



Review Article

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A review of chemical constituents and pharmacological properties of *Hibiscus sabdariffa* L.

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ABSTRACT

Hibiscus sabdariffa is a medicinal plant that is consumed for its health benefits, juice/concoction prepared from the plant is taken as a preventive/curative measures against diabetes and hypertension. The antihypertensive and other pharmacological properties such as antibacterial, anti-oxidant, nephro- and hepato-protective, renal/diuretic effect, anti-cholesterol, and anti-diabetic effects of *Hibiscus sabdariffa* have been demonstrated in several studies. Constituents of different plant parts of *Hibiscus sabdariffa* include phenolic acids, organic acid, flavonoids and anthocyanins which may contribute to the pharmacological effects of the plant. *Hibiscus sabdariffa* is relatively safe as LD₅₀ of its extract in rats was found to be above 5000 mg/kg. Therefore, *Hibiscus sabdariffa* because of its pharmacological and nutritional benefits could be exploited in the management of various pathological conditions such as cardiovascular disease, cancer, neurological disorders and diabetes.

Introduction

There is growing market for nutraceutical and functional foods, while study on natural sources of antioxidants and their potential as nutraceutical and functional foods is on the increase (Cevalls - Casals and Cisneros-Zealous, 2003). One plant that have attracted much attention over the years for its health benefits is roselle (*Hibiscus sabdariffa*), many studies on the plant, its numerous preparation and constituents focused on its antioxidant properties. *Hibiscus sabdariffa* L. (roselle) belongs to the family Malvaceae. It exists as herbs or shrubs, often with fibrous stems, (Eno, 2000). The leaves are deeply three- to five-lobed, 8–15 cm long, arranged alternately on the stems.

Vernacular names, in addition to roselle, in English-speaking regions are rozelle, sorrel, red sorrel, and Florida cranberry. In North Africa and the Near East *Hibiscus sabdariffa* is called karkadé or carcadé (Morton, 1987). *Hibiscus sabdariffa* is believed to have originated from India and Malaysia, where it is commonly cultivated, and must have been carried at an early date to Africa (Morton, 1987). Two main types of *Hibiscus sabdariffa* L. exist. The more important economically is *Hibiscus sabdariffa* variety *altissima* Wester, an erect, sparsely branched annual plant which is cultivated for its jute-like fiber in India, the East Indies, Nigeria and to some extent in tropical America. The other distinct type of roselle, *Hibiscus sabdariffa* variety *sabdariffa*,

embraces shorter, bushy forms which have been described as races: *bhagalpuriensi*, *intermedius*, *albus*, and *ruber*, all breeding true from seed (Morton, 1987).



Fig. 1: *Hibiscus sabdariffa* L.

In India, Africa and Mexico, all above-ground parts of the *Hibiscus sabdariffa* plant are valued in native medicine. Infusions of the leaves or calyxes are regarded as diuretic, cholerectic, febrifugal and hypotensive, decreasing the viscosity of the blood and stimulating intestinal peristalsis. The fresh calyx of *Hibiscus sabdariffa* is eaten raw in salads, is cooked and used as a flavouring in cakes, presently, it is consumed worldwide as a cold beverage and as a hot drink (sour tea) (Facciola, 1990; Wang et al., 2000; Ross, 2003). The red anthocyanin pigments in the calyxes are used as food colouring agents (Esselen and Sammy, 1975). Seeds of *Hibiscus sabdariffa* are used in oily soups, sauces and coffee substitute (Duke, 1983; Kunkel, 1984; Facciola, 1990). Root of *Hibiscus sabdariffa* is edible but very fibrous, mucilaginous, without very much flavour (Cribb, 1976).

Constituents of *Hibiscus sabdariffa*

There are many published reports on the constituents of different plant parts of *Hibiscus sabdariffa*.

Organic acids

Citric and malic acids are the major organic acids in aqueous extracts of the flowers of *Hibiscus sabdariffa* (Buogo and Picchinenna, 1937) this finding was collaborated by earlier works

(Indovina and Capotummino, 1938; Reaubourg and Monceaux, 1940). Lin, (1975) detected tartaric acid along with citric and oxalic acids, by paper chromatography in flower extracts of *Hibiscus sabdariffa*. High concentrations of oxalic, malic, tartaric and succinic acids were also reported to be present in the calyx of *Hibiscus sabdariffa* with the latter predominating (Wong et al., 2002). Khafaga and Koch (1980) detected citric, hibiscus, malic and tartaric acids in the calyxes of five strains of *Hibiscus sabdariffa* var. *sabdariffa*. Ascorbic acid was also reported to be present in aqueous extracts of *Hibiscus sabdariffa* (Buogo and Picchinenna, 1937; Reaubourg and Monceaux, 1940; Wong et al., 2002).

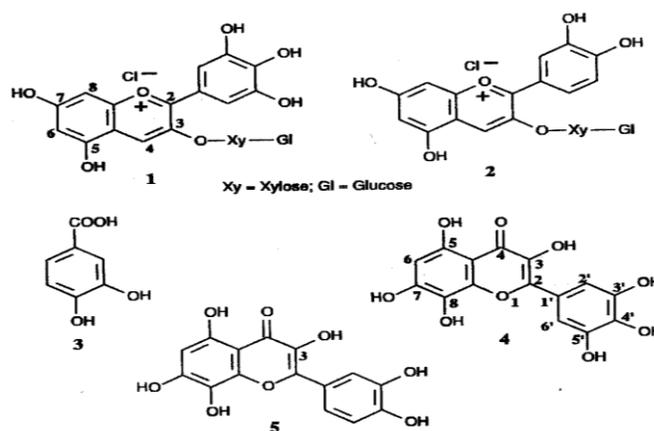


Fig. 2: Some constituents of *Hibiscus sabdariffa*: 1. Delphinidin-3-sambubioside, 2. Cyanidin-3-sambubioside, 3. Hibiscus protocatechuic acid, 4. Hibiscetin, 5. Gossypetin. Modified from Ali et al. (2005).

Anthocyanins

Most of the chemical investigations of the flower constituents have been directed towards characterization of their pigments. Yamamoto and Oshima (1932) isolated an anthocyanin, to which they assigned the structure, cyanidin-3-glucoside this was later changed to delphinidin-pentoside-glucoside (Yamamoto and Oshima, 1936). Delphinidin and cyanidin were reported as major constituents of plants grown in Trinidad (Forsyth and Simmonds, 1954). These pigments were further examined by Du and Francis (1973), who also isolated delphinidin-3-sambubioside (major component), delphinidin-3-monoglucoside and cyanidin-3-monoglucoside, but, in addition, characterized cyanidin-3-sambubioside as the

second most abundant anthocyanin in the extract. Shibata and Furukawa (1969) had earlier studied the pigments of Taiwanese roselle and reported the presence of delphinidin-3-sambubioside, along with small amounts of delphinidin-3-monoglucoside, cyanidin-3-monoglucoside and delphinidin. More recently, anthocyanins in *Hibiscus sabdariffa* had been quantified with HPLC and their relative percentage determined: delphinidin-3-sambubioside (56%); delphinidin-3-glucoside (4%); cyanidin-3-sambubioside (33%) and cyanidin-3-glucoside (3%) (Hong and Wrostad, 1990; Sukwattanasinit et al., 2007; Owoade et al., 2015)

Carbohydrate content

The petals of *Hibiscus sabdariffa* was reported to yield 65% (dry weight) of mucilage, and this yielded galactose, galacturonic acid and rhamnose on hydrolysis (El-Hamidi et al., 1966). Three water-soluble polysaccharides have been extracted from the flower buds of *Hibiscus sabdariffa*. The neutral compounds are composed of arabinans and arabinogalactans of low relative molecular mass. The major fraction was shown to be a pectin-like molecule ($M_r = 10^5$ Da). The main chain is composed of α -1, 4-linked galacturonic acid (24% methyl esterified) and α -1, 2-linked rhamnose. Side chains are built of galactose and arabinose and are connected to the main chain via C-4 of every third rhamnose (Muller and Franz, 1992).

Table 1. The General Composition of fresh leaf of *Hibiscus sabdariffa*. Modified from Duke (1983).

Component	Amount (% fresh leaf wt.)
Water	85.6
Protein	3.3
Fat	0.3
Total Carbohydrate	9.2
Fiber	1.6
Ash	1.6
Calcium	0.213
Phosphorus	0.093
Iron	0.0048
β -Carotene Equivalent	0.0041
Ascorbic Acid	0.054
Thiamine	0.00017
Riboflavin	0.00045
Niacin	0.0012

Table 2. The General Composition of fresh fruit of *Hibiscus sabdariffa*. Modified from Duke (1983).

Component	Amount (% fresh fruit wt.)
Water	84.5
Protein	1.9
Fat	0.1
Total Carbohydrate	12.3
Fiber	2.3
Ash	1.2
Calcium	0.0017
Phosphorus	0.057
Iron	0.0029
β -Carotene Equivalent	0.0003
Ascorbic Acid	0.014

Lipid content

The sterols of the seed oil of *Hibiscus sabdariffa* were studied by Salama and Ibrahim (1979), who reported the presence of cholesterol, campesterol, stigmasterol, β -sitosterol, α -spinasterol and ergosterol. The seed oil has also been found to be good source of lipid-soluble antioxidant, α -tocopherol 25% β -tocopherol 74.5% and γ -tocopherol 0.5% (Mohamed et al., 2007) while the component acids of the seed lipids were 2.1% myristic, 35.2% palmitic, 2.0% palmitoleic, 3.4% stearic, 34.0% oleic, 14.4% linoleic, and 3 unusual HBr-reacting fatty acids (cis-12, 13-epoxy-cis-9-octadecenoic (12,13-epoxoleic) 4.5%; sterculic, 2.9%; and malvalic, 1.3%) (Ahmad et al., 1979)

Polyphenols: Flavonoids and phenolic acids

In the last few decades there has been great interest in plant polyphenolic flavonoids and phenolic acids due to their antioxidant activity and protective effect against the development of cardiovascular disease and cancer (Schroeter, 2002; Fraga, 2007). Hibiscitrin, gossypitrin and sabdaritrin have been isolated from the flower petals of *Hibiscus sabdariffa*. Further studies on these compounds proved Hibiscitrin to be the 3-monoglucoside of hibiscetin, and gossypitrin to be the 7-glucoside of gossypetin while sabdaritrin on acid hydrolysis, yielded hydroxyflavonesabdaretin (Rao and Seshadri, 1942; Rao and Seshadri, 1948). Owoade et al., (2016), using TLC, HPLC and LCMS analysis showed the presence of ferulic

acid, chlorogenic acid, naringenin, rutin and quercetin in *Hibiscus sabdariffa* extracts. Also, protocatechuic acid, catechin, epigallocatechin, epigallocatechin gallate and caffeic acid have been identified with HPLC in an extract of *Hibiscus sabdariffa* (Lin, 2005; Owoade et al., 2016). Earlier workers have also isolated protocatechuic acid (Tseng et al., 1996), eugenol (Chen et al., 1998) and quercetin (Takeda and Yasui, 1985) in *Hibiscus sabdariffa*.

Pharmacological properties

Effect on blood pressure

Intravenous injection of aqueous extracts of *Hibiscus sabdariffa* calyx to anaesthetized cats (Ali et al., 1991) and anaesthetized rats (Adegunloye et al., 1996) lowered blood pressure in a dose-dependent manner. More recently, the antihypertensive action of *Hibiscus sabdariffa* has been confirmed in rats with experimental hypertension (Odigie et al., 2003; Mojiminiyi et al., 2007) and in spontaneously hypertensive rats (Onyenekwe et al., 1999) given the aqueous extracts at doses of 250–1000 mg/kg for up to 14 weeks. Dietary supplementation with *Hibiscus sabdariffa* has been shown to have blood pressure reducing effects in patients with moderate essential hypertension, (Haji-Faraji and Haji-Tarkhani, 1999; Herrera-Arellano et al., 2004; Herrera-Arellano et al., 2007). This hypotensive action of *Hibiscus sabdariffa* extracts was due to inhibition of angiotensin-converting enzyme (Jonadet et al., 1990; Herrera-Arellano et al., 2007). The inhibition of angiotensin-converting enzyme has also been demonstrated *in vitro* with a crude hydroethanol extract of *Hibiscus sabdariffa* calyces, and was ascribed to flavones present in the extract. In addition, a beneficial cardioprotective effect of this extract was shown *in vivo*, and was attributed to flavonoids and anthocyanins (Jonadet et al., 1990).

Lipid-lowering effects

Blood lipids and lipoproteins circulating in the blood in the form of LDL are decreased in response to treatment with *Hibiscus sabdariffa*. Ethanol extract of *Hibiscus sabdariffa* has been shown to reduce cholesterol, VLDL-cholesterol and LDL-cholesterol in alloxan – diabetic rats (Farombi and

Ige, 2007). Dietary supplementation with *Hibiscus sabdariffa* was effective in lowering serum concentrations of triglycerides, total cholesterol and LDL-cholesterol in hypercholesterolemic rabbits (Chen et al., 2003), and hypercholesterolemic rats (Hirunpanich et al., 2006). In addition, thiobarbituric acid reactive substances (TBARS) and conjugated dienes formed during oxidation of LDL by CuSO₄, CCl₄ were reduced (Hirunpanich et al., 2006; Owoade and Adetutu, 2015). Similar study using *Hibiscus* anthocyanins (HAs) extracts shown the extracts decrease the relative electrophoretic mobility of oxLDL, inhibit fragmentation of Apo B, reduced TBARS formation in the Cu²⁺-mediated oxidize LDL and scavenge over 95% of free DPPH radicals (Chang et al., 2006). Lipid fractions in plasma, heart, brain, kidney and liver were lowered in hypercholesterolaemia rats fed with *Hibiscus sabdariffa* calyx (5% or 10%) for 9 weeks (El-Saadany et al., 1991).

Anticancer effect

In vitro studies have shown that *Hibiscus sabdariffa* extracts can induce apoptosis in cancer cells. *Hibiscus* polyphenol-rich extracts (HPE) induce cell death in human gastric carcinoma (AGS) in a concentration-dependent manner (Lin et al., 2005; Lin et al., 2007), this effect of HPE on AGS cells was mediated via p53 signalling and p38 MAPK/FasL cascade pathway (Lin et al., 2005). Also, *Hibiscus* anthocyanins extract (a group of natural pigments existing in the dried calyx of *Hibiscus sabdariffa* L.) caused cancer cell apoptosis, in HL-60 cells (Chang et al., 2005; Sowemimo et al., 2007), similarly Delphinidin 3-sambubioside (Dp3-Sam), isolated from the dried calices of *Hibiscus sabdariffa* L. induce apoptosis in human leukemia cells (HL-60) (Hou et al., 2005). Anticlastogenic effects of *Hibiscus sabdariffa* extract has been demonstrated against sodium arsenite-induced micronuclei formation in erythrocytes in mouse bone marrow (Adetutu et al., 2004). Various studies on *Hibiscus* protocatechuic acid has demonstrated its ability to inhibit the carcinogenic action of various chemicals in different tissues of the rat, including diethyl nitrosamine in the liver (Tanaka et al., 1993), 4-nitroquinoline-1-oxide in the oral cavity (Tanaka et al., 1994), azoxymethane in the colon (Kawamori et al., 1994), N-methyl-N-nitrosourea in glandular

stomach tissue (Tanaka et al., 1995) and Nbutyl-N-(4-hydroxybutyl)nitrosamine in the bladder (Hirose et al., 1995). Tseng et al. (2000) also demonstrated that Hibiscus protocatechuic acid inhibits the survival of human promyelocytic HL-60 cells in a concentration- and time-dependent manner. The data presented by Tseng et al. (2000) suggest that the compound is an apoptosis inducer in human leukaemia cells and that RB phosphorylation and Bcl-2 protein may play a crucial role in the early stage.

Renal effects

Oral administration of *Hibiscus sabdariffa* extracts significantly normalizes the level of ammonia, urea, uric acid, creatinine and non-protein nitrogen in the blood of ammonium chloride-induced hyperammonemic rats (Essa and Subramanian, 2007). Consumption of *Hibiscus sabdariffa* extract in normal human subject significantly decreased the urinary concentrations of creatinine, uric acid, citrate, tartrate, calcium, sodium, potassium and phosphate, but not oxalate (Kirdpon et al., 1994). Also, low dose of *Hibiscus sabdariffa* (16 g/day) caused a more significant decrease in salt output in the urine than a high dose (24 g/day) (Kirdpon et al., 1994). Dietary supplementation with dried calyx of *Hibiscus sabdariffa* in rats resulted in a significant uricosuric action (Caceres et al., 1987; Mojiminiyi et al., 2000).

Scavenging of ROS

Hibiscus sabdariffa extracts and its constituents, Protocatechuic acid, anthocyanins demonstrated the ability to scavenge the 1, 1-diphenyl-2-picrylhydrazyl (DPPH) and 2,2-azino-bis(3-ethylbenzthiazoline-6-sulfonic acid) (ABTS) free radicals using a cell free system (Chang et al., 2006; Owoade et al., 2015; 2016). *Hibiscus sabdariffa* extracts and its constituents have also been observed to scavenge the t-butyl hydroperoxide radical and hence prevent oxidative damage in rat primary hepatocytes (Tseng et al., 1996; Tseng et al., 1997; Wang et al., 2000; Liu et al., 2002). The extracts have been shown to scavenge hydroxyl radical (OH[•]) and Hydrogen peroxide (H₂O₂) (Farombi and Fakoya, 2005). The extracts also showed strong inhibitory effect on xanthine oxidase activity and superoxide (•O₂⁻) radical

(Tseng et al., 1997, Owoade, et al., 2016). Hibiscus protocatechuic acid isolated from *Hibiscus sabdariffa* inhibits lipopolysaccharide-induced rat hepatic damage (Lin et al., 2003) and inhibits oxidation of low-density lipoprotein induced by either copper or a nitric acid donor (Lee et al., 2002). *Hibiscus sabdariffa* anthocyanins were effective in significantly mitigating the pathotoxicity induced by paracetamol in mice (Ali et al., 2003), it also protects against DNA damage induced by tert-butylhydroperoxide in rat smooth muscle and hepatoma cells (Lazze et al., 2003). In view of the established strong antioxidant and anti-lipidperoxidation actions of *Hibiscus sabdariffa* extracts and the compounds they contain (Tseng et al., 1997; Wang et al., 2000; Suboh et al., 2004), anthocyanins and Hibiscus protocatechuic acid may potentially be useful in ameliorating or preventing these diseases and conditions.

Effects on endogenous antioxidant defences

Dietary supplementations with *Hibiscus sabdariffa* extracts has been shown to significantly reduce carbon tetrachloride (CCl₄) induced liver damage in rats (Liu et al., 2006; Owoade et al., 2015; 2016), acetaminophen and Fe²⁺ induced liver damage in mice and rats (Olaleye and Rocha, 2007 & b) and cadmium induce liver toxicity (Asagba et al., 2007). Also, aqueous extracts of *Hibiscus sabdariffa* demonstrated protective effect against azathioprine-induced hepatotoxicity. Animals pre-treated with the extracts not only failed to show necrosis of the liver after azathioprine administration, but also retained livers that, for the most part, were histologically normal (Amin and Hamza, 2005). In all studies *Hibiscus sabdariffa* extracts significantly decreased the elevation of aspartate aminotransferase (AST) and alanine aminotransferase (ALT) in plasma (Amin and Hamza, 2005; Liu et al., 2006; Asagba et al., 2007; Olaleye and Rocha, 2007, Owoade et al., 2016). In a similar study oral administration of the ethanol extracts of *Hibiscus sabdariffa* significantly decreased sodium arsenite - induced malondialdehyde (MDA) formation in rat's liver, the extract also attenuated sodium arsenite induced reduction in the serum level of vitamin C (Usuh et al., 2005). In all these studies it was demonstrated that pre-treatment of animal with *Hibiscus sabdariffa* extracts prevented GSH

depletion, while other endogenous antioxidant enzymes (SOD, catalase and glutathione peroxidase) activity were increased couple with decrease in lipid peroxidation (Amin and Hamza, 2005; Usoh et al., 2005; Liu et al., 2006; Essa and Subramanian, 2007).

Effect on smooth muscle

Hibiscus sabdariffa have been shown to have relaxation effect on the smooth muscles, and this has been proposed to be partially responsible for its hypotensive action (Ajay et al., 2007). The extracts of *Hibiscus sabdariffa* calyces inhibited the tone of various isolated muscle preparations that included rabbit and rat aortic strip (Obiefuna et al., 1994; Ajay et al., 2007) and rat ileal strip (Salah et al., 2002). The extract also rhythmically contracted rat uterus, guinea-pig tracheal chain and rat diaphragm. The same extract stimulated quiescent rat uterus and frog rectus abdominus muscle (Ali et al., 1991; Fouda et al., 2007). The tonic effects on rat uterus were partially reduced by hydrocortisone and indomethacin. The overall effect is a direct relaxation of the smooth muscles. The relaxant response was related to endothelium-dependent and endothelium-independent mechanisms (Obiefuna et al., 1994), or mediated through calcium channels, possibly generated by constituents such as quercetin and eugenol (Salah et al., 2002; Ajay et al., 2007). However, the presence of stimulatory substance(s) in the extract has also been demonstrated using the frog rectus abdominus preparation (Ali et al., 1991).

Toxicological effect

The extract of *Hibiscus sabdariffa* was found to be relatively and virtually non-toxic with LD50 in rats to be above 5000 mg/kg (Onyenekwe et al., 1999).

Conclusion

The information from in vitro and in-vivo studies shows a wide range of potentially new health applications and therapeutic targets for *Hibiscus sabdariffa*. *Hibiscus sabdariffa* is relatively safe and virtually non-toxic. Many pharmacological properties of *H. sabdariffa* may be attributed to the presence of a plethora of phytochemicals in the plant. The potent antioxidant activity of *Hibiscus sabdariffa* may be linked to the presence different

antioxidants compounds with differing sites and mechanisms of action which may act alone or in concert with one another. Therefore, dietary supplementation of *Hibiscus sabdariffa* plant extract may be beneficial in reducing the risk of developing various pathological conditions such as cardiovascular disease, cancer, neurological disorders and diabetes.

Conflict of interest statement

Authors declare that they have no conflict of interest.

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