

International Journal of Phytomedicine 2018;10(3):148-152



A phytopharmacological review on an Arabian medicinal plant: *Caralluma flava* N.E.Br

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Abstract

Caralluma flava (N.E.Br.) is a medicinal plant which is gaining renowned attention in scientific community due to its potential curative powers. Traditionally, it is used in Arab regions as an anti-diabetic and anti-ulcer. It is also used as an anti-inflammatory, anti-parasitic and anti-pyretic. Due to its pharmacological values, initial biological activities of *C. flava* have been carried out recently and some pregnane glycoside molecules have been isolated and identified from its sap. The plant lacks the reports of phytochemical studies and pharmacological activities which support its further investigation and validation of the traditional therapeutic claims. The compiled data in this review is intended to serve as a reference to many researchers and may help them to focus on the priority areas of research yet to be discovered.

Keywords: Caralluma flava; pregnane glycoside; anticancer activity; Desmidorchis flava

Introduction

Caralluma flava (N.E.Br.) Meve & Liede is a succulent herb which grows wildly in many regions of Sultanate of Oman, Yemen and United Arab Emirates [1-5]. It burgeons after rain and mainly found on stony sides of the wadies, dry riverbeds, and hillsides. The fresh juicy stems of this cactus-like plant are collected by the natives and consumed as food. Due to its extremely bitter taste, it is eaten with an addition of spices and lemon. It has bright yellow flowers which are quite unmistakable to identify and are also edible. These flowers have an unpleasant odor and a distinctive flavor with a sweet aftertaste [6-8]. The plant is available in the vegetable and local markets of Arab regions in fresh and dried powdered forms. It is very well known for its medicinal properties [9]. Taxonomical classification of *C. flava* is given in Table-1.

Kingdom	: Plantae
Phylum	: Magnoliophyta
Class	: Magnoliopsida
Order	: Gentianales
Family	: Apocynaceae
genus	: Caralluma
specie	: flava
Plant's Name	: Caralluma flava (N.E.Br.) Meve & Liede
Synonyms	: Desmidorchis flava (N.E.Br.) Meve & Liede
	: Crenulluma flava (N.E.Br.) Plowes
	: Desmidorchis flavus
Vernacular	: <i>Zoja</i> (Jibbali Arabic)
names	
	: <i>Duru-el-kebeh</i> (Dhofari Arabic)

Botanical description

C. flava is a succulent stemmed herb forming clumps up to 30 cm high and 50 cm across. The stems are greyish-green often red-tinged, erect, branched and 4-angled with prominent white callus scars frequent on the stem edges. Young stems are covered with disk-shaped trichomes and more or less grow straight. Inflorescences are umbellate with 15-30 flowers [2,10]. Its pedicles are slender and 1-2 cm long. The flowers are yellow in color, all about 0.8-1.0 cm in diameter. The calyxes are five-lobed and

Full list of author information is available at the end of the article. Received: 03 Apr 2018, Accepted:09 Aug 2018



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each lobe is 2-3 mm long and free almost to base. Its bright yellow to greenish yellow corollas are shortly cup-shaped below with 5 spreading narrowly triangular lobes. It has pale yellow coronas which are 1-2 mm across. Fruits are straight, narrow cylindrical bladders up to 10 cm long. Seeds are brown with a tuft of long white hairs at the tip [10-13].

Traditional Uses

The plant of *C. flava* is traditionally used in Arabian medicine to treat peptic ulcer and diabetes. It is consumed freshly by many people in the treatment of leprosy, gastric ulcer, rheumatism and as antipyretic. Drops of its juice are used for ear inflammation [13-15]. The sap inside its stems is externally applied to cuts and wounds to speed convalescence. The succulent stems are rubbed on burns to soothe them and used as a cooling agent on sunburns and itchy-skin. Fresh stems are eaten raw with an addition of lemon and salt for constipation. Tea prepared from the powdered of dried plant is taken for liver disorders, gas trouble, high blood pressure and diabetes. It is also used by the natives to suppress hunger and quench thirst [16-18].

Phytochemical studies

The phytochemistry of several species of *Caralluma* is reported with pregnane and its glycosides [19]. In common with other Caralluma species, many pregnane glycoside molecules are identified from this species. In 2015, Raees et al. and his coworkers published the first phytochemical report on C. flava in which two new pregnane glycosides, namely desmiflavasides A (1) and B (2) (Figure 2), were isolated from the plant sap [20]. Later on, the same group reported seven more new pregnane glycosides viz., desmiflavasides C (3) and D (4), nizwaside (5) and desflavasides A-D (6-9) from the same source [21-23]. Glucose, digitalose, cymarose, allomerose and thevetose sugar units were detected in these molecules. Compounds 3-8 had a pair of benzoyl groups attached to the aglycon portion. The structural characterization of all the isolates was achieved by using different multidimensional NMR techniques. 1D-TOCSY experiments were performed to determine the correlations between all protons within each sugar units. So far, no phytochemical constituent has been reported from the crude extract of the plant.

Pharmacological studies

Anti-proliferative activity

Anti-proliferative effects of isolated chemical constituents from the sap of *C. flava* were studied using MTT assays and results confirmed their worth in the treatment of cancer [20-22]. Compounds **3** and **4** showed concentration-dependent antiproliferative effects on MDA-MB231 breast cancer cells and demonstrated significant growth inhibition of the malignant cells, after 24 hours, with apparent IC_{50} values of 19.97 μ M and 25.84 μ M, respectively. Compound **5** possessed strong antiproliferative activity (IC_{50} :23.5 μ g/ml) on the same cancer cell line and was found to be more active than the standard doxorubicin. Treatment of normal breast epithelium cells (MCF10-2A) with these compounds (**3-5**) demonstrated no remarkable growth inhibition and indicated selective cytotoxicity towards the cancer cells only [21-22].

Anti-ulcer activity

In 2017, Al-Naqeb published the first report on the antiulcerative potential of the methanolic extract of *C. flava* against ethanol-induced gastric ulcer in rats model. The results showed significant anti-ulcerative effect comparable to omeprazole used as a standard. The extract reduced the incidence and severity of ulceration by 68.39% protection at a dose of 500mg/kg. Whereas, omeprazole exhibited 47.11% of inhibition at a dose of 20mg/kg. In PH examination, the treated rats with *C. flava* methanolic extract (3.74 \pm 0.42) showed significant less acidity as compared to the control rats (1.56 \pm 0.04) which revealed its protective behavior against gastric ulcer [14].

Anti-hyperglycemic activity

C. flava is reported to possess hypoglycemic activity. Its aqueous extract showed significant effects in adjusting the raised blood glucose level in streptozotocin-induced diabetic rats. After 28 days of extract administration, the blood glucose level was low-ered from 14.23 mg/100 ml to 8.45 mg/100 ml at a dose of 0.5g/kg [15].

Acute toxicity

Although, *C. flava* is consumed freshly by many people in Oman, Yemen, and the United Arab Emirates as a traditional but no scientific research was performed until recently to evaluate the toxicity of the plant. In 2017, Al-Naqeb evaluated the safety profile of *C. flava* plant for the very first time by treating rats with a single dose of its methanolic extract (3 g/kg) for 14 days. The results showed no significant toxicological symptoms and signs of abnormalities such as behavioral changes and alteration in body weight [14].

Antioxidant activity

According to a study undertaken by Karthishwaran et al., *C. flava* possesses strong antioxidant activity. The antioxidant potential of its ethanolic extract was evaluated *in-vitro* by using 2, 2'-diphenyl-1-picrylhydrazyl (DPPH), 2,2-azino-bis-3-ethylbenzothiazoline-6-sulfonic acid (ABTS), nitric oxide radi



Figure 1 Caralluma flava N.E.Br. Meve & Liede (Snapshot was taken in the mountains of Jebel Akhdar in Sultanate of Oman by Rashid Al-Harrasi; 11hr 45min/16 April 2016)



cal and hydroxyl radical scavenging assays [24]. The methanolic extract of the plant also reduced DPPH radical very strongly and the results were similar to the standard L-ascorbic acid [14].

Conclusion

The review of literature signifies the medicinal importance of *C*. *flava* which is being used as an antidiabetic and antiulcer in the

Arab traditional medicine. The phytoconstituents present in the plant are mainly pregnane glycosides which can be attributed to its medicinal actions. Further pharmacological studies of *C*. *flava* crude extract and its active compounds need to be carried out in order to confirm its medicinal potentials and also to discover its other pharmacological actions. Less information is available regarding the chemical constituents of the plant. So,

there is a good scope to isolate and identify new compounds from *C. flava* considering its wide pharmacological profile.

Acknowledgments

The author is grateful to Professor Ahmed Al-Harrasi and Mr. Mohammad Abdullah Al-Broumi for their support.

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