

***CALOTROPIS GIGANTEA: A COMPREHENSIVE REVIEW***

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### **Summary**

Plants used in traditional medicine represent a priceless tank of new bioactive molecules. *Calotropis gigantea* R.Br. is one of the important plant from traditional system of medicine found all over the world. *Calotropis gigantea* R.Br. Asclepiadaceae, commonly known as milkweed or swallow-wort, is a common wasteland weed, has been reported to possess potent pharmacological properties like hepatoprotective, anti-diarrhoeal, analgesic, anti-inflammatory, anti-ulcer and anti-asthmatic activities, etc. The various chemical constituents like cardiac glycosides, triterpenoids, proteases, flavonoid glycosides, triterpenoids, volatile long chain fatty acids,  $\beta$ -sitosterol and many others were identified in this plant. This review gives a bird's eye view mainly on the pharmacognostic characteristics, traditional uses, phytochemistry and Pharmacological actions.

**Keywords:** *Calotropis gigantea*, swallow-wort, phytochemistry, pharmacological actions.

### **Introduction**

There exists a plethora of knowledge and information and benefits of herbal drugs in our ancient literature of Ayurvedic and Unani medicine. One of the earliest treatises of Indian medicine, the Charaka Samhita (1000 B.C.) mentions the use of over 2000 herbs for medicinal purpose. According to the WHO survey 80% of the populations living in the developing countries rely almost exclusively on traditional medicine for their primary health care needs.

Exploration of the chemical constituents of the plants and pharmacological screening may provide us the basis for developing the leads for development of novel agents. In addition, herbs have provided us some of the very important life saving drugs used in the armamentarium of modern medicine. However, among the estimated 250,000-400,000 plant species, only 6% have been studied for biological activity, and about 15% have been investigated phytochemically.(1, 2) This shows a need for planned activity guided phyto-pharmacological evaluation of herbal drugs.

### **General Information**

*Calotropis gigantea* belonging to family Asclepiadaceae.

### **Vernacular names**

Marathi- Ruvi

Bengali – Akanda/ Gurtakand/ swetakand

English - Swallowwort/ Maddar

Gujrathi - Akdo/Aakado

Hindi – Madar/ Ag/ Ark/ak

Tamil –Aerukku/ Erukkam

Oriya- Akondo/ kotuki

Sanskrit- Aditya/arka/mandara

Telgu - Mandaramu/Jilleedudoodi(floss)/ nallajilleedu. (3, 4)

Habitat is wild throughout India, in comparatively drier and warmer areas, upto to an altitude of 1050 meters. (5)

### **Taxonomical hierarchy of *Calotropis gigantea* (6, 7)**

Kingdom - Plantae

Subkingdom - Tracheobionta

Superdivision - Spermatophyta

Division - Magnoliophyta

Class - Magnoliopsida

Subclass - Asteridae

Order - Gentianales

Family - Asclepiadoideae

Genus - *Calotropis*.

Species - *C. gigantea*

### **Botanical description**

A much – branched , hardy, erect, woody shrub, 1-5 m in height, native to India, found growing up to an altitude of 1050 m throughout India including the Andamans. Stems woody, round, tender ones covered with soft, loosely appressed, whitish, waxy or sometimes powdery pubescence; bark thick, light yellow or ash- grey, soft, corky, deeply fissured; leaves fleshy, cuneate- obovate or obovate – oblong, with a narrow cordate or often amplexicaule base, 10.0-20.0 cm X 2.5- 7.5 cm, smooth above, cottony below; flowers lilac or pale rose or purple, rarely light greenish yellow or white, in simple or compound cymose- corymbs; follicles 2 or 1 fleshy, or recurved, 7 -10 cm long ; seeds brown, numerous, broadly ovate, flattened with 2.5-3.2 cm long, white, tuft of silky hair (coma) at the pointed end. (4)



**Fig.1. Leaves & flowers of *Calotropis gigantea***



Fig.2. Flowers of *Calotropis gigantea*

### Phytochemistry

#### Root

##### Major

Mudarol, akundarol (8,9), uscharidin (10), Calotropin (10,11), Frugoside (11,12), 4'-O- $\beta$ -D-glucopyranosylfrugoside (11,12), Calotroposides A to G. (12,13)

##### Others

$\alpha$ -Amyrin,  $\beta$ -amyrin, taraxasterol and its  $\Psi$ -isomer,  $\beta$ -sitosterol,  $\alpha$ -amyrin methylbutazone,  $\beta$ -amyrin methylbutazone,  $\alpha$ -amyrin acetate,  $\beta$ -amyrin acetate, taraxasteryl isovalerate, taraxasteryl acetate (14), calotroplupenyl acetate A, lupeol acetate B, gigantursenyl acetate A, gigantursenyl acetate B. (12,15)

#### Flowers

##### Major

$\alpha$ -Calotropeol,  $\beta$ -Calotropeol, (9) rutin. (16)

##### Others

$\beta$ -amyrin, hyperoside.(16)

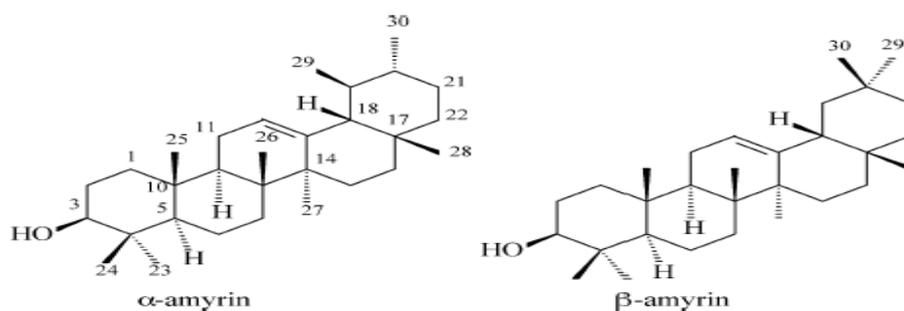
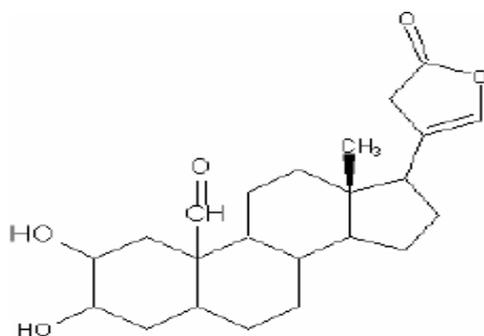
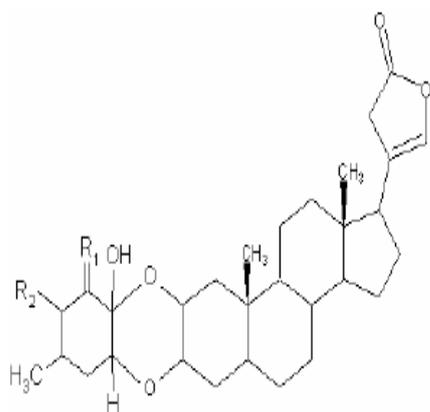
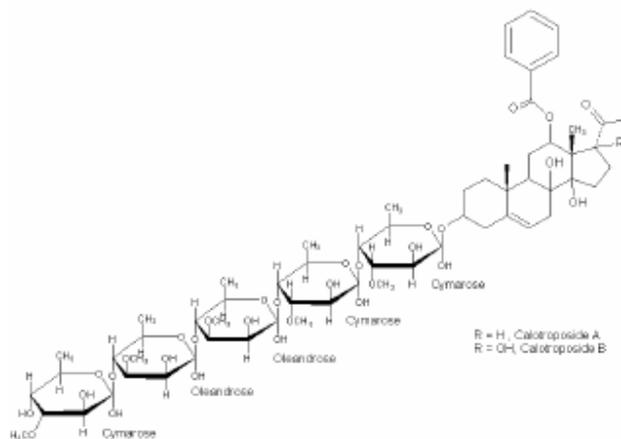


Fig.3. Structure of  $\alpha$ -amyrin and  $\beta$ -amyrin



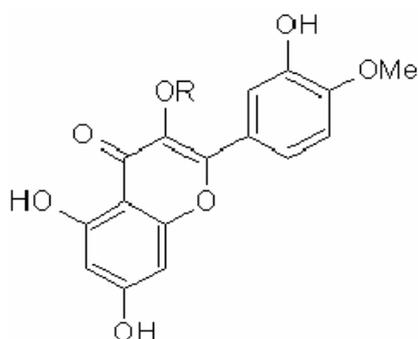
Calotropogenin (17)

**9-Calotroposide A, 10- Calotroposide B (18)**



Structure No.	Nomenclature	R <sub>1</sub>	R <sub>2</sub>
2	Calotropin	$\alpha$ - OH, $\beta$ - H	H
3	Uscharin	$\left\{ \begin{array}{l} S-CH_2 \\ N-CH_2 \end{array} \right.$	H
4	Calotoxin	$\gamma$ -H, $\gamma$ -OH	H
5	Calactin	$\alpha$ - H, $\beta$ - OH	H

(17)



(19)

Structure No.	Nomenclature	R
11	Isorhamnetin -3-O-rutinoside	Glu-ORha O Gal
12	Isorhamnetin 3-O-glucoside	Glucose
13	Isorhamnetin rhamnoglucoside	Glucose-O-Rhamnose

## Pharmacological Actions

### Analgesic activity

The ethanolic extract of *C. gigantea* flowers showed significant analgesic activity in the Eddy's hot plate method (20, 21) and acetic acid induced writhing. (22, 23) In acetic acid induced writhing test, an inhibition of 20.97% and 43.0% in the number of writhes was observed at the doses of 250 and 500 mg/kg, respectively. In the hot plate method the paw licking time was delayed. The analgesic effect was observed after 30 min of dose administration which reached its maximum after 90 min. The ethanolic extract of *C. gigantea* flower produced a significant decrease in the number of writhings and paw licking time.

### CNS activity

#### 1. Assessment of analgesic activity

##### a. Hot plate method.

Alcoholic extract of peeled roots of *C. gigantea* R.Br. (Asclepiadaceae) was tested orally in albino rats at the dose level of 250 and 500 mg/kg body weight for CNS activity. Prominent analgesic activity was observed in Eddy's hot plate method. The paw licking time was delayed. (24)

##### b. Acetic acid induced writhing.

Alcoholic extract of peeled roots of *C. gigantea* R.Br. (Asclepiadaceae) was tested orally in albino rats at the dose level of 250 and 500 mg/kg body weight for CNS activity. Prominent analgesic activity was observed in acetic acid induced writhings and the numbers of writhings were greatly reduced. (25, 26)

## **2. Assessment of locomotor activity**

Alcoholic extract of peeled roots of *C. gigantea* R.Br. (Asclepiadaceae) was tested orally in albino rats at the dose level of 250 and 500 mg/kg body weight for CNS activity and there was a decrease in the locomotor activity. (27,28)

## **3. Assessment of antianxiety activity**

Alcoholic extract of peeled roots of *C. gigantea* R.Br. (Asclepiadaceae) was tested orally in albino rats at the dose level of 250 and 500 mg/kg body weight for CNS activity. The extract treated rats spent more time in the open arm of elevated plus maze (EPM) showing its antianxiety activity. (28, 26)

## **4. Assessment of skeletal muscle relaxant activity (motor coordination)**

Alcoholic extract of peeled roots of *C. gigantea* R.Br. (Asclepiadaceae) was tested orally in albino rats at the dose level of 250 and 500 mg/kg body weight for CNS activity and the fall off time (motor coordination) was also decreased. (27,28)

## **5. Assessment of anticonvulsant activity**

Alcoholic extract of peeled roots of *C. gigantea* R.Br. (Asclepiadaceae) was tested orally in albino rats at the dose level of 250 and 500 mg/kg body weight for CNS activity. Significant anticonvulsant activity was seen as there was a delay in the onset of pentylenetetrazole induced convulsions as well as decrease in its severity. (27, 28)

## **6. Repellant activity**

Repellant activity of the milkweed plant, *C. gigantea* R.Br. was evaluated against important storage pests such as *Callosobruchus maculatus* (Fab.), *Sitophilus oryzae* Linn, and *Tribolium castaneum* (Herbst). Leaf, flower, stem, root and whole plant were Soxhlet extracted by using petroleum ether solvent and repellency test was carried out using glass olfactometer. Whole plant extract at 5 % concentration had maximum repellent effect followed by leaf, flower, stem and root extracts. (29)

## **7. Pregnancy interceptive activity**

The ethanolic extract of the roots of *C. gigantea* Linn. exhibited 100% pregnancy interceptive activity in rats when administered as a single oral dose of 100 mg/kg on Day 1 postcoitum. The extract also exhibited 100% efficacy at the dose of 12.5 mg/kg when administered in the Days 1–5 and 1–7 postcoitum schedules. When administered during the peri-cum-early postimplantation period (i.e., Days 5–7 postcoitum at 250 mg/kg), most of the implantations showed signs of resorption. On fractionation, the chloroform fraction showed 100% activity at 100 mg/kg in the single-day (Day 1 postcoitum) schedule, whereas the hexane, n-butanol-soluble and n-butanol-insoluble fractions were found to be inactive at this dose. At autopsy on Day 10 postcoitum, 7–25% loss in body weight was recorded at the minimum effective contraceptive dose (MED) in rats treated with the ethanolic extract as well as its chloroform-soluble fraction on Days 1–7, 1–5 and

1 postcoitum, in comparison with the 6–7% increase in body weight observed in vehicle control rats. There was however no mortality in any of the treatment groups. The active ethanolic extract and its chloroform fraction were devoid of any estrogen agonistic or antagonistic activity at their respective MEDs in the ovariectomized immature rat bioassay. Efforts are being made to isolate the active principles devoid of effect on body weight. (30)

### **Traditional uses**

The plant is analgesic, purgative, alexipharmic, anthelmintic; cures leprosy, leucoderma, ulcers, tumours, piles, diseases of the spleen, the liver, and the abdomen; the juice is anthelmintic and laxative; cures piles and “kapha”. The root bark is diaphoretic; cures asthma and syphilis. The flower is sweet, bitter, anthelmintic, analgesic, astringent. The milk is bitter, heating, oleaginous, purgative; cures leucoderma, tumours, ascites, diseases of the abdomen. (31), cures toothache and earache (32, 33), in sprain (34), in anxiety and pain (35, 36), in epilepsy (37) and in mental disorders. (38)

### **Conclusion**

Medicinal plants have provided copious leads to combat diseases, from the dawn of civilization. Herbal medicines are in great demand in the developed as well as developing countries for primary healthcare because of their wide biological and medicinal activities, higher safety margins and lesser costs. (39, 40) The extensive survey of literature revealed that *C. gigantea* is important medicinal plant with diverse pharmacological spectrum. The plant shows presence of many chemical constituents, which are responsible of the various activities of the plant. *C. gigantea* embibing a tremendous potential deserves a special attention of the scientific fraternity to emerge as a milestone for medical science of this millennium due to its various medicinal uses. Further evaluation needs to be carried out on *C. gigantea* in order to explore the concealed areas and their practical clinical applications, which can be used for the welfare of the mankind.

### **References**

- 1) Balandrin MF, Klocke JA, Wrtle ES, Boilinger WH. Content and purity of extract solasodine in some available species of Solanum. Sci & Culture 1985; 56 (5): 214-216.
- 2) Cragg GM, Newman DJ, Sander KM. Natural products in drug discovery and development. J Nat Prod 1997; 60: 52-60.
- 3) GuptaAK. Quality standards of Indian medicinal plants. Indian council of medicinal research, New delhi 2005; Vol- 2:35.

- 4) The Wealth of India, A Dictionary of Indian Raw Materials and Industrial Products, Revised edition, by Publication and Information Directorate, CSIR, Hillside road, New Delhi. Vol-3: 78.
- 5) Khare CP. Encyclopedia of Indian medicinal plants, Rational western therapy, Ayurvedic & other traditional usage, Botany. Springer publication: 120.
- 6) "[http://en.wikipedia.org/wiki/Calotropis\\_gigantea](http://en.wikipedia.org/wiki/Calotropis_gigantea)".
- 7) The Ayurvedic pharmacopoeia of India, I<sup>st</sup> edition, part-I, 1999; vol –II: 169-170.
- 8) Mutri PBR, Seshadri T. Wax and resin components of *Calotropis gigantea* Part III Root bark. Proc Acad Sci 1945; 21A:147-154.
- 9) Mutri PBR, Seshadri T. Chemical composition of *Calotropis gigantea* Part VI. Flowers. A comparison of the composition of the various parts of the plant. Proc Indian Acad Sci 1945; 22 A:304-309.
- 10) Datta SK. Separation and HPLC – identification of two cardiac glycosides from *Calotropis gigantea* (Linn.) R. Br. ex Ait. Indian Drugs 1988; 25:167-168.
- 11) Kiuchi F, et al. Cytotoxic principles of a Bangladeshi crude drug, Akond Mul (roots of *Calotropis gigantea* L.). Chem pharm Bull 1998; 46:528-530.
- 12) Kitagawa I, et al. Indonesian medicinal plants I. Chemical structures of calotroposides A and B, two new oxypregnane- Oligoglycosides from the roots of *Calotropis gigantea* (Asclepiadaceae). Chem pharm Bull 1992; 40: 2007-2013.
- 13) Shibuya H, Indonesian medicinal plants V. Chemical structures of calotroposides C, D, E, F, and G, five additional new oxypregnane- Oligoglycosides from the roots of *Calotropis gigantea* (Asclepiadaceae). Chem pharm Bull 1992; 40: 2647-2653.
- 14) Anjaneyulu V and Row LR. The triterpenes of *Calotropis gigantea* Linn. Curr sci 1968; 6: 156-157.
- 15) Mohd A and Gupta J. New pentacyclic triterpenic esters from the root of *Calotropis gigantea*. Indian J Chem 1999; 38B:877-881.
- 16) Subramanian SS and Nair AG R. Flavonoids of some Asclepiadaceous plants. Phytochem 1968; 7:1703-1704.
- 17) Seiber N, Caroltn J, Nelson J, Mark Lee S. Cardenolides in the latex and leaves of seven *Asclepias* species and *Calotropis procera*. Phytochem 1982; 9: 2243-2248.
- 18) Kitagawa J, et al. Indonesian medicinal plants. I. Chemical structures of calotroposides A and B, two new oxypregnane-oligoglycosides from the root of *Calotropis gigantea* (Asclepiadaceae). Chem Pharm Bull 1992; 40(8): 2007-2013.

- 19) Sen S, Niranjana S, Sahu P, Mahato S. Flavonol glycosides from *Calotropis gigantea*. *Phytochem* 1992; 31(8): 2919-2921.
- 20) Eddy NS. Analgesic activity of *Calotropis gigantea* flowers. *J Pharmacol* 1932; 45:339.
- 21) Kamath JV and Rana AC. Pharmacological activities of ethanolic extract of *Calotropis procera* roots. *Indian Drugs* 2003; 40:29.
- 22) Sonavane GC, Sarveiya V, Kasture V, Kasture BS. Behavioural actions of *Myristica fragrans* seed. *Indian J Pharmacol* 2001; 33: 417-424.
- 23) Dewan S, Sangraula H and Kumar VL. Preliminary studies on the analgesic activity of latex of *Calotropis procera*. *J Ethnopharmacol* 2000; 73: 307.
- 24) Kamath JV, Rana AC. Pharmacological activities of ethanolic extract of *Calotropis procera* roots. *Indian Drugs* 2003; 40: 292–295.
- 25) Dewan S, Sangraula H, Kumar VL. Preliminary studies on the analgesic activity of latex of *Calotropis procera*. *J Ethnopharmacol* 2000; 73: 307–311.
- 26) Sonavane GC, Sarveiya V, Kasture V, Kasture BS. Behavioural actions of *Myristica fragrans* seed. *Indian J Pharmacol* 2001; 33: 417–424.
- 27) Kulkarni SK. *Handbook of Experimental Pharmacology*, 3rd ed. Vallabh Prakashan, New Delhi. 1999; 133–137, 191, 199–223.
- 28) Turner RA. *Screening Methods in Pharmacology*. Academic Press, New York. 1965; 70, 75, 78, 164.
- 29) Arulprakash R and Veeravel R. Studies on the repellent properties of *Calotropis gigantea* R. Br. Plant parts against important storage insect pests. *Madras Agric J* 2005; 92 (4-6): 308-310.
- 30) Srivastava SR, Keshria G, Bhargava B, Singh C, Singha MM. Pregnancy interceptive activity of the roots of *Calotropis gigantea* Linn. in rats. *Contraception* 2007; 75: 318– 322.
- 31) Kirtikar KR and Basu BD. *Indian Medicinal Plants*. Sudhindra Nath Basu, Allahabad. 1995; 3: 1609.
- 32) Aminuddin and Girach RD. Observations on ethnobotany of the Bhunjia—a tribe of Sonabera plateau, Kalahandi, Orissa. *Ethnobotany* 1993; 5: 84.
- 33) Allen TF. *Handbook of Materia Medica and Homeopathic Therapeutics*. Jain Publishers (P) Ltd., New Delhi. 1994: 251.
- 34) Manandhar MP. Folk-lore medicine of Chitwan district, Nepal. *Ethnobotany* 1990; 2: 33.

- 35) Boericke W. Pocket Manual of Homeopathic Materia Medica and Repertory. Jain Publishers (P) Ltd., New Delhi 1999:157.
- 36) Sharma V. Dravyaguna V, Chaukhambala B. Academy, Varanas. 2001; Vol. 2: 435.
- 37) Jain SK, Sinha BK, Saklani A. Medicinal plants known among tribal societies of India. Ethnobotany 1989; 1: 92.
- 38) Upadhyaya AS, Vartak VD, Kumbhojkar MS. Ethnomedicobotanical studies in western Maharashtra, India. Ethnobotany 1994; 6: 28.
- 39) Chattopadhyay RR and Bhattachryya SK. *Terminalia chebula*: An update. Pharmacog Rev 2007; 1(1): 151-157.
- 40) Shinde VM and Dhalwal K. Pharmacognosy: Changing Scenario. Pharmacog Rev 2007 1: 1-6.