ABSTRACT: Most diets in developed nations fall far short of recommended fruit and vegetable, and thus key nutrient, intake. Momordica cochinchinensis (gac), indigenous to Southeast Asia, contains high concentrations of lycopene and beta-carotene. In Vietnam, gac is prized by natives for promoting longevity and vitality. In a supplementation trial among Vietnamese children, gac increased serum vitamin A levels more than synthetic beta-carotene. Rosa roxburghii (cili), native to southwest China and traditionally used to combat stress and aging, is very rich in vitamin C and other phytonutrients. In humans, cili enhanced immunity and raised erythrocyte superoxide dismutase (SOD). Lycium chinense (wolfberry), also of Chinese origin, and traditionally used for longevity, wellness, and vision, is very rich in highly utilisable zeaxanthin. In animals, wolfberry showed immune modulating and antioxidant actions, and in humans, significantly increased blood SOD. Hippophae rhamnoides (sea buckthorn), native to Siberia and the Himalayas and traditionally used to expel phlegm and promote digestion, was shown in animals to have liver-protective and antioxidant activity. In humans, sea buckthorn greatly increased the dietary intake of flavonoids and vitamin C, and showed cardiovascular benefits. Thus, due to their nutritional value, intake of these fruits may greatly enhance the diet's healthfulness.

KEY WORDS: Antioxidants, Carotenoids, Hippophae rhamnoides, Lycium chinense, Momordica cochinchinensis, Rosa roxburghii

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INTRODUCTION

Over the past 25 years, a large number of human studies have led to the consensus that eating 5-9 servings of fruits and vegetables can help prevent many chronic diseases, and that most people do not attain this level of intake. Many health benefits can be attributed to the vitamins, minerals, antioxidants, carotenoids and other phytonutrients present in these foods. Thus, it would be desirable to add fruits with very high levels of these nutrients to the diet. There are four lesser-known fruits that have extraordinary nutritional value: Momordica cochinchinensis (gac), Rosa roxburghii (cili), Lycium Chinense (wolfberry), and Hippophae rhamnoides (sea buckthorn). These are evaluated in this review for antioxidant potency, nutritional content, traditional folklore usage, and scientific documentation of health benefits.

GAC (MOMORDICA COCHINCHINENSIS)

Traditional uses and characteristics

To native people, gac is valued for many important health benefits. Grown in Vietnam and other areas in Southern Asia, gac is esteemed as “the fruit from heaven” (Voung, 2001), and is prized for its ability to promote longevity, vitality, and health. A large, bright-red fruit (Kuhnlein, 2004), gac is known as “sweet gourd”. A perennial climber, the gac vine produces up to 60 fruits weighing 1-3 kg each in one season (Shadeque and Sharma Baruah, 1984). The seeds, known in traditional Chinese medicine (TCM) as “Mubiezi”, are thought to have resolvent and cooling properties, and are used for liver and spleen disorders, wounds, hemorrhoids, bruises, swelling, and pus (De Shan, et al., 2001). The Vietnamese often use gac in a dish called “xoi gac”, prepared by mixing gac seed and pulp with cooked rice to impart a red color and distinct flavor (Voung, 2001; Vo Van Chi, 1997; Vu Dinh Trac, 1986). Since xoi gac is served at festive occasions such as weddings, the lunar New Year, and other important celebrations (Do, 1999), it is essential to mask the white color of rice, since white is considered the color of death (Vuong, 2001). In addition to their use in xoi gac, the seed membranes are used to make a tonic for children and lactating or pregnant women, and to treat “dry eyes” (xerophthalmia) and night blindness (Guichard and Bui, 1941). Gac is not well known worldwide,
but to indigenous people, gac is familiar and easy to grow, yet is seasonal and only available for three months each year, where harvesting begins in September and lasts until December. In Vietnam, the gac vine is often seen growing on lattices going to the entrances of rural homes (Vuong et al., 2002). Gac may be underutilized in the diet worldwide because of its seasonality, regional lack of availability, and lack of awareness of its potential health value. (Vuong 2002; Vuong and King, 2003).

**Nutritional content**

Gac is becoming known as a premier source of carotenoids, especially beta-carotene and lycopene. Carotenoids were first identified in gac in 1941 (Guichard and Bui, 1941). However, it was only recently found that gac has beta-carotene and lycopene at very high levels, with those of lycopene being up to 308 µg/g in the seed membrane, about 10-fold higher than in other lycopene-rich fruits and vegetables (Vuong, 2001; Aoki et al., 2002; Vuong et al., 2003; Vuong et al., 2005). In gac aril (pulp) mean levels of lycopene registered even higher, at 2227 µg/g of fresh material. The aril also comprised of high levels of fatty acids, ranging from 17% to 22% by weight (Vuong and King, 2003; Ishida et al., 2004). Oil extracted from the gac fruit aril (gac oil) showed a total carotenoid concentration of 5700 µg/ml, with 2710 µg of that being beta-carotene. This oil also included high levels of vitamin E (Vuong and King, 2003; Kuhnlein, 2004). The fatty acids in the aril are important for the absorption of fat-soluble nutrients including carotenoids in a diet typically low in fat (Kuhnlein, 2004; Vuong, 2002). In addition, gac oil was readily accepted by women and children of Vietnam, and consumption of the oil reduced lard intake (Vuong et al., 2002; Vuong and King, 2003). Thus, gac provides an acceptable source of high levels of valuable antioxidants that have good bioavailability.

**The potential health benefits of lycopene**

Since the gac aril has over 70 times the lycopene/gram than tomatoes, this has interesting implications for prostate health. Epidemiological studies have shown that high intakes of tomatoes and tomato products, rich in lycopene, as well as high blood levels of lycopene, are significantly associated with decreased prostate cancer risk (Deming et al., 2002; Giles et al., 1997; Giovanucci et al., 1995; Giovanucci et al., 2002; Lu et al., 2001; Vogt et al., 2002). In a double-blind study of 26 men with newly diagnosed prostate cancer, lycopene was administered at 15 mg/day for three weeks before prostatectomy. A significantly increased number of subjects with tumors <4 mm in size, and decreased number with cancer of extra-prostatic tissue were observed (Kucuk et al., 2001). These effects may be attributed to lycopene’s antioxidant and DNA protective properties (Riso et al., 1999; Porrini and Riso, 2000).

**Clinical research**

Human research has now confirmed that the beta-carotene in gac is highly bioavailable. In a double-blind study with 185 Vietnamese preschool children, some were given Xoi Gac containing 3.5 mg/day beta-carotene, while others were given an identical-lookish dish containing 5 mg beta-carotene powder, for 30 days. At the end, the former group had significantly greater plasma levels of beta-carotene than the latter. Increases in plasma retinol, alpha-carotene, zeaxanthin, and lycopene levels were also significantly greater in children given gac (Vuong et al., 2002). This is of great importance, because worldwide, vitamin A deficiency continues to be a major health problem and the leading cause of blindness in children (Maurin and Renard, 1997; Underwood and Arthur, 1996; Pinnock, 1995).

It is likely that the fatty acids in gac are what make its beta-carotene more bioavailable than that of the synthetic form (Vuong et al., 2002). Conversely, consumption of certain betacarotene-rich foods has been shown to produce little increase in plasma beta-carotene or retinol concentrations (de Pee, et al., 1998; Vuong, et al., 2002).

**CILI (ROSA ROXBURGHII)**

**Traditional uses**

Cili, also known as sweet chestnut rose (van Rensburg, et al., 2005), is a wild plant of southwest China (the Guizhou province), and has traditionally been used for longevity, cancer, immunity, and atherosclerosis (Zhang, et al., 2001). Like gac, cili is of limited availability (van Rensburg, et al., 2005).

**Nutritional content**

Cili has many components with antioxidant activity, which may contribute to its health benefits. Protecatechusic acid, a biphenol found in cili, displayed antioxidant activity, (Psotova et al., 2003; Sroka and Cisowski, 2003; Yeh and Yen, 2003), and inhibited DNA scission by scavenging hydroxyl radicals (Ueda et al., 1991). Cili is also rich in vitamins C and E, biotin, polyphenols, polysaccharides, zinc, strontium, and SOD (Zhang et al., 2001). This antioxidant milieu may aid in the preservation of cili’s vitamin C (Yoshida et al., 1987), and contribute to cili’s cardiovascular benefits (Zhang et al., 2001; van Rensburg et al., 2005).

**Preclinical studies**

Animal research has indicated that cili may have potential cardiovascular, antioxidant, and life extending benefits. In cholesterol-fed rabbits and quails, cili juice significantly ameliorated atherosclerosis, reduced plasma lipids, and increased red blood cell SOD activity (Zheng et al., 2001; Hu, et al, 1994). In fruit flies given cili, the life span increased significantly, compared to a controls (Ma et al, 1997).

**Clinical research**

Two interesting double-blind, placebo-controlled studies with cili have shown antioxidant, cardiovascular, and mental benefits. In the first study, 10 ml of a cili extract was administered twice daily to 60 healthy men and women aged 50-75 years for two months, while a placebo was given to 61 others of the same age range. After cili supplementation,
natural killer (NK) cell activity, and erythrocyte catalase, SOD and glutathione (GSH) levels all increased, while serum lipid peroxides decreased. Various indices of cardiovascular function and microcirculation improved, and peripheral vascular resistance was reduced. Simple and selective light reaction time decreased, and memory quotient improved. All these changes were significant, with no such changes in the control group (Ma et al., 1997).

In another placebo-controlled, randomized, single-blind trial, 36 young, healthy, non-smoking subjects received either a placebo or encapsulated cili supplement for five weeks. The latter significantly increased reduced:oxidized GSH ratios in the blood, and plasma antioxidant capacity. This suggests that cili’s antioxidants protect reduced GSH, thus producing the observed shift in the glutathione redox state (van Rensburg et al., 2005).

WOLFBERRY (LYCIUM BARBARUM OR L. CHINENSE)

Traditional uses

Wolberry has a long history of traditional use, where in Northern and Central China, wolberry has been used in TCM for 2000 years, for longevity, vision, wellness, and headaches. Additionally, since the Tang Dynasty (1000-1400 AD), wolberry has been noted for its multiple traditional health benefits, such as nourishing the yin, strengthening the liver and kidney, and sustaining the blood. Anciently, people used wolberries to make soup, stew, tea, and wine, or chew them like raisins.

Preclinical studies

In animals and in vitro, wolberry has been shown to have interesting antioxidant, immune-enhancing, radioprotective, anti-aging, and other health benefits. In vitro, flavonoids from this fruit scavenged the potent superoxide and hydroxyl radicals in a concentration-dependent manner (Ren et al., 1995), and in rats, wolberry inhibited the lipid peroxidation of RBC membranes (Li et al., 2002). In S180 tumor-bearing mice, this fruit augmented the immune system (Gan et al., 2004), and a glycoconjugate from wolberry, named LbGp2, showed marked immunoenzymatic activity (Peng and Tian, 2001). When given at high doses i.p. for seven consecutive days before irradiation, both the roots and aerial parts of wolberry protected bone marrow from death in mice due to radiation (500 but not 250 mg/kg). This may have worked through enhancing the regeneration of hematopoietic stem cells due to enhanced post-irradiation repair or increased proliferation (Hsu et al., 1999). In male and female fruit flies (Drosophila melanogaster), wolberry increased the maximal lifespan, and in males, increased the average lifespan. In d-galactose-administered aged mice, wolberry at 20 mg/kg/d increased the serum and liver levels of SOD and GSH peroxidase to levels higher than that of aged mice not given d-galactose. Wolberry reduced lipofuscin levels below that found in control aged mice, and reduced MDA content dose-dependently, where the high dose reduced it to levels found in young mice (Wang et al., 2002). Containing naturally occurring free-radical scavenging flavonoids, wolberry has also shown potential hypotensive, hepatoprotective, anti-asthmatic (Huang et al., 1998), hypoglycemic, and hypolipidemic effects (Luo et al., 2004).

Human studies

With aging, oxidative stress is known to increase, and wolberry has shown anti-aging effects in humans. In 25 Chinese subjects aged 64-80 years, who could care for themselves, ingestion of 50 g/d dry wolberry fruit for 10 days significantly increased blood SOD and hemoglobin by 48 and 12%, respectively, and decreased blood lipid peroxides by 65% (Li et al., 1991). Wolberry also improved some conditions associated with age. In another study, thirty-six of 42 healthy subjects (35 males and 7 females), aged an average of 68 years, were given 50 mg wolberry extract twice daily for two months, and the other six served as controls. The wolberry group showed dramatically decreased dizziness, fatigue, chest distress, sleep problems, and anorexia, while the control group showed much less pronounced changes (significance not assessed) (Li 1989).

Wolberry is very rich in zeaxanthin, which showed good bioavailability. In a randomized, single-blind cross-over study with 12 volunteers, a balanced breakfast was co-administered with 5 mg 3R-3’R-zeaxanthin, either in non-esterified (synthetic), or in esterified (dipalmitate) form found in wolberry, suspended in 150g yogurt. This was followed by a 3-week zeaxanthin-depletion period, after which treatments were reversed. Plasma zeaxanthin 24h area-under-the-curve, was significantly (~2-fold) greater with the esterified form (Breithaupt, et al., 2004). Wolberry has significant implications for ocular health. A number of studies support the protective role for zeaxanthin in the prevention of age-related eye diseases. For example, reduced risks of cataracts (Brown et al., 1999; Chasan-Taber et al., 1999) and ARMD (Seddon et al., 1994) have been associated with high intakes of leafy green vegetables, which are rich sources of lutein and zeaxanthin. In addition, lower levels of serum (Bernstein et al., 1999; Chasan-Taber et al., 1999) and ARMD (Bone et al., 2002; Seddon et al., 1994) zeaxanthin were found in ARMD patients compared to healthy controls. Many other studies have also indicated that zeaxanthin is important for ocular health (Breithaupt et al., 2004; Bone et al., 1985; Bone et al., 1992; Bone et al., 1993; Bone et al., 2003; Mozafarzadeh et al., 2003; Gale et al., 2003; Johnson et al., 2002).

SEA BUCKTHORN (SB, HIPPOPHAE RHAMNOIDES)

Traditional uses and history

In Russia and China, SB has a rich history of many uses. In the Chinese Pharmacopoeia, SB is recognized for the following indications: expelling phlegm, arresting coughs, promoting digestion, removing food stagnancy, and promoting blood flow to remove blood stasis (Thomas et al., 2004). The seed and pulp oils of SB have been traditionally used in Russia and China to treat skin disease (Yang et al., 2000). In Russia, active
Compounds have been investigated in the plant’s fruits, leaves, and bark since the 1940s. SB reportedly had use as a fruit in the diet of Russian cosmonauts and the oil in a cream to protect them from solar radiation (Thomas et al., 2004).

Preclinical studies

SB has been shown to have a wide variety of antioxidant and cell-protective benefits. In lymphocytes, SB inhibited chromium-induced free radical production, apoptosis, and DNA fragmentation, and restored antioxidant status to that of control cells. In addition, SB arrested the chromium-induced inhibition of lymphocyte proliferation. These observations suggest that SB has marked cytoprotective properties, which can be attributed to its antioxidant activity (Geetha et al., 2002; Geetha et al., 2003). SB also protected the functional integrity of mitochondria from radiation-induced oxidative stress (Goel et al., 2005).

In albino rats, the leaf extract of SB exerted significant protection against chromium-induced oxidative injury (Geetha et al., 2003). In rats given i.p. nicotine for three weeks, SB and vitamin E prevented nicotine-induced increases in MDA. SB but not vitamin E also prevented nicotine-induced reductions in erythrocyte SOD. In addition, compared with rats given nicotine only or nicotine plus vitamin E, those given SB had increased erythrocyte GSH-Px activity. Plasma vitamin A levels were higher in both the vitamin E and SB groups, compared with the nicotine and control groups (Suleyman et al., 2002). A study with mice demonstrated that SB may have liver-protective ability. In animals treated with carbon tetrachloride, acetalaminophen, and ethyl alcohol, SB significantly reduced the increases in SGOT, SGPT, and MDA induced by these agents, and the decrease in reduced glutathione induced by acetalaminophen. This indicates reduced injury to the liver from lipid peroxidation (Lipkan and Oliinyk, 2000; Cheng 1990; Cheng et al., 1992; Cheng et al., 1994).

SB’s free-radical scavenging activity is due in large part to its phenolic fractions (Gao et al., 2000), with the predominant polyphenols being flavonoids, and catechins and phenolic acids representing minor portions. Vitamin C is also a major antioxidant found in SB (Rosch et al., 2003).

Clinical research

In humans, SB or its components have been shown to have cardiovascular benefits, and to increase the intake of important dietary nutrients, especially flavonoids and vitamin C. The total flavones of SB, when given for 8 weeks to 35 hypertensive patients, prevented supine isometric exercise-induced increases in heart rate, blood pressure, and plasma catecholamine concentrations (Zhang et al., 2001). SB pulp oil at 5 grams/day induced a significant increase in serum HDL-C in patients with atopic dermatitis (Yang et al., 1999). In 20 healthy male volunteers given either SB juice or a placebo for 8 weeks, SB increased daily intakes of beta-carotene, alphatocopherol, flavonoids, and vitamin C by 1.0, 3.2, 355, and 462 mg/d, respectively (Eccleston et al., 2002).

CONCLUSIONS

Gac, cili, wolfberry, and sea buckthorn are all very highly nutritious fruits that will, individually or in combination, greatly enhance the healthfulness of a diet. Clinical research has shown that one or more of these fruits supported normal vitamin A levels and enhanced bodily antioxidant levels, cardiovascular and mental health, and immunity. Gac and wolfberry, with their high level of bioavailable carotenoids, may also promote prostate health and protect the eyes from age-related macular degeneration and cataracts. Cili, with its powerful antioxidant milieu, may help fight some of the signs of aging, as supported by its extension of life in fruit flies, and, in older adults, its enhancement of some aspects of cardiovascular and mental function, and reduction of oxidative stress. Extensive preliminary evidence suggests that SB has powerful antioxidant effects and thus the ability to reduce oxidative stress in many situations, such as exhaustive exercise, driving in heavy traffic, smoking, weight loss, etc. SB has also been shown to protect organs such as the liver and circulatory system. It appears that consumption of these fruits can provide a wide range of potential health benefits, and thus, the increase of public awareness and intake of these fruits may be of significant value to society.

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