is a strong cAMP stimulator that activates hormone-sensitive lipase, causing release of fatty acids from adipose tissue. A few limited clinical trials have examined its effect on weight loss and found a more significant effect in males than females. A placebo-controlled, double-blind clinical trial involving 15 obese males treated with 500 mg/day forskolin extract (10%) for 12 weeks found a significant reduction in body fat in the forskolin group and an increase in lean body mass without any stated side effect, although the basal metabolic rate remained unchanged [22, 23, 63].

## Conjugated linoleic acid

Conjugated linoleic acid is a natural essential omega-6 fatty acid derived from linoleic acid, found mostly in meat and dairy products. Animal studies have shown its many beneficial effects including immune enhancement, reduction of atherosclerosis biomarkers and altered body composition (lower fat mass, higher lean mass). In humans, dietary conjugated linoleic acid supplements may improve insulin sensitivity and lipid metabolism by decreasing plasma levels of triglycerides and low-density cholesterol [24, 64]. Conjugated linoleic acid proved to reduce hunger in adults, with no remarkable side effects [25].

## ***Ephedra sinica***

*Ephedra sinica* is a plant that occurs naturally in Asia, although it is cultivated in other parts of the world. For thousands of years, *Ephedra sinica* has been used in Chinese medicine. Ephedrine is the bioactive ingredient of *Ephedra sinica* associated with weight loss; it is mostly taken with caffeine. In clinical trials, ephedra and ephedrine demonstrate only short-term weight-loss effects. The proposed mechanism of action involves an increase in the metabolic rate and stimulation of fat burning. For these reasons, ephedrine is used in weight-loss supplements [26].

## **Fucoxanthin**

Fucoxanthin is a marine carotenoid widespread in nature and mostly isolated from diatoms and seaweeds. Several preclinical trials have investigated a mechanism of action of fucoxanthin in the treatment of obesity and associated cardiometabolic alterations. Fucoxanthin decreases plasma and hepatic concentrations of triglycerides. It also reduces expression of acetyl-CoA carboxylase, thus decreasing malonyl-CoA formation, fatty acid synthase expression and saturated long-chain fatty acid synthesis [27-29].

Research studies have reported that fucoxanthin may downregulate low density lipoprotein receptor expression in the liver [28]. It also downregulates expression of the *C/EBP $\alpha$*  (CCAAT/enhancer-binding protein-alpha), *PPAR- $\gamma$*  (peroxisome proliferator-activated receptor gamma) and *SREBP-1c* (sterol regulatory element-binding protein 1c) genes during intermediate to late adipocyte differentiation stages, whereas during the initial adipocyte differentiation stages, fucoxanthin enhances expression of the proteins *C/EBP $\alpha$* , *PPAR- $\gamma$* , A-FABP (adipocyte fatty acid-binding protein), *SREBP-1c* (sterol-regulatory element binding protein-1C), glucose transporter 4 (GLUT4) and lipoprotein lipase [30]. Similarly fucoxanthin has been known to stimulate UCP-1 (uncoupling protein1) expression in white adipocytes, thereby increasing energy expenditure and thermogenesis [31].

In a study of 151 obese premenopausal women who were given supplements containing fucoxanthin from brown seaweed and pomegranate seed oil extracts at various doses for almost 16 weeks, the group receiving 300 mg pomegranate seed oil and 300 mg seaweed extract (2.4 mg fucoxanthin) had a statistically significant decrease in waist circumference, body weight and body fat, without any side effects. Moreover, the group that received more than 2.4mg fucoxanthin showed an increase in resting energy expenditure compared to the placebo [65].

## ***Garcinia cambogia***

*Garcinia cambogia* is a plant native to Asia, Polynesia, Africa, and Australia. It has various significant anti-in-

flammatory, antineoplastic, hypolipidemic and anti-diabetic effects on the body, and its extracts have anorexic effects. *Garcinia* has also been analyzed for weight management. Its rind is high in hydroxycitric acid, believed to be a bioactive component that causes weight loss by inhibiting extra-mitochondrial citrate lyase or ATP-citrate lyase that influences the synthesis of cholesterol and fatty acids. As ATP-citrate lyase is the major enzyme involved in the synthesis and storage of fatty acid in cells, hydroxycitric acid may inhibit lipogenesis [10, 32, 33].

## **Glucosamin**

Glucosamin is a water-soluble polysaccharide dietary fiber that is easily extracted in large amounts from softwoods, tubers, plant bulbs and roots. Most glucosamin is extracted from the tuber of konjac (*Amorphophallus konjac*), an Asian plant for use as an herbal remedy. Glucosamin high molecular weight polysaccharide is composed of  $\beta$ -(1-4)-linked D-mannose and D-glucose monomers and is considered a soluble fiber. This highly viscous dietary fiber can absorb 50 times its weight in water. Since human salivary and pancreatic amylase cannot break  $\beta$ -1,4 glycosidic bonds, glucosamin reaches the colon almost unchanged and is fermented by gut microbiota. Glucosamin is lately being analyzed for its beneficial effects on weight loss, blood glucose, and dyslipidemia, amongst other uses [34, 35]. Some clinical trials question its effectiveness in weight loss [35, 66]. Thus, new studies will be needed to confirm glucosamin positive action on weight reduction.

Various mechanisms of action could explain the effects of glucosamin on weight loss: it may cause satiety through greater mastication effort, by prolonging gastric emptying, and by shortening the time food remains in the small intestine. Fecal energy loss may be another mechanism of action, as soluble fiber decreases absorption of fats and protein. Sood et al. reported statistically significant weight loss (-0.79 kg) after almost 5.2 weeks in humans taking glucosamin [36].

## **Green coffee**

Green coffee extract is obtained from unroasted green coffee beans and marketed in caffeinated and decaffeinated forms. The bioactive ingredients of green coffee extract include chlorogenic acid, a polyphenol of the phenolic acid subfamily [67]. Different mechanisms of action of green coffee extract on weight loss may be caused by a reduction in pancreatic lipase activity, a lipolytic effect on adipocytes, inhibition of hydroxymethylglutaryl-CoA (HMG-CoA) reductase, acyl-CoA-cholesterol acyltransferase (ACAT) and fatty acid synthase (FASN), an increase in  $\beta$ -oxidation and increased expression of *PPAR- $\alpha$*  in liver, a nutrient sensor that regulates genes important in peroxisomal and mitochondrial  $\beta$ -oxidation, and fatty acid transport. A meta-analysis



reported statistically significant weight loss after green coffee extract supplementation (180-200 mg/day) for a period of 4-12 weeks, with no side effects [37-40].

## Green tea

Green tea is made from the steamed and pan-fried unfermented (unoxidized) leaves of *Camellia sinensis*. Green tea has caffeine that is thought to contribute to the suppression of appetite and stimulation of thermogenesis. Antioxidants in green tea like the catechin epigallocatechin-3-gallate, inhibit norepinephrine breakdown that in turn causes an increase in calories burned [26, 41]. A randomized, double-blind, placebo-controlled clinical trial reported that green tea extract reduces body weight without side effects on more than 100 women [68].

## Guar gum

Guar gum, a dietary fiber obtained from seeds of the plant *Cyamopsis tetragonolobus*, is used in various food products particularly as a thickener and emulsifier in baked items [10]. It consists of high molecular weight galactomannan polysaccharides in linear,  $\beta$ -1,4-linked D-mannopyranosyl chains with  $\alpha$ -1,6 D-galactopyranosyl side chains [69]. Guar gum may cause weight loss by virtue of its bulking characteristics in the gut which delay gastric emptying [7, 70]. Guar gum supplements have therefore been used to reduce food intake and to increase satiety, but a meta-analysis did not sustain its effectiveness, and side effects were reported including diarrhea, flatulence, and cramps [42, 70].

## *Hoodia gordonii*

*Hoodia gordonii*, also known as Bushman's hat, is a leafless succulent plant with medicinal properties occurring naturally in Namibia, Botswana, and South Africa. Researchers have studied *Hoodia gordonii* as a weight-loss adjuvant due to its appetite suppressing properties, although its use was questioned due to several adverse effects, among which were nausea and skin reactions [45]. Metabolites such as pregnane glycosides that contain 6-deoxy and 2,6-dideoxy sugars have been isolated from *Hoodia gordonii*. While the active compound causing the anorexigenic effect of *Hoodia gordonii* is not yet clear, P57AS3 (P57), an oxypregnane steroidal glycoside, is commonly considered to be the metabolite responsible for these effects [43-45].

Research into the effects of P57 in vivo have shown that intraventricular injection of purified P57 in rats decreases food intake and significantly increases hypothalamic ATP production, which may decrease the appetite response. According to an in-vitro study, P57 stimulates cholecystokinin secretion in human enteroendocrine cells, while cholecystokinin has been studied for insights into its appetite-suppressing effect through

the vagus nerve. However, since oral administration of *Hoodia gordonii* is known to cause gastric breakdown of P57, the extract must be taken in high doses to obtain significant clinical effects [45, 69, 71], or easy to absorb derivatives developed.

## *Irvingia gabonensis*

*Irvingia gabonensis*, also known as African wild mango, is a native to western and central Africa. Its seed is high in saturated fatty acids and its flesh is rich in polyphenols, specifically flavonoids. Oben et al. treated adipocyte cultures with *Irvingia gabonensis* extract and observed inhibition of PPAR- $\gamma$  (peroxisome proliferator-activated receptor gamma) expression, a decrease in leptin protein levels and upregulation of adiponectin expression, thus inhibiting lipogenesis [46, 47]. Other randomized double-blind clinical trials analyzed the weight loss effects of *Irvingia gabonensis* extracts [72, 73]. Most of these studies were systematically reviewed by Onakpoya et al., who reported that administration of 200-3150 mg/day *Irvingia gabonensis* extract for 4-10 weeks could lead to statistically and clinically significant weight loss and decreased waist circumference compared to a placebo group. Side effects of the extract included headache and sleep difficulty [74].

*Irvingia gabonensis* emerged from these studies as a useful adjuvant dietary supplement in weight reduction management. However, many of these clinical studies were conducted on relatively small samples of black people from Africa. The research should be extended to larger and more diverse populations [72].

## Raspberry

Raspberry ketones [4-(4-hydroxyphenyl)-2-butanone] are natural aromatic substances found, for example, in red raspberries and rhubarb. Extracted from raspberries, they are used as flavoring in the food industry. Several *in-vitro* studies on the effect of these ketones on adipocytes have reported increased fatty acid oxidation, reduced lipid accumulation and enhanced secretion of adiponectin [48]. Molecular studies have established that raspberry ketones downregulate expression of various genes associated with adipogenesis, such as C/EBP $\alpha$ , PPAR- $\gamma$ , ACC1 (acetyl-CoA carboxylase1), SCD1 (steroyl-CoA desaturase1), A-FABP 2 (adipocyte fatty acid-binding protein 2) and FAS (fatty acid synthase), whereas it increased the mRNA levels of genes involved in the process of fatty acid oxidation, such as HSL (hormone-sensitive lipase), CPT (carnitine palmitoyl transferase) and ATGL (adipose triglyceride lipase) was observed [75]. Moreover, in-vivo studies on rodents fed a high-fat diet have shown that raspberry ketones prevented any increase in body weight or visceral adipose tissue. The mechanism of this effect presumably involves stimulation of brown and white adipose tissue and inhibition of pancreatic lipase activity [76].

These properties suggest that raspberry ketone supplementation could be an alternative weight-loss measure, but sufficient clinical evidence is lacking and certain teratogenic and cardiotoxic effects of raspberry ketones identified by in-silico research, indicate the need for further detailed study [77].

## Phenylpropylamine

Phenylpropylamine (PPA) is a sympathomimetic agent that has structural similarity to ephedrine and amphetamine. It may act as a decongestant and appetite suppressant and induce significant weight loss. It is used as an over-the-counter supplement for weight loss. Researchers believe that PPA acts through the  $\alpha$ -1 adrenergic receptor ( $\alpha$ 1-AR) [49]. In a double-blind placebo-controlled study, Scheingart administered a 1200 kcal diet and 75 mg/day PPA to 101 overweight subjects for 6 to 20 weeks and reported greater weight loss (2.59 kg) in the PPA-supplemented group than in the placebo group, without untoward side effects [78].

## Pyruvate

Pyruvate is the simplest alpha-keto acid having a carboxylic and a ketone functional group. It is the end product of glycolysis but is also obtained from other sources. It breaks down glucose and produces energy and is also a key intermediate in several cell metabolic pathways. Pyruvate helps lipid metabolism by reversible conversion to phosphoenolpyruvate, increasing glucose uptake by skeletal muscle. It may therefore promote weight loss [50]. By virtue of these properties, pyruvate has been used to promote weight loss in subjects with obesity, to improve athletic performance, and in the treatment of cataracts, high cholesterol and cancer. A meta-analysis which analyzed six clinical trials concluded that pyruvate is effective in reducing body weight, and that adverse effects include gas, bloating and diarrhea [79]. Pyruvate should be administered with thiamine, a cofactor of pyruvate dehydrogenase, an essential enzyme that allows pyruvate to enter the Krebs's cycle [80].

## White kidney bean

Extract of white kidney bean (*Phaseolus vulgaris* L.) contains alpha-amylase inhibitor (phaseolamin) that inhibits pancreatic amylase activity, lowering glycaemia and calorie absorption by delaying or preventing complex carbohydrate digestion. It is therefore used as an over-the-counter dietary supplement for weight loss [51]. Several proof-of-concept clinical trials have established a dose-dependent reduction in glucose absorption after supplementation with white kidney bean. However, weight-loss effects were more prominent in some relatively small, short-term studies that used 1.5-6 g/day of white kidney bean extract. For instance, a

clinical trial involving 60 subjects with obesity reported greater reductions in fat mass, body weight and thigh, hip and waist circumference in the group taking white kidney bean extract than in the placebo group, without any side effect [81, 82].

## Conclusions

Obesity is a prevalent nutritional disease and a significant public health issue in industrialized countries. Due to their low toxicity profile, dietary supplements are considered an attractive alternative to traditional therapy, which usually comprise surgical and pharmacological treatments. Dietary supplements for weight loss or obesity managements may act through several mechanisms, among which are reduction of lipogenesis, appetite, and nutrient absorption, or increasing lipolysis and energy expenditure. Among the many ingredients presented in this study, chitosan and green tea showed promising results with limited side effects. Weight loss and absence of side effects were reported for both ingredients by meta-analysis. More clinical studies on chitosan and green tea, as well as other promising ingredients, will be needed to increase our knowledge on the efficacy and on the mode of actions of dietary supplements for weight loss and obesity management.

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## Conflicts of interest statement

Authors declare no conflict of interest.

## Author's contributions

Gabriele Bonetti, Karen L. Herbst: These authors contributed equally to this work.

MB: study conception, editing and critical revision of the manuscript; GB, KLH, Kevin D, Kristjana D, AKK, BA, VV, GM, AI: literature search, editing and critical revision of the manuscript. All authors have read and approved the final manuscript.

## References

- [1] Oppert JM. Obesity: Epidemiology, Pathophysiology and Extra-Respiratory Complications. *Rev Pneumol Clin* 2002;58:63-70.
- [2] Vettori A, Paolacci S, Maltese PE, Herbst KL, Cestari M, Michelini S, Michelini S, Samaja M, Bertelli M. Genetic determinants of the effects of training on muscle and adipose tissue homeostasis in obesity associated with lymphedema. *Lymphat Res Biol* 2021;19:322-33. <https://doi.org/10.1089/lrb.2020.0057>

- [3] Vettori A, Pompucci G, Paolini B, del Ciondolo I, Bressan S, Dunder M, Kenanoglu S, Unfer V, Bertelli M. Geneob Project genetic background, nutrition and obesity: a review. *Eur Rev Med Pharmacol Sci* 2019;23:1751-61. [https://doi.org/10.26355/eurev\\_201902\\_17137](https://doi.org/10.26355/eurev_201902_17137)
- [4] Obesity: Preventing and Managing the Global Epidemic. Report of a WHO Consultation. *World Health Organ Tech Rep Ser* 2000;894:1-253.
- [5] Pagano C, Marin O, Calcagno A, Schiappelli P, Pilon C, Milan G, Bertelli M, Fanin E, Andrighetto G, Federspil G, Vettor R. Increased serum resistin in adults with Prader-Willi Syndrome is related to obesity and not to insulin resistance. *J Clin Endocrinol Metab* 2005;90:4335-40. <https://doi.org/10.1210/jc.2005-0293>
- [6] Camilleri G, Kiani AK, Herbst KL, Kaftalli J, Bernini A, Dhuli K, Manara E, Bonetti G, Stuppia L, Paolacci S, Dautaj A, Bertelli M. Genetics of fat deposition. *Eur Rev Med Pharmacol Sci* 2021;25:14-22. [https://doi.org/10.26355/eurev\\_202112\\_27329](https://doi.org/10.26355/eurev_202112_27329)
- [7] Ríos-Hoyo A, Gutiérrez-Salmeán G. New dietary supplements for obesity: what we currently know. *Curr Obes Rep* 2016;5:262-70. <https://doi.org/10.1007/s13679-016-0214-y>
- [8] Bhutani S, vanDellen MR, Cooper JA. Longitudinal weight gain and related risk behaviors during the COVID-19 Pandemic in adults in the US. *Nutrients* 2021;13:671. <https://doi.org/10.3390/nu13020671>
- [9] Dhuli K, Ceccarini MR, Precone V, Maltese PE, Bonetti G, Paolacci S, Dautaj A, Guerri G, Marceddu G, Beccari T, Michelini S, Bertelli M. Improvement of quality of life by intake of hydroxytyrosol in patients with lymphedema and association of lymphedema genes with obesity. *Eur Rev Med Pharmacol Sci* 2021;25:33-42. [https://doi.org/10.26355/eurev\\_202112\\_27331](https://doi.org/10.26355/eurev_202112_27331)
- [10] Batsis JA, Apolzan JW, Bagley PJ, Blunt HB, Divan V, Gill S, Golden A, Gundumraj S, Heymsfield SB, Kahan S, Kopatsis K, Port A, Prout Parks E, Reilly CA, Rubino D, Saunders KH, Shean R, Tabaza L, Stanley A, Tchang BG, Gundumraj S, Kidambi S. A Systematic Review of Dietary Supplements and Alternative Therapies for Weight Loss. *Obesity* 2021;29:1102-13. <https://doi.org/10.1002/oby.23110>
- [11] Gregg EW. Secular trends in cardiovascular disease risk factors according to body mass index in US adults. *JAMA* 2005;293:1868. <https://doi.org/10.1001/jama.293.15.1868>
- [12] Walter S, Kunst A, Mackenbach J, Hofman A, Tiemeier H. Mortality and disability: the effect of overweight and obesity. *Int J Obes* 2009;33:1410-8. <https://doi.org/10.1038/ijo.2009.176>
- [13] Office of Dietary Supplements. Mission, Origin, and Mandate. Available at: <https://ods.od.nih.gov/About/MissionOriginMandate.aspx#:~:text=The%20mission%20of%20ODS%20is,health%20for%20the%20U.S.%20population>. Accessed on: 02/07/2022.
- [14] Cloetens L, Ulmius M, Johansson-Persson A, Åkesson B, Önnig G. Role of dietary Beta-Glucans in the prevention of the Metabolic Syndrome. *Nutr Rev* 2012;70:444-58. <https://doi.org/10.1111/j.1753-4887.2012.00494.x>
- [15] el Khoury D, Cuda C, Luhovyy BL, Anderson GH. Beta Glucan: health benefits in Obesity and Metabolic Syndrome. *J Nutr Metab* 2012;1-28. <https://doi.org/10.1155/2012/851362>
- [16] Kaats GR, Miller H, Preuss HG, Stohs SJ. A 60day double-blind, placebo-controlled safety study involving Citrus Aurantium (Bitter Orange) extract. *Food Chem Toxicol* 2013;55:358-62. <https://doi.org/10.1016/j.fct.2013.01.013>
- [17] Zhu W, Cai D, Wang Y, Lin N, Hu Q, Qi Y, Ma S, Amarasekara S. Calcium plus Vitamin D3 supplementation facilitated fat loss in overweight and obese college students with very-low calcium consumption: a randomized controlled trial. *Nutr J* 2013;12:8. <https://doi.org/10.1186/1475-2891-12-8>
- [18] Patrulea V, Ostafe V, Borchard G, Jordan O. Chitosan as a starting material for wound healing applications. *European Eur J Pharm Biopharm* 2015;97:417-26. <https://doi.org/10.1016/j.ejpb.2015.08.004>
- [19] Brownley KA, Boettiger CA, Young L, Cefalu WT. Dietary chromium supplementation for targeted treatment of diabetes patients with comorbid depression and binge eating. *Med Hypotheses* 2015;85:45-8. <https://doi.org/10.1016/j.mehy.2015.03.020>
- [20] Vincent JB, Lukaski HC. Chromium. *Advances Nutr* 2018;9:505-6. <https://doi.org/10.1093/advances/nmx021>
- [21] Barišić V, Kopjar M, Jozinović A, Flanjak I, Ačkar Đ, Miličević B, Šubarić D, Jokić S, Babić J. The chemistry behind chocolate production. *Molecules* 2019;24:3163. <https://doi.org/10.3390/molecules24173163>
- [22] Kavtiha C, Rajamani K, Vadivel E. Select record coleus forskohlii - a comprehensive review on morphology, phytochemistry and pharmacological aspects. *J Med Plant Res* 2010;4:278-85.
- [23] Godard MP, Johnson BA, Richmond SR. Body composition and hormonal adaptations associated with forskolin consumption in overweight and obese men. *Obes Res* 2005;13:1335-43. <https://doi.org/10.1038/oby.2005.162>
- [24] Abenhaim L, Moride Y, Brenot F, Rich S, Benichou J, Kurz X, Higenbottam T, Oakley C, Wouters E, Aubier M, Simonneau G, Bégaud B. Appetite-suppressant drugs and the risk of primary pulmonary hypertension. *N Engl J Med* 1996;335:609-16. <https://doi.org/10.1056/NEJM199608293350901>
- [25] Kamphuis MMJW, Lejeune MPGM, Saris WHM, Westerterp-Plantenga MS. Effect of conjugated linoleic acid supplementation after weight loss on appetite and food intake in overweight subjects. *Eur J Clin Nutr* 2003;57:1268-74. <https://doi.org/10.1038/sj.ejcn.1601684>
- [26] Diepvens K, Westerterp KR, Westerterp-Plantenga MS. Obesity and thermogenesis related to the consumption of caffeine, ephedrine, capsaicin, and green tea. *Am J Physiol Regul Integr Comp Physiol* 2007;292:R77-R85. <https://doi.org/10.1152/ajpregu.00832.2005>
- [27] Peng J, Yuan J-P, Wu C-F, Wang J-H. Fucoxanthin, a marine carotenoid present in brown seaweeds and diatoms: metabolism and bioactivities relevant to human health. *Mar Drugs* 2011;9:1806-28. <https://doi.org/10.3390/md9101806>
- [28] Hu X, Li Y, Li C, Fu Y, Cai F, Chen Q, Li D. Combination of fucoxanthin and conjugated linoleic acid attenuates body weight gain and improves lipid metabolism in high-fat diet-induced obese rats. *Arch Biochem Biophys* 2012;519:59-65. <https://doi.org/10.1016/j.abb.2012.01.011>
- [29] Gammone M, D'Orazio N. Anti-obesity activity of the marine carotenoid fucoxanthin. *Mar Drugs* 2015;13:2196-214. <https://doi.org/10.3390/md13042196>
- [30] Kang S-I, Ko H-C, Shin H-S, Kim H-M, Hong Y-S, Lee N-H, Kim S-J. Fucoxanthin exerts differing effects on 3t3-l1 cells according to differentiation stage and inhibits glucose uptake in mature adipocytes. *Biochem Biophys Res Commun* 2011;409:769-74. <https://doi.org/10.1016/j.bbrc.2011.05.086>
- [31] Maeda H, Hosokawa M, Sashima T, Funayama K, Miyashita K. Fucoxanthin from edible seaweed, undaria pinnatifida, shows antiobesity effect through UCP1 expression in white adipose tissues. *Biochem Biophys Res Commun* 2005;332:392-7. <https://doi.org/10.1016/j.bbrc.2005.05.002>
- [32] Heymsfield SB, Allison DB, Vasselli JR, Pietrobelli A, Greenfield D, Nunez C. Garcinia Cambogia (hydroxycitric acid) as a potential antiobesity agent. *JAMA* 1998;280:1596. <https://doi.org/10.1001/jama.280.18.1596>
- [33] Astell KJ, Mathai ML, Su XQ. Plant extracts with appetite suppressing properties for body weight control: a systematic review of double blind randomized controlled clinical trials. *Complement Ther Med* 2013;21:407-16. <https://doi.org/10.1016/j.ctim.2013.05.007>
- [34] Alonso-Sande M, Teijeiro-Osorio D, Remuñán-López C, Alonso MJ. Glucomannan, a promising polysaccharide for biopharmaceutical purposes. *Eur J Pharm Biopharm* 2009;72:453-62. <https://doi.org/10.1016/j.ejpb.2008.02.005>
- [35] Keithley JK, Swanson B, Mikolaitis SL, DeMeo M, Zeller



- JM, Fogg L, Adamji J. Safety and efficacy of glucomannan for weight loss in overweight and moderately obese adults. *J Obes* 2013;1-7. <https://doi.org/10.1155/2013/610908>
- [36] Sood N, Baker WL, Coleman CI. Effect of Glucomannan on plasma lipid and glucose concentrations, body weight, and blood pressure: systematic review and meta-analysis. *The Am J Clin Nutr* 2008;88:1167-75. <https://doi.org/10.1093/ajcn/88.4.1167>
- [37] Mullin GE. Supplements for weight loss. *Nutrition in Clinical Practice* 2015;30:311-2. <https://doi.org/10.1177/0884533615572655>
- [38] Flanagan J, Bily A, Rolland Y, Roller M. Lipolytic activity of svetol®, a decaffeinated green coffee bean extract. *Phytother Res* 2014;28:946-8. <https://doi.org/10.1002/ptr.5085>
- [39] Cho A-S, Jeon S-M, Kim M-J, Yeo J, Seo K-I, Choi M-S, Lee M-K. Chlorogenic acid exhibits anti-obesity property and improves lipid metabolism in high-fat diet-induced-obese mice. *Food Chem Toxicol* 2010;48:937-43. <https://doi.org/10.1016/j.fct.2010.01.003>
- [40] Onakpoya I, Terry R, Ernst E. The use of green coffee extract as a weight loss supplement: a systematic review and meta-analysis of randomised clinical trials. *Gastroenterol Res Pract* 2011;1-6. <https://doi.org/10.1155/2011/382852>
- [41] Carrasco-Pozo C, Cires MJ, Gotteland M. Quercetin and epigallocatechin gallate in the prevention and treatment of obesity: from molecular to clinical studies. *J Med Food* 2019;22:753-70. <https://doi.org/10.1089/jmf.2018.0193>
- [42] Pasman W, Westerterp-Plantenga M, Muls E, Vansant G, van Ree J, Saris W. The effectiveness of long-term fibre supplementation on weight maintenance in weight-reduced women. *Int J Obes* 1997;21:548-55. <https://doi.org/10.1038/sj.ijo.0800439>
- [43] Russell PJ, Swindells C. Chemical characterisation of hoodia gordonii extract. *Food Chem Toxicol* 2012;50:S6-S13. <https://doi.org/10.1016/j.fct.2011.02.020>
- [44] van Heerden FR. Hoodia gordonii: a natural appetite suppressant. *J Ethnopharmacol* 2008;119:434-7. <https://doi.org/10.1016/j.jep.2008.08.023>
- [45] Smith C, Krygsman A. Hoodia Gordonii: to eat, or not to eat. *J Ethnopharmacol* 2014;155:987-91. <https://doi.org/10.1016/j.jep.2014.06.033>
- [46] Oben JE, Ngondi JL, Blum K. Inhibition of Irvingia Gabonensis seed extract (OB131) on adipogenesis as mediated via down regulation of the PPARgamma and leptin genes and up-regulation of the adiponectin gene. *Lipids Health Dis* 2008;7:44. <https://doi.org/10.1186/1476-511X-7-44>
- [47] Yamoneka J, Malumba P, Blecker C, Gindo M, Richard G, Fauconnier M-L, Lognay G, Danthine S. Physicochemical properties and thermal behaviour of african wild mango (*Irvingia Gabonensis*) seed fat. *LWT - Food Science and Technology* 2015;64:989-96. <https://doi.org/10.1016/j.lwt.2015.06.035>
- [48] Park K. Raspberry ketone increases both lipolysis and fatty acid oxidation in 3T3-L1 adipocytes. *Planta Med* 2010;76:1654-8. <https://doi.org/10.1055/s-0030-1249860>
- [49] Greenway F, Heber D, Raum W, Morales S. Double-blind, randomized, placebo-controlled clinical trials with non-prescription medications for the treatment of obesity. *Obes Res* 1999;7:370-8. <https://doi.org/10.1002/j.1550-8528.1999.tb00420.x>
- [50] Miller AT, Thomas BM. Pyruvate metabolism in obesity. *Am J Clin Nutr* 1956;4:619-24. <https://doi.org/10.1093/ajcn/4.6.619>
- [51] Obiro WC, Zhang T, Jiang B. The nutraceutical role of the phaseolus vulgaris  $\alpha$ -Amylase inhibitor. *Br J Nutr* 2008;100:1-12. <https://doi.org/10.1017/S0007114508879135>
- [52] Sima P, Vannucci L, Vetvicka V.  $\beta$ -Glucans and Cholesterol (Review). *Int J Mol Med* 2018. <https://doi.org/10.3892/ijmm.2018.3411>
- [53] Stohs SJ, Preuss HG, Shara M. The safety of citrus aurantium (Bitter Orange) and its primary protoalkaloid p -Synephrine. *Phytother Res* 2011;25:1421-8. <https://doi.org/10.1002/ptr.3490>
- [54] Allison DB, Cutter G, Poehlman ET, Moore DR, Barnes S. Exactly which synephrine alkaloids does citrus aurantium (Bitter Orange) contain? *Int J Obes* 2005;29:443-6. <https://doi.org/10.1038/sj.ijo.0802879>
- [55] Bitter Orange. In: *Drugs and Lactation Database (LactMed)*. Bethesda (MD): National Library of Medicine (US) 2021. Available at: <https://pubmed.ncbi.nlm.nih.gov/30000952/>. Accessed on: 12/08/2022.
- [56] van Bennekum AM, Nguyen Dv, Schulthess G, Hauser H, Phillips MC. Mechanisms of cholesterol-lowering effects of dietary insoluble fibres: relationships with intestinal and hepatic cholesterol parameters. *Br J Nutr* 2005;94:331-7. <https://doi.org/10.1079/BJN20051498>
- [57] Ernst E, Pittler M. Chitosan as a treatment for body weight reduction: a meta-analysis. Available at: <https://www.ncbi.nlm.nih.gov/books/NBK67453/#:~:text=The%20meta%2Danalysis%20implies%20that,raised%20about%20the%20original%20studies>. Accessed on: 12/08/2022.
- [58] Woodgate DE, Conquer JA. Effects of a stimulant-free dietary supplement on body weight and fat loss in obese adults: a six-week exploratory study. *Curr Ther Res Clin Exp* 2003;64:248-62. [https://doi.org/10.1016/S0011-393X\(03\)00058-4](https://doi.org/10.1016/S0011-393X(03)00058-4)
- [59] Pereira A, Guedes AD, Verreschi ITN, Santos RD, Martinez TLR. A obesidade e sua associação com os demais fatores de risco cardiovascular em escolares de itapetinga, Brasil. *Arq Bras Cardiol* 2009;93. <https://doi.org/10.1590/S0066-782X2009000900009>
- [60] Willoughby D, Hewlings S, Kalman D. Body composition changes in weight loss: strategies and supplementation for maintaining lean body mass, a brief review. *Nutrients* 2018;10:1876. <https://doi.org/10.3390/nu10121876>
- [61] Greenberg JA, O'Donnell R, Shurpin M, Kordunova D. Epicatechin, procyanidins, cocoa, and appetite: a randomized controlled trial. *Am J Clin Nutr* 2016;104:613-9. <https://doi.org/10.3945/ajcn.115.129783>
- [62] León-Flores P, Nájera N, Pérez E, Pardo B, Jimenez F, Diaz-Chiguer D, Villarreal F, Hidalgo I, Ceballos G, Meaney E. Effects of cacao by-products and a modest weight loss intervention on the concentration of serum triglycerides in overweight subjects: proof of concept. *J Med Food* 2020;23:745-9. <https://doi.org/10.1089/jmf.2019.0201>
- [63] Insel PA. Forskolin as a tool for examining adenylyl cyclase expression, regulation, and G Protein Signaling. *Cell Mol Neurobiol* 2003;23:305-14. <https://doi.org/10.1023/A:1023684503883>
- [64] Lee KN, Kritchevsky D, Parizaa MW. Conjugated linoleic acid and atherosclerosis in rabbits. *Atherosclerosis* 1994;108:19-25. [https://doi.org/10.1016/0021-9150\(94\)90034-5](https://doi.org/10.1016/0021-9150(94)90034-5)
- [65] Abidov M, Ramazanov Z, Seifulla R, Grachev S. The effects of xanthigen™ in the weight management of obese premenopausal women with non-alcoholic fatty liver disease and normal liver fat. *Diabetes Obes Metab* 2010;12:72-81. <https://doi.org/10.1111/j.1463-1326.2009.01132.x>
- [66] Zalewski BM, Szajewska H. No effect of Glucomannan on body weight reduction in children and adolescents with overweight and obesity: a randomized controlled trial. *J Pediatr* 2019;211:85-91. <https://doi.org/10.1016/j.jpeds.2019.03.044>
- [67] Tsao R. Chemistry and biochemistry of dietary polyphenols. *Nutrients* 2010;2:1231-46. <https://doi.org/10.3390/nu2121231>
- [68] Chen I-J, Liu C-Y, Chiu J-P, Hsu C-H. Therapeutic effect of high-dose green tea extract on weight reduction: a randomized, double-blind, placebo-controlled clinical trial. *Clin Nutr* 2016;35:592-9. <https://doi.org/10.1016/j.clnu.2015.05.003>
- [69] MacLean DB, Luo L-G. Increased ATP content/production in the Hypothalamus may be a signal for energy-sensing of satiety: studies of the anorectic mechanism of a plant steroidal glycoside. *Brain Res* 2004;1020:1-11. <https://doi.org/10.1016/j.brainres.2004.04.041>



- [70] Pittler MH, Ernst E. Guar gum for body weight reduction: meta-analysis of randomized trials. *Am J Med* 2001;110:724-30. [https://doi.org/10.1016/S0002-9343\(01\)00702-1](https://doi.org/10.1016/S0002-9343(01)00702-1)
- [71] le Nevé B, Foltz M, Daniel H, Gouka R. The steroid Glycoside H.g.-12 from Hoodia Gordonii activates the human bitter receptor TAS2R14 and induces CCK release from HuTu-80 cells. *Am J Physiol Gastrointest Liver Physiol* 2010;299:G1368-G1375. <https://doi.org/10.1152/ajpgi.00135.2010>
- [72] Ngondi JL, Oben JE, Minka SR. The Effect of Irvingia Gabonensis seeds on body weight and blood lipids of obese subjects in Cameroon. *Lipids Health Dis* 2005;4:12. <https://doi.org/10.1186/1476-511X-4-12>
- [73] Oben JE, Ngondi JL, Momo CN, Agbor GA, Sobgui C. The use of a Cissus Quadrangularis/Irvingia Gabonensis combination in the management of weight loss: a double-blind placebo-controlled study. *Lipids Health Dis* 2008;7:12. <https://doi.org/10.1186/1476-511X-7-12>
- [74] Onakpoya I, Davies L, Posadzki P, Ernst E. The efficacy of Irvingia Gabonensis supplementation in the management of overweight and obesity: a systematic review of randomized controlled trials. *J Diet Suppl* 2013;10:29-38. <https://doi.org/10.3109/19390211.2012.760508>
- [75] Park KS. Raspberry Ketone, a naturally occurring phenolic compound, inhibits adipogenic and lipogenic gene expression in 3T3-L1 adipocytes. *Pharm Biol* 2015;53:870-5. <https://doi.org/10.3109/13880209.2014.946059>
- [76] Morimoto C, Satoh Y, Hara M, Inoue S, Tsujita T, Okuda H. Anti-obese action of Raspberry Ketone. *Life Sci* 2005;77:194-204. <https://doi.org/10.1016/j.lfs.2004.12.029>
- [77] Bredsdorff L, Wedebye EB, Nikolov NG, Hallas-Møller T, Pilegaard K. Raspberry Ketone in food supplements – high intake, few toxicity data – a cause for safety concern? *Regul Toxicol Pharmacol* 2015;73:196-200. <https://doi.org/10.1016/j.yrtph.2015.06.022>
- [78] Schteingart DE. Effectiveness of phenylpropranolamine in the management of moderate obesity. *Int J Obes Relat Metab Disord* 1992;16:487-93.
- [79] Onakpoya I, Hunt K, Wider B, Ernst E. Pyruvate supplementation for weight loss: a systematic review and meta-analysis of randomized clinical trials. *Crit Rev Food Sci Nutr* 2014;54:17-23. <https://doi.org/10.1080/10408398.2011.565890>
- [80] Jauhari P, Sankhyan N, Vyas S, Singhi P. Thiamine responsive pyruvate dehydrogenase complex deficiency: a potentially treatable cause of leigh's disease. *J Pediatr Neurosci* 2017;12:265-7. [https://doi.org/10.4103/jpn.JPN\\_191\\_16](https://doi.org/10.4103/jpn.JPN_191_16)
- [81] Vinson JA, al Kharrat H, Shuta D. Investigation of an amylase inhibitor on human glucose absorption after starch consumption. *Open Nutraceuticals J* 2009;2:88-91. <https://doi.org/10.2174/1876396000902010088>
- [82] Celleno L, Tolaini MV, D'Amore A, Perricone Nv, Preuss HG. A dietary supplement containing standardized phaseolus vulgaris extract influences body composition of overweight men and women. *Int J Med Sci* 2007;45-52. <https://doi.org/10.7150/ijms.4.45>

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