Potential of Traditional Medicinal Plantsfor Treating Obesity: A Review

Mahnaz Kazemipoor¹, Che Wan Jasimah Wan Mohamed Radzi²⁺, Geoffrey A. Cordell³, Iman Yaze⁴

^{1,2}Department of Science & Technology Studies, Faculty of Science, University of Malaya, 50603 Kuala Lumpur, Malaysia

³ Natural Products Inc., Evanston, IL 60203, USA ⁴Faculty of medicine

Abstract. Obesity is a global health concern associated with high morbidity and mortality. Therapeutic strategies include synthetic drugs and surgery, which may entail high costs and serious complications. Plant-based medicinal agents offer an alternative approach. A review of the studies on accessible botanical sources for the treatment of obesity is provided, which attempts to explain how these medicinal plants act to cause weight loss, and which approach is safer and more efficient. Information was gathered forthe period of 1991 to 2012. Five basic mechanisms, including stimulating thermogenesis, lowering lipogenesis, enhancing lipolysis, suppressing appetite, and decreasing the absorption of lipids may be operating. Consumption of standardized medicinal plant extracts may be a safe treatment for obesity. However, some combinations of medicinal plants may result in either lower efficacy or cause unexpected side-effects.

Keywords: Medicinal plants, Adipose-tissue differentiation, Fat absorption, Slimming aids, Dietary supplements

1. Introduction

By 2005, obesity had affected 400 million adults [66], and since 1997, WHO has cited obesity as a global epidemic [3, 8]. More than half of the adult population in OECD countries is overweight (body mass index [BMI]>25 Kg/m²) [10]. According to WHO, obesity is related to cardiovascular diseases, hypertension, diabetes mellitus, gallbladder disease, cancer, endocrine and metabolic disturbances, osteoarthritis, gout, pulmonary diseases, as well as psychological issues, including social bias, prejudice, discrimination, and overeating [65]. Economically, obesity and its health consequences place enormous costsnow and for future health care [13, 49, 67]. Being overweight is a cosmetic issue, a major health risk factor [33], and maydecrease life expectancy [46]. A proliferation of high-cost, anti-obesity products in the market[25]. However, they exhibit side effects, such as gastrointestinal and kidney problems [25, 55], and only *Orlistat*and *Sibutramine* can be used long-term, in spite of issues regarding weight loss and tolerance[50, 51]. The use of natural remedies for weight loss has increased, based on reliability, safety, and cost compared with synthetic drugs [11] or surgical procedures[12], which may have limitations [40].

1.1. Natural Medications

Natural ingredients and medicinal plant preparationsmay enhance satiety, boost metabolism, and speed up weight loss [34, 43]. Including these foods in the diet may therefore assistslow, individual weight loss. However, doubts about human application remain[58]. Despite the global market for satiety, fat burning,

E-mail address: (jasimah@um.edu.my)

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⁺ Corresponding author. Tel.: + (0379675182); fax: + (0379674396).

dietary supplements and other weight management remedies, patient awareness of these products is insufficient [15]. Here, a brief review of natural medicinal agents and their anti-obesity potential is presented which could aidpatients in selecting botanical product to develop a healthy body.

2. Methods of Data Collection

Data were acquired from various databases, includingScience Direct, Pub-Med, Scopus, Web of Science, and from books and theses for the period from 1991 until January 2012. Key search words included: traditional medicine, medicinal herbs, plant extracts, anti-obesity, weight loss, overweight, botanical remedy, complementary therapy, natural, alternative, phytonutrients, phytochemicals, efficacy, safety, bioactive compounds, appetite, satiety, metabolism, thermogenesis, lipolysis, lipogenesis, adipocytes and anthropometric indices. Papers on human and animal studies, clinical trials, and related toplant-based obesity medication are discussed.

3. Results and Discussion

3.1. Mechanism of Action

Natural anti-obesity preparations can induce weight loss through several mechanisms. Their functions can be classified into five major categories, as shown in Table 1.

No.	Anti-obesity function	Anti-obesity preparations		
1	Inhibiting pancreatic lipase activity	chitosan [7, 27], levan [28], mate tea [41], oolong tea [23] jasmine tea [45], green tea [32]		
2	Enhancing thermogenesis	sea weed [36, 38, 39], bitter orange [14, 19, 52], soybean [24]		
3	Preventing adipocyte	turmeric [1], capsicum [22], palm oil [62], banana leaf [4, 31], brown algae [37],		
	differentiation	garlic [2], flaxseed [60], black soybean [29]		
4	Enhancing lipid metabolism	herb teas [45], cinnamon [57]		
5	Decreasing appetite	pine nut [48], pomegranate leaf [35], ginseng [30], Hoodia gordonii[63]		

Table 1: Different functions of anti-obesity medicinal plants in humans

Based onpancreatic lipase activity inhibition[5], some medicinal plants will prevent the intestinal lipid absorption, and non-absorbed fat will be excreted through oily faeces. Certain bioactive components can increase the metabolic rate[20] which enhances thermogenesis and helps burning calories and excess body fat. Prevention of adipocyte differentiation [62] throughmedicinal plant may inhibit adipogenesis and fat cellformation, and based on enhanced lipolysissome medicinal plantscan increase lipolysis through inducing β-oxidation or noradrenaline secretion in fat cells [45]. Other anti-obesity ingredients may suppress appetite and induce satiety [16], allowing for appetite control. These differing functions of antiobesity medicinal plants will result in a reduction of food and energy intake [19].

3.2. Approaches in Medicinal Plant Preparation with Maximum Efficacy and Safety

Medicinal plant samples may be the whole plant, or plant parts (stem, bark, leaf, flowers, and roots). These materials may be presented as powder or capsules, although the medicinal plants showing antiobesity properties were aqueous or alcoholic extracts. In this way, components inhibiting the anti-obesity compounds may be removed. Extraction, partial purification, and/or the isolation of active principle(s) could increase bioactive constituent bioavailability in the extracts, and enhance medicinal agent efficacy in weight loss[9, 56]. However (Table 2), different antiobesity preparations in combination with other medicinal plantsmay produce unexpected side-effects.

The application of single medicinal plants has not caused any adverse events. The undesiredhuman effects may be due to interactions between the different phytochemical constituents present in the different plants[21].

4. Summary and Conclusions

Based on a number of *in vivo* studies regarding the efficacy of anti-obesity medicinal plant preparations, they may act by stimulating thermogenesis, lowering lipogenesis, enhancing lipolysis, suppressing appetite, and decreasing lipid absorption. Single and mixed anti-obesity medicinal plant preparations may have

different effects. The botanical sources, route of administration, presence of various bioactive components and their respective functions, experimental methods used, treatment dosage, study design, treatment duration, andsafety and efficacy of the plant are also factors.

In conclusion, the dietary intake of the singlemedicinal plantsmayprovide a higher degree of safety and efficacy than mixed medicinal plant preparations. These findings support health organization recommendation regarding the regular consumption of vegetables and selected herbs, such as turmeric, capsaicin, ginger, and green tea. Improving knowledge on the use of anti-obesity medicinal preparations, and encouraging obese patients to consume them along with an enhanced exercise regimen and a healthy diet should be continued. Additional chemical, biological, and clinical studies are needed on the effectiveness of selected medicinal plants, particularly those used as spices and condiments, in ameliorating and treating obesity in humans. Such anti-obesity data would be useful for food and drugmanufacturers as new products are developed, and to governments in the regulation of food products as a way to promote and enhance public health.

Table 2: Comparison of safety and efficacy of single and mixed medicinal plant anti-obesity preparations

Medicinal Plant ingredient	*Result	Adverse effects	Combination formula	Result	Adverse effects
Rhubarb (rheum) [26]	+	Not reported	In combination with ginger, astragulus, red sage, turmeric, and gallic acid[18, 53]	greater weight gain in intervention group/ (-)	Musculoskeletal, gastrointestinal, oral, dermatologic, vaginal irritation, headache, etc.
Green tea (Camellia sinensis)[44]	+ (p<0.05)	Not reported	In combination with asparagus, black tea, guarana, kidney bean, <i>Garcinia cambogia</i> and chromium yeast [47]	no inter group difference in weight (-)	Gastrointestinal complaints
Bitter orange Citrus aurantium) [59]	+	Not reported	In combination with pantothenic acid, green tea leaf extract, guarana, white willow bark and ginger root [17]	greater weight gain in intervention group (p<0.04)	Hypertension, musculoskeletal, neurological, migraine, anxiety
Kidney bean (<i>Phaseolus</i> vulgaris) [61]	+ (p=0.07)	Not reported	In combination with green tea extract[6]	inter group diff. (+)	Flatulence, soft stool, constipation
Garcinia cambogia[42]	+ (p=0.03)	Not reported	in combination with natural caffeine [54]	No inter group diff. (p=0.3)	Not reported
Glucomannan fiber [64]	+ (p<0.005)	Not reported	in combination with chitosan, fenugreek, <i>Gymemna sylvestre</i> , vitamin C [68]	inter group differences (+) (p<0.01)	Constipation, headache, indigestion

^{*} Results indicate the efficacy and intergroup differences

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6. References

- [1] J. Ahn, H. Lee, S. Kim *et al.* Curcumin-induced suppression of adipogenic differentiation is accompanied by activation of Wnt/β-catenin signaling. *American Journal of Physiology-Cell Physiology*. 2010, 298 (6): C1510-C1516.
- [2] S. Ambati, J. Y. Yang, S. Rayalam *et al.* Ajoene exerts potent effects in 3T3 L1 adipocytes by inhibiting adipogenesis and inducing apoptosis. *Phytotherapy Research.* 2009, 23 (4): 513-518.

- [3] M. C. Auld, and L. M. Powell. Economics of food energy density and adolescent body weight. *Economica*. 2009, 76 (304): 719-740.
- [4] N. Bai, K. He, M. Roller *et al.* Active compounds from Lagerstroemia speciosa, insulin-like glucose uptake-stimulatory/inhibitory and adipocyte differentiation-inhibitory activities in 3T3-L1 cells. *Journal of agricultural and food chemistry.* 2008, 56 (24): 11668-11674.
- [5] R. B. Birari, and K. K. Bhutani. Pancreatic lipase inhibitors from natural sources: unexplored potential. *Drug discovery today.* 2007, 12 (19-20): 879-889.
- [6] G. S. Birketvedt, Composition for treating obesity comprising extract from white kidney beans, red kidney beans, and green tea leaves, USA, Google Patents, 2009, US 7,579,027 B2.
- [7] G. Bondiolotti, S. R. Bareggi, N. G. Frega *et al.* Activity of two different polyglucosamines, L1120 and FF450, on body weight in male rats. *European journal of pharmacology.* 2007, 567 (1-2): 155-158.
- [8] B. Caballero. The global epidemic of obesity: an overview. Epidemiologic reviews. 2007, 29 (1): 1-5.
- [9] J. Calixto. Efficacy, safety, quality control, marketing and regulatory guidelines for herbal medicines (phytotherapeutic agents). *Brazilian Journal of Medical and Biological Research*. 2000, 33 (2): 179-189.
- [10] M. Cecchini, F. Sassi, J. A. Lauer *et al.* Tackling of unhealthy diets, physical inactivity, and obesity: health effects and cost-effectiveness. *The Lancet.* 2010, 376 (9754): 1775-1784.
- [11] J. Chang. Medicinal herbs: drugs or dietary supplements? Biochem Pharmacol. 2000, 59 (3): 211-9.
- [12] A. Clegg, J. Colquitt, M. Sidhu *et al.* Clinical and cost effectiveness of surgery for morbid obesity: a systematic review and economic evaluation. *International journal of obesity*. 2003, 27 (10): 1167-1177.
- [13] G. A. Colditz. Economic costs of obesity and inactivity. Medicine & Science in Sports & Exercise. 1999, 31 (11): S663-S667.
- [14] C. Dallas, A. Gerbi, G. Tenca *et al.* Lipolytic effect of a polyphenolic citrus dry extract of red orange, grapefruit, orange (SINETROL) in human body fat adipocytes. Mechanism of action by inhibition of cAMP-phosphodiesterase (PDE). *Phytomedicine*. 2008, 15 (10): 783-792.
- [15] A. Esmaillzadeh, and L. Azadbakht. Major dietary patterns in relation to general obesity and central adiposity among Iranian women. *The Journal of nutrition*. 2008, 138 (2): 358-363.
- [16] P. Geoffroy, B. Ressault, E. Marchioni *et al.* Synthesis of Hoodigogenin A, aglycone of natural appetite suppressant glycosteroids extracted from Hoodia gordonii. *Steroids*. 2011, 76 (7): 702-708.
- [17] F. Greenway, L. de Jonge-Levitan, C. Martin *et al.* Dietary herbal supplements with phenylephrine for weight loss. *Journal of medicinal food.* 2006, 9 (4): 572-578.
- [18] F. Greenway, Z. Liu, C. Martin *et al.* Safety and efficacy of NT, an herbal supplement, in treating human obesity. *International journal of obesity.* 2006, 30 (12): 1737-1741.
- [19] S. Haaz, K. Fontaine, G. Cutter *et al.* Citrus aurantium and synephrine alkaloids in the treatment of overweight and obesity: an update. *Obesity reviews.* 2006, 7 (1): 79-88.
- [20] J. C. Hansen, A. P. Gilman, and J. Ø. Odland. Is thermogenesis a significant causal factor in preventing the "globesity" epidemic? *Medical hypotheses*. 2010, 75 (2): 250-256.
- [21] D. Heber. Herbal preparations for obesity: are they useful? *Primary care*. 2003, 30 (2): 441-463.
- [22] C. L. Hsu, and G. C. Yen. Effects of capsaicin on induction of apoptosis and inhibition of adipogenesis in 3T3-L1 cells. *Journal of agricultural and food chemistry*. 2007, 55 (5): 1730-1736.
- [23] T. Hsu, A. Kusumoto, K. Abe *et al.* Polyphenol-enriched oolong tea increases fecal lipid excretion. *European journal of clinical nutrition*. 2006, 60 (11): 1330-1336.
- [24] K. Ishihara, S. Oyaizu, Y. Fukuchi *et al.* A soybean peptide isolate diet promotes postprandial carbohydrate oxidation and energy expenditure in type II diabetic mice. *The Journal of nutrition*. 2003, 133 (3): 752-757.
- [25] B. Jacobs, and K. Gundling, ACP evidence-based guide to complementary and alternative medicine, p.^pp. 330-351: Amer College of Physicians, 2009.
- [26] H. Jin, and D. Jiao. Effect of jiang-zhi jian-fei yao on gastro-intestinal movement and adipose cell of abdominal wall. *Chinese journal of integrated traditional and Western medicine/Zhongguo Zhong xi yi jie he xue hui, Zhongguo Zhong yi yan jiu yuan zhu ban.* 1994, 14 (4): 230-231.
- [27] S. Jun, E. Jung, D. Kang *et al.* Vitamin C increases the fecal fat excretion by chitosan in guinea pigs, thereby reducing body weight gain. *Phytotherapy Research.* 2010, 24 (8): 1234-1241.
- [28] S. A. Kang, K. Hong, K. H. Jang *et al.* Altered mRNA expression of hepatic lipogenic enzyme and PPARα in rats fed dietary levan from *Zymomonas mobilis*. *The Journal of Nutritional Biochemistry*. 2006, 17 (6): 419-426.
- [29] H. J. Kim, I. Y. Bae, C. W. Ahn *et al.* Purification and identification of adipogenesis inhibitory peptide from black soybean protein hydrolysate. *Peptides.* 2007, 28 (11): 2098-2103.
- [30] J. H. Kim, D. H. Hahm, D. C. Yang *et al.* Effect of Crude Saponin of Korean Red Ginseng on High Fat Diet-Induced Obesity in the Rat. *Journal of pharmacological sciences*. 2005, 97 (1): 124-131.
- [31] G. Klein, J. Kim, K. Himmeldirk *et al.* Antidiabetes and anti-obesity activity of Lagerstroemia speciosa. *Evidence Based Complementary and Alternative Medicine*. 2007, 4 (4): 401-408.
- [32] S. I. Koo, and S. K. Noh. Green tea as inhibitor of the intestinal absorption of lipids: potential mechanism for its lipid-lowering effect. *The Journal of Nutritional Biochemistry*. 2007, 18 (3): 179-183.
- [33] P. G. Kopelman. Obesity as a medical problem. NATURE-LONDON-. 2000, 635-643.
- [34] N. I. Larson, M. T. Story, and M. C. Nelson. Neighborhood Environments:: Disparities in Access to Healthy Foods in the US. *American journal of preventive medicine*. 2009, 36 (1): 74-81. e10.

- [35] F. Lei, X. N. Zhang, W. Wang *et al.* Evidence of anti-obesity effects of the pomegranate leaf extract in high-fat diet induced obese mice. *Int J Obes (Lond).* 2007, 31 (6): 1023-9.
- [36] H. Maeda, M. Hosokawa, T. Sashima *et al.* Dietary combination of fucoxanthin and fish oil attenuates the weight gain of white adipose tissue and decreases blood glucose in obese/diabetic KK-Ay mice. *J. of Agricultural and Food Chemistry.* 2007, 55 (19): 7701-7706.
- [37] H. Maeda, M. Hosokawa, T. Sashima *et al.* Fucoxanthin and its metabolite, fucoxanthinol, suppress adipocyte differentiation in 3T3-L1 cells. *International journal of molecular medicine*. 2006, 18 (1): 147-152.
- [38] H. Maeda, M. Hosokawa, T. Sashima *et al.* Fucoxanthin from edible seaweed, Undaria pinnatifida, shows antiobesity effect through UCP1 expression in white adipose tissues. *Biochemical and biophysical research communications*. 2005, 332 (2): 392-397.
- [39] H. Maeda, T. Tsukui, T. Sashima *et al.* Seaweed carotenoid, fucoxanthin, as a multi-functional nutrient. *Asia Pac J Clin Nutr.* 2008, 17 (S1): 196-199.
- [40] L. K. Mahan, and S. Escott-Stump, Krause's food & nutrition therapy: Saunders/Elsevier, 2008.
- [41] F. Martins, T. M. Noso, V. B. Porto *et al.* Maté tea inhibits in vitro pancreatic lipase activity and has hypolipidemic effect on high-fat diet-induced obese mice. *Obesity.* 2009, 18 (1): 42-47.
- [42] R. D. Mattes, and L. Bormann. Effects of (-)-hydroxycitric acid on appetitive variables. *Physiology & behavior*. 2000, 71 (1): 87-94.
- [43] M. A. McCrory, B. R. Hamaker, J. C. Lovejoy *et al.* Pulse consumption, satiety, and weight management. *Advances in Nutrition: An International Review Journal.* 2010, 1 (1): 17-30.
- [44] T. Nagao, T. Hase, and I. Tokimitsu. A Green Tea Extract High in Catechins Reduces Body Fat and Cardiovascular Risks in Humans&ast. *Obesity*. 2007, 15 (6): 1473-1483.
- [45] H. Okuda, L. Han, Y. Kimura *et al.* Anti-Obesity Action of Herb Tea.(Part 1). Effects or Various Herb Teas on Noradrenaline-Induced Lipolysis in Rat Fat Cells and Pancreatic Lipase Activity. *Japanese Journal of Constitutional Medicine*. 2001, 63 (1/2): 60-65.
- [46] S. J. Olshansky, D. J. Passaro, R. C. Hershow *et al.* A potential decline in life expectancy in the United States in the 21st century. *New England Journal of Medicine*. 2005, 352 (11): 1138-1145.
- [47] T. Opala, P. Rzymski, I. Pischel *et al.* Efficacy of 12 weeks supplementation of a botanical extract-based weight loss formula on body weight, body composition and blood chemistry in healthy, overweight subjects-a randomised double-blind placebo-controlled clinical trial. *European journal of medical research.* 2006, 11 (8): 343-350.
- [48] W. J. Pasman, J. Heimerikx, C. M. Rubingh *et al.* The effect of Korean pine nut oil on in vitro CCK release, on appetite sensations and on gut hormones in post-menopausal overweight women. *Lipids in Health and Disease*. 2008, 7 (10): 7-10.
- [49] J. Picot, J. Jones, J. Colquitt *et al.*, The clinical effectiveness and cost-effectiveness of bariatric (weight loss) surgery for obesity: a systematic review and economic evaluation. *Health Technology Assessment*.2009, 13(41), 1-190, 215-357, iii-iv.
- [50] M. H. Pittler, and E. Ernst. Dietary supplements for body-weight reduction: a systematic review. The American journal of clinical nutrition. 2004, 79 (4): 529-536.
- [51] H. G. Preuss, D. DiFerdinando, M. Bagchi *et al.* Citrus aurantium as a thermogenic, weight-reduction replacement for ephedra: an overview. *Journal of medicine*. 2002, 33 (1-4): 247-264.
- [52] A. T. Roberts, C. K. Martin, Z. Liu *et al.* The safety and efficacy of a dietary herbal supplement and gallic acid for weight loss. *Journal of medicinal food.* 2007, 10 (1): 184-188.
- [53] D. Rothacker, and B. Waitman. Effectiveness of a Garcinia cambogia and natural caffeine combination in weight loss: a double-blind placebo-controlled pilot study. International Journal of Obesity. 1997, 21 (suppl 2): 53.
- [54] D. Rucker, R. Padwal, S. K. Li *et al.* Long term pharmacotherapy for obesity and overweight: updated meta-analysis. *Bmj.* 2007, 335 (7631): 1194-1199.
- [55] V. Schulz, R. Hänsel, and V. E. Tyler, *Rational phytotherapy: a physician's guide to herbal medicine*: Routledge, 2001.
- [56] X. Sheng, Y. Zhang, Z. Gong *et al.* Improved insulin resistance and lipid metabolism by cinnamon extract through activation of peroxisome proliferator-activated receptors. *PPAR Research.* 2008, 581348; 1-9.
- [57] S. Smyth, and A. Heron. Diabetes and obesity: the twin epidemics. *Nature Medicine*. 2006, 12 (1): 75-80.
- [58] S. J. Stohs, H. G. Preuss, and M. Shara. The Safety of Citrus aurantium (Bitter Orange) and its Primary Protoalkaloid p Synephrine. *Phytotherapy Research*. 2011, 25 (10): 1421-1428.
- [59] S. Tominaga, T. Sugahara, S. Nishimoto *et al.* The effect of secoisolariciresinol on 3T3-L1 adipocytes and the relationship between molecular structure and activity. *Bioscience, biotechnology, and biochemistry.* 2009, 73 (1): 35-39.
- [60] J. Udani, M. Hardy, and D. C. Madsen. Blocking carbohydrate absorption and weight loss: a clinical trial using Phase 2TM brand proprietary fractionated white bean extract. *Alternative medicine review*. 2004, 9 (1): 63-69.
- [61] H. Uto-Kondo, R. Ohmori, C. Kiyose *et al.* Tocotrienol suppresses adipocyte differentiation and Akt phosphorylation in 3T3-L1 preadipocytes. *The Journal of nutrition.* 2009, 139 (1): 51-57.
- [62] F. Van Heerden. Hoodia gordonii: A natural appetite suppressant. *Journal of Ethnopharmacology*. 2008, 119 (3): 434-437.
- [63] D. E. Walsh, V. Yaghoubian, and A. Behforooz. Effect of glucomannan on obese patients: a clinical study. *Int J Obes.* 1984, 8 (4): 289-93.

- [64] World Health Organization, Obesity: preventing and managing the global epidemic. Report of a WHO Consultation. WHO Technical Report Series, No. 894. WHO: Geneva, 2000.
- [65] World Health Organization. "Obesity and overweight," March 4, 2009; URL: http://www.who.int/mediacentre/factsheets/fs311/en/index.html
- [66] A. M. Wolf, and G. A. Colditz. Current estimates of the economic cost of obesity in the United States. *Obesity Research*. 1998, 6 (2): 97-106.
- [67] D. E. Woodgate, and J. A. Conquer. Effects of a stimulant-free dietary supplement on body weight and fat loss in obese adults: a six-week exploratory study. *Current Therapeutic Research*. 2003, 64 (4): 248-262.