

## Review Article

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## A Review on King of Bitters: *Andrographis paniculata*

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### ABSTRACT

Kalmegh, The King of Bitter is traditionally used plants in ayurvedic system of medicine. Its biological name is *Andrographis paniculata* (Nees) belonging to the family Acanthaceae. From the beginning, it has been used in ayurveda as anti-inflammatory, Antioxidant, Antibacterial, Antidiabetic, Antispasmodic, anti-carcinogenic, antipyretic, antidiarrhoeal, hepatoprotective, nematocidal, anti-HIV and in many infectious diseases ranging from malaria to dysentery. Medicinal value of this plant is due to the presence of a diterpenoid derivative Andrographolide and Neoandrographolide. Traditionally, its decoction has been used as hepatoprotective in treatment of jaundice. The roots of the plant were used to prepare decoction and thus the chemical acting as hepatoprotective must be present in roots of the plant.

**Key-words:** Kalmegh, Bitter, Acanthaceae, Hepatoprotective.

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## Introduction

Kalmegh is annual to aborescent herb with intense medicinal uses. Plants is upto 1m high and with linear to lanceolate leaves. stem of plants is woody to semi woody with several cross armed branches. The stem also bears slender hard adventitious roots at nodes. Leaves are simple, opposite and exstipulate, thin membranous, lanceolate and somewhat undilated, glabrous and dark green. The plants is zygomorphic bracteate, pedicellate with pink to purple colored with purple dots. Fruit of plant is capsulated narrow from both ends, slightly flattened.

## Distribution

*Andrographis paniculata* is widely distributed herb in tropical Asian countries. Plant being gregarious grows abundantly throughout moist shady topical regions in waste grounds. However it can be seen in a variety of habitats, such as plains, hillsides, coastlines. Native populations of *Andrographis paniculata* are found to be present throughout south India and Sri Lanka. This represents the centre of origin and species diversity. The distribution of 28 species of small annual herb belonging to the genus *Andrographis* Wall. was found in tropical Asia. Only few species of kalmegh were found to possess medicinal uses. Use of kalmegh as medicinal herb has been for centuries in several Ayurvedic traditional systems of medicine. It is also extensively used in Unani and Siddha medicines as home remedy for various diseases like jaundice, hepatitis, fever in tribal medicine in India and some other countries for multiple clinical applications.

## Synonyms

Kariyat, King of bitters

## Classification of *Andrographis paniculata*

Kingdom: Plantae

Order: Lamiales

Family: Acanthaceae

Genus: *Andrographis*

Species: *A. paniculata*

## Vernacular names

Beng. - kalmegh

Eng. - Creat, King of bitters

Guj. - Kariyattu, Kiriyaati, Kirayata

Hind. - Charayetah, Mahatita, Mahatela

Punj. - Korbutti

Tel. – Nelavemu

Kan. – Nelaberu

Mar. – OliKiriaat

### **Cultivation**

It is cultivated in rainy season as Kharif crop in India. Hot and humid conditions are prerequisite as the climatic requirement for the plant with ample sunshine. *Kalmegh* does not require one particular soil and can be cultivated on ample number of soils from loam to lateritic soils with moderate fertility. Its cultivation can also be carried out on shady wastelands. Flowering of the plants begins with the onset of monsoon. The plant grows luxuriantly with the moderation in temperature after end of monsoon. The propagation of plants is done by seeds. Vegetative propagation can also be carried out for certain special cases through layering as each node is capable of producing enough roots. Seeds are small and dormancy remains for 5-6 months. Liberal use of organic manure in nursery is suggested for cultivation of healthy seedlings. The seeds should be covered with thin layer of soil and compost mixture. Suitable mulch should be used to cover the bed properly and irrigated regularly with water fountain till seedlings emerge (6-7 days). The seedlings should be raised in shade in order to protect them from heat if possible. Irrigation of the beds should be done immediately after transplanting. It can be grown on poor to moderate fertile soil but application of nitrogen fertilizers helps to increase the yield of the herb. Maximum herb yield can be obtained in 90-100 days after which leaves begin to shed. The crop keeps dormant in winter. At the time of flowering, the active principle andrographolide is maximum in leaves. Since, the whole plant consists of active principles, plant material is dried in shade and powdered. A well protected crop grown during monsoon season yields 3.5 to 4 tonnes/ha of dried plant.

### **Chemical constituents**

The principal bitter compound that makes plant King of bitters is 'Kalmeghin' which is a bitter crystalline diterpene lactone viz. andrographolide flavanoids which possess following structure. Two flavonoids identified as 5, 7, 2', 3'-tetramethoxyflavanone and 5-hydroxy-7, 2', 3'-trimethoxyflavone were identified from the whole plant, while 12 new flavonoids and 14 diterpenoids have been isolated from the aerial parts. The recent reports showed 2 new flavonoid glycosides and a new diterpenoid (andrographic acid) and two new ent-labdane diterpenoid glycosides were isolated from the aerial parts.

Thus flavanoids occur in plants mainly in roots but also in leaves in small quantity.<sup>1</sup> The main lactone derivative constituents present are Andrographalide (C<sub>20</sub> H<sub>30</sub> O<sub>5</sub>, mp 230-239°C). B, chuanxinlian A (deoxyandrographolide), C (neoandrographolide) and D (14-deoxy-11,12-didehydroandrographolide). Other lactones isolated are Ninandrographolide, Isoandrographolide, Andropanolide etc. Moreover, other constituents present are oxygenated flavones, Volatile oils; Carvacrol, eugenol, myristic acid, caffeic acid and chlorogenic acid.

## Traditional uses

*Andrographis paniculata* was reported to be used in the Ayurvedic system of medicine for decades and was found to have antibacterial, antifungal, antiviral, antioxidant, anti-inflammatory, antihyperglycaemic, hypocholesterolemic, emollient, diuretic, carminative, astringent, hepatoprotective, antipyretic and adaptogenic. It has "blood purifying" activity so it is recommended in the treatment of leprosy, gonorrhoea, scabies, boils, skin eruptions, and chronic and seasonal fevers. Juice or decoction of fresh leaves is administered to infants to comfort griping, irregular bowel habits, and loss of appetite. It is represented as so-called "bitter and cold", is expressed to be antipyretic, detoxicant, anti-inflammatory, and detumescent, and believed to mitigate "pathogenic heat" from the blood. *Andrographis paniculata* is used to treat pharyngolaryngitis, diarrhoea, dysentery, cough with thick sputum, carbuncle, sores, and snake bites. Various Ayurvedic and unani preparations and compound formulas of the herb have been used for the treatment of infectious and non-infectious diseases, and there were significant effective rates described for conditions such as epidemic encephalitis B, suppurative otitis media, neonatal subcutaneous annular ulcer, vaginitis, cervical erosion, pelvic inflammation, herpes zoster, chicken pox, mumps, neurodermatitis, eczema, and burns.

## Anti-inflammatory activity

It is reported that inflammation induced by histamine, dimethyl benzene and adrenaline was significantly reduced by dehydroandrographolide followed by neoandrographolide and andrographolide.<sup>1</sup>

## Hepatoprotective activities

In the Indian system of medicine *Andrographis paniculata* was exclusively used as hepatoprotective. Under the Ayurvedic system of medicine, it is part of many Ayurvedic formulations used to treat jaundice and hepatitis like Livokas Ayurvedic proprietary medicine. The main role of kalmegh in these preparations being a hepatoprotectant is well established in Ayurveda. Studies for kalmegh as hepatoprotective drug have been conducted. Comparative studies for hepatoprotective actions have been conducted between silymarin and andrographolide which indicates the significant choleretic effects produced by andrographolide against acetaminophen-induced reduction in volume and contents of bile have been better than silymarin. Chowdhury *et al* shows the pre and post treatment of adult rat with kalmegh extract is found to be effective as protective against ethanol-induced increase in serum transaminase. Activity of the leaf extract on evaluation revealed the protection against carbon tetrachloride-induced microsomal lipid peroxidation which was in vitro inhibited completely by the extract but not by andrographolide which indicates the activity of hepatoprotection of the kalmegh was not solely due to the presence of andrographolide.<sup>2</sup> Flavonoids, acknowledged as 5,7,2',3'-tetramethoxyflavanone and 5-hydroxy-7,2',3'-trimethoxyflavone, as well as several other flavonoids, andrographolide diterpenoids, and polyphenols have been acquired from the phytochemical investigation of the whole plant of kalmegh. With the aid of spectroscopic methods, the structures of these compounds have been established.<sup>2</sup>

### **Antimicrobial and Antiparasitic Activities**

The aqueous extract of the *Andrographis paniculata* has been found to have the antimicrobial and antiparasitic activity against bacteria, viruses etc. This activity of drug attributed to combined effect of andrographolides and arabinogalactan proteins.<sup>3</sup> Crude aqueous extract of leaves of the plant found to have similar antimicrobial activity against gram's positive *S. aureus* and methicillin resistant *S. aureus* and gram's positive *Pseudomonas aeruginosa*. However it lack in activity against *E. coli* and *Klebsiella*.<sup>4</sup> The ethanolic extract of plant found to inhibit adherence of *Streptococcus mutans* ATCC 25175 and *S. mutans* TPF-1 *in vitro* at an effective concentrations of about 0.5%.<sup>5</sup> The further investigations carried out for antibacterial activity of plant by filter plate disc-agar diffusion procedure against common bacterial pathogens such as *Pseudomonas aeruginosa*, *Clostridium perfringens*, *Serratia marcescens*, *Bacillus subtilis*, *Enterobacter aerogenes*, *Shigella flexneri*, *Staphylococcus aureus* and *Salmonella typhi*. The hexane and chloroform extract was found to inhibit most of the pathogens but the activity was found to be highest against Methicillin resistant *S. aureus*. Presence of terpenoids, steroids and coumarins were found in the extract by phytochemical screening.<sup>6</sup>

### **Anti-inflammatory and Antioxidant activity**

Various investigators reported the Anti-inflammatory activity of the plants, as the inflammation is caused by the release of histamine was found to significantly reduced by the dehydroandrographolide followed by neoandrographolide and andrographolide. This anti-inflammatory action of dehydroandrographolide was found due to its property to enhance the release of adrenocorticotrophic hormone (ACTH) from the pituitary gland which signals adrenal gland to release cortisol which in turn is potent endogenous anti-inflammatory agent.<sup>7</sup>

It was reported that the nicotine-induced suppression of mitochondrial electron chain complexes and the conclusive increase in nitric oxide (NO) in different parts of rats' brains was averted by concurrent treatment with the water and ethanol extracts of *A. paniculata* or andrographolide; the water extract was found to exhibit higher antioxidant activity than the ethanol extract. Phytochemical analysis was found to reveal higher flavonoid but lower phenol contents in water extract than in ethanol extract.<sup>8</sup>

### **Anti-Hyperglycaemic Effects**

It was reported that oral intake of Ilogen-Excel (an Ayurvedic formulation entrenched with this plant, 50 mg/kg and 100 mg/kg) for 60 days results in significant reducing of blood glucose and raising levels of plasma insulin, hepatic glycogen and total hemoglobin. The ethanolic extract of whole plant was effective for anti-hyperglycaemic property and abates oxidative stress in diabetic rats.<sup>9</sup> Moreover, after *in vitro* experiments, it was concluded that the hypoglycemic effect of *Andrographis paniculata* is due to insulin release from pancreatic cells through ATP sensitive potassium channels, similar to other insulin tropic antidiabetic agents.<sup>10,11</sup>

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