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TINOSPORA CORDIFOLIA: ETHNOBOTANY, PHYTOPHARMACOLOGY AND PHYTOCHEMISTRY ASPECTS

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ABSTRACT: Tinospora cordifolia (Thunb.) Miers, (Guduchi) is an evergreen perennial climber. This deciduous and dioecious plant belongs to the family Menispermaceae. It is a plant of significant medicinal importance in the Indian system and designated as Rasayana. Its efficacy has been also recognized by the modern system of medicine. The whole plant is used medicinally however; the stem is approved for use in medicine. This is due to higher alkaloid content in the stems than in the leaves. This plant has been known to possess immunomodulatory, hypoglycaemic, antioxidant, antihyperglycaemic, antiallergic, anti-inflammatory, diabetes hypogycemia and several other properties also. The plant mainly contains alkaloids, glycosides, steroids, diterpenoid lactones, sesquiterpenoid, aliphatic compound and other miscellaneous compound. The present review is an attempt to highlight the various ethnobotanical and traditional uses as well as the various pharmacological, phytochemical and clinical reports on T. cordifolia.

INTRODUCTION: *Tinospora cordifolia* (Thunb.) Miers, (Guduchi) is one of the important dioecious plants. In Hindi, the plant is commonly known as *Giloe* ¹ which is a Hindu mythological term that refers to the heavenly elixir that has saved celestial beings from old age and kept them eternally young. In Ayurveda, it is designated as Rasayana drug recommended to enhance general body resistance, promote longevity and as antistress and adaptogen ²⁻³. The fact that it is called "Amrita" signifies its use for revitalization and its importance in Ayurveda. This significant plant is also mentioned in important Pharmacopoeias ⁴⁻⁶.

Distribution: It is indigenous to areas of India, Myanmar, Sri Lanka, China, Thailand, Philippines, Indonesia, Malaysia, Borneo, Vietnam, Bangladesh, North Africa, West Africa, and South Africa ⁷⁻¹⁰. It typically grows in deciduous and dry forests at elevations up to 1000ft.

Origin and Habitat: *T. cordifolia* is a climbing shrub native to lower elevation in tropical areas of the Indian subcontinent and climbs numerous types of trees¹¹. It prefers wide range of soil, acid to alkaline and it needs moderate level of soil moisture.

The Family: *T. cordifolia* belongs to the family Menispermaceae ¹²⁻¹⁴ which consists of about 70 genera and 450 species that are found in tropical lowland regions. They are generally climbing or twining, rarely shrubs. This family is a rich source of alkaloid and terpenes ¹⁵.

The Genus: The genus *Tinospora* Miers (Menispermaceae) has about 32 species distributed in tropical Africa, Madagascar, Asia to Australia and the Pacific Islands ¹⁶⁻¹⁸. In India, the genus is represented by four species; two species, *T. cordifolia* (Thunb.) Miers and *T. sinensis* (Lour.) Merr., are known to occur

in South India and other two *T. crispa* (L.) Hook.f. & Thomson and *T. glabra* (Burm.f.) Merr., are reported from Northeast India and the Andaman Islands ¹⁹.

The Species: Tinospora cordifolia (Thunb.) Miers

Synonyms:

- 1. *Menispermum crispum* Linnaeus, Sp. Pl., ed. 2, 2: 1468. 1763
- 2. Tinospora gibbericaulis Handel-Mazzetti
- 3. Tinospora mastersii Diels
- 4. Tinospora rumphii Boerlage
- 5. Tinospora thorelii Gagnepain.

Vernacular Names: The common names are Gilo (Arabic); Amarlata (Assamese); Gadancha, Guluncha, Giloe (Bengali); K'uan chu Hsing (Chinese); Culancha (French); Tinospora (English); Gado, Galo, Gulo (Gujerati); Giloe, Gulbel, Gurcha (Hindi); Amrytu, Sittamrytu (Malayalam); Ambarvel, Giroli, Gulvel (Marathi), Garjo (Nepali); Gulancha (Oriya); Gulbel (Persian); Gilo (Punjabi, Kashmiri), Amrita, Guduchi, (Sanskrit); Gurjo (Sikkikim); Amridavalli, Niraidarudian (Tamil); Guduchi, Iruluchi (Telugu) and Guruch (Urdu)²⁰⁻

Botanical Description: *T. cordifolia* is a large extensively spreading glabrous, dioecious perennial deciduous climber, grows on wide range of hedges and trees. It is reported to bear distinct male and female flowers ²¹⁻²³. Its stem, when fresh, have a green succulent bark covered by a thin brown bark and are studded with warty lenticels when dry, the stem shrinks and the bark separate from the wood. Branches are sending down slender pendulous fleshy roots, terete, striate, with tubercled, pale sometimes shining or glabrous bark.

Leaves membranous, 7-9 nerved, 5-10 cm, roundish, cordate or heart shaped (giving name *cordifolia* to the plant) with a 2.5- 7.0 cm petiole. The flower bloom in summer. Racemes is rather lax, 5.0 cm, elongating and often longer than leaves. The male flowers are small, yellow or green in colour, and occur in clusters in the axils of small subulate bracts. Sepals are 6, 3 outer very small, ovate-oblong, acute, the inner 3 larger, membranous, broadly elliptical, concave, yellow.

Petals are 6, equal, broadly spathulate, each loosely embracing a stamen, claw cuneate, reflexed to apex, pistillode. Female flowers usually solitary, similar to male, but sepals green, margins not reflexed, staminode short, linear. Carpels 1-3, widely separated on the short fleshy gynophores, dorsally convexed, and scarlet. The fruit are the size and shape of a large pea and turn from green to red when ripe in winter and mucilaginous ²⁴.

Nutritive Composition of *Tinospora cordifolia*: *T. cordifolia* contains high fibre (15.9%), sufficient protein (4.5%-11.2%), sufficient carbohydrate (61.66%), and low fat (3.1%). Its nutritive value is 292.54 calories per 100 g. It has high potassium (0.845%), high chromium (0.006%), sufficient iron (0.28%) and sufficient calcium (0.131%), important in various regulatory functions ²⁵.

Properties – in Ayurveda: Ayurveda advocates the treatment for the entire body as a single unit. With modern medicine it gives a synergistic activity and antagonises and minimise the toxicity of modern drug. T. cordifolia referred to as Guduchi (plant which protects from diseases, Sanskrit), has been described in ancient textbooks of Ayurveda including Sushrut Samhita and Charak Samhita. Other synonyms used for Guduchi which refer to its various properties and uses, include Chhinnaruha/Chhinnodbhava (for the plant's capacity to grow from the cut side), Vatsadini (eaten by grazing animals), Pittaghni (bile destroying), Amruta (imparts immortality), Rasayana (capacity to improve quality of rasa, the primordial tissue which in turn strengthens all other tissues), Chakrangi/Chakraakshana (for its wheel-like appearance), Jwaranashi/ Jwarari (potent antipyretic), Vayastha (prevents ageing), Amrutsambhava (ambrosia), Bhishakpriya (favourite of physicians), Saumya (not harmful) and Tikta (bitter taste).

The stems, leaves, and roots are used in medicine ^{11, 26-27}. In Ayurveda, *T. cordifolia* is used as "rasayana" which in Sanskrit implies circulation of "rasa"- the nutrient. The ancient Indian physician, Charaka described rasayana as anti aging, increasing the life span, promoting intelligence, improving memory and freedom from diseases, indicating an immunostimulant effect ²⁸ and also used as an antispasmodic and an anti-diarrhoeal agent ²⁹.

Herbalist Sebastian Pole writes that "those growing up neem trees are said to be the best synergy between these two bitter plants enhances guduchi's efficacy." It is a traditional belief that *Guduchi satva* obtained from the *Guduchi* plant growing on *neem* tree (*Azadirachta indica*) is bitterer and more efficacious and is said to incorporate the medicinal values of *neem* ³⁰.

T. cordifolia has been described in Ayurveda in various dosages form. These include Swaras (juice from the fresh stem: 10-20 mL/day), Kalka (paste of fresh stem: 10 g/day), Churna (powdered dry stem: 1-3 g/day), Kwatha (hot water extract from ground dried stem: 20-30 mL two to three times a day), Fant (hot water infusion: 10-20 mL/day), Arishta (stable processed formulation from a decoction of T. cordifolia containing self-generated alcohol), Satwa (sedimented starchy extract of the stem: 750 mg to 2 g/day) Ghana (solidified aqueous extract: 500 mg to 1 g three to four times a day), along with lipid formulations of T. cordifolia processed in ghee or oil (Guduchi Ghrita: 10-20 g/day) and Guduchi Taila (for external application) ³¹. Guduchi is recognized in the Indian Pharmacopoeia and also used in several compound formulations as an ingredient for the treatment of general weakness,

fever, dyspepsia, dysentery, gonorrhoea, secondary syphilis, urinary diseases, impotency, gout, viral hepatitis, skin diseases, and anaemia ³²⁻³⁴, leprosy, fever, asthma, anorexia, jaundice, gout, skin infections, diabetes, chronic diarrhoea, dysentery ³⁵⁻⁴².

Ethnobotanical, Folk and Tribal Uses: There are over 400 different tribal and other ethnic groups in India. Each tribal group has its own tradition, folk language, beliefs and knowledge about the use of natural resources as medicines ⁴³. T. cordifolia finds a special mention for its use in tribal or folk medicine in different parts of the country. Almost all the parts of the plant are documented to be useful in ethno botanical surveys conducted by ethno botanists 44-45. In folk and tribal medicine, whole plant, powdered root and stem bark, decoction of root and stem, juice of the root, juice or paste of the leaves, and stem of the T. cordifolia are being used to treat various ailments viz. fever, jaundice, diarrhoea, dysentery, general debility, cough, asthma, leucorhea, skin diseases, fractures, eye disorders, bites of poisonous insects, venomous snake etc. Ethnobotanical uses of T. cordifolia are given in Table 1.

TABLE: 1. ETHNOBOTANICAL USES OF TINOSPORA CORDIFOLIA

S. No.	Plant part	Ethnobotanical uses		
1	Leaves	Used in the treatment of gout and ulcer		
2	Stem	The stem is bitter stomachic, stimulates bile secretion, diuretic, enriches the blood, cures jaundice, useful in skin diseases, The juice is useful in diabetes, vaginal and urethral discharges low fevers and enlarged spleen (Stem as an infusion) used to drunk as a vermifuge, jaundice, against intestinal worms (Stem as decoction) used for washing sore eyes and syphilitic sores, antipyretic, antimalarial Starch (statue) obtained from stem used for chronic diarrhoea and some form of obstinate chronic dysentery, deal with intestinal problems and improve digestion 48		
3	Stem +Root	Combination with other drugs as an antidote to snake bite and scorpion sting 49-51		
4	Fruit	Dried fruit with ghee or honey used as tonic and treatment of jaundice and rheumatism.		
5	Bark	Anti-allergic, anti-spasmodic, anti-leprotic 52-54		
	NS	Urinary diseases, syphilis, skin diseases, bronchitis 55		
	NS	Promote longevity and increase body's resistance ⁵⁶ . Stimulate the immune system ⁵⁷⁻⁵⁹		

NS=not specify

Pharmacology: As evident from the ayurvedic and ethnobotanical reports, *T. cordifolia* has various medicinal uses. In present times, the extracts/fractions (mainly aqueous and alcoholic sometimes methanolic, chloroform, ether, petroleum ether, acetone, successive, bitter or non-bitter and other fractions)

even chemical constituents of this drug have been subjected for numerous pharmacological, pre-clinical and clinical investigations. The medicinal traits and biological activities endorsed are presented in **Table 2**.

TABLE 2: PHARMACOLOGICAL ACTIVITIES OF TINOSPORA CORDIFOLIA

Immunomodulatory 57,60-68	Antioxidant 10,69-75	Anti-hyperglycaemic 76-79
Anti- diabetic 80-82	Antimalarial ⁸⁰⁻⁸²	Anti-inflammatory 43,53,73,83-88
Antitumor 53,64,72,89,90-92	Anti-allergic ^{73, 93}	Antineoplastic activity 89,94
Antipyretic 53,72,80-82	Antiamoebic, Antihelmentic 53, 80-82	Anti hyperlipidimic ⁹⁵
Immunobiological activity 96-99	Antigout, Antiasthmatic 80-82	Antiasthmatic , Antigonorrhoeal 80-82
Antigonorrhoeal 80-82	Antiperiodic ⁸⁰⁻⁸²	Antiperiodic 80-82
Mental disorder ¹⁰⁰	Androgenic ¹⁰¹	Radioprotective ¹⁰²
Antifertility effects ¹⁰³	Anticoagulant ⁷²	Antiemetic and Antiicteric 80-82
Antigonorrhoeal 80-82	Antiatherogenic 53,80-82	Diuretic ¹⁰⁴
Anticancer 89,105-107	Learning and memory enhance 108	Anti-depression ¹⁰⁹
Anti-stress 110-113	Anti-ischemic ¹¹⁴	Anti hyperlipidimic ⁹⁵
Hypolipidaemic 72, 115-116	Antileprotic 43	Antispasmodic 43,117-118
Antimicrobial ⁷²	Antiulcer 119-120	Analgesic 53,80-82
Help to dissolve urinary calculi 121	Malaria ¹²²	Obstructive jaundice 123-124
Infections 125	Hepatoprotective ^{64,126}	Prevent hepatotoxicity 127-128
Phagocytes ¹²⁹	Hepatic and splenic injury 130,131	Diabetes mellitus ¹³²

Phytochemistry: A variety of constituents have been isolated from *T. cordifolia* plant. They belong to different classes such as alkaloids, diterpenoid lactones, glycosides, steroids, sesquiterpenoid, phenolics, aliphatic compounds and polysaccharides.

Three major groups of compounds; protoberberine alkaloids, terpenoids and polysaccharides are considered as putative active constituents of *T. cordifolia* ⁴⁶⁻⁴⁷. These chemical constituents are listed in **Table 3** and their role in various therapeutic actions is listed in **Table 4**.

TABLE: 3. PHYTOCHEMISTRY OF TINOSPORA CORDIFOLIA

Part	Chemical type	Active principle	Ref.
	Alkaloids	Berberine, palmatine D, choline D, tinosporine, Magnoflorine, tetrahydropalmatine, isocolumbin	133-138
		18-norclerodane glycoside	
Stem	Glycosides	Furanoid diterpene glycoside	
		Tinocordiside	142-143
		Syringin	144-145
		Syringin-apiosylglycoside,	139-140
		Tinocordifolioside, cordioside, cordifolioside A, cordifolioside B, palmatoside C31,	141,143-144, 146-148
		palmatoside F31, cordiofoliside B2, cordifoliside D2, cordifoliside	
	Sesquiterpenoid	Tinocordifolin	150
Root	Alkaloid	Palmatine	136,149
Aerial parts	Steroids b-sitosterol, d-sitosterol, g-sitosterol b-hydroxyecdysone, ecdysterone, makisterone giloinsterol jateorine, columbin		151-152
	Diterpenoid lactones	Furanolactone, tinosporon , columbin	153-155
Whole	Aliphatic compound	Octacosanol, heptacosanol	156
plant	Miscellaneous compound	Nonacosan-15-one 3, (a,4-dihydroxy-3-methoxy-benzyl)-4-(4-hydroxy-3-methoxy-benzyl)-tetrahydrofuran, Tinosponidine, 6 cordifol, 6 Cordifelone, 6 Jatrorrhizine	157-158

TABLE 4: MECHANISM OF ACTION OF SOME IMPORTANT PHYTOCHEMICALS 159-172

Phytochemicals	Activity	Mechanism of action
Quinones	Antimicrobial	Binds to adhesions, complex with cell wall, inactivate enzymes
	Antimicrobial	Complex with cell wall, binds to adhesins Inhibits release of autacoids and prostaglandins,
Flavanoids	Antidiarrhoeal	Inhibits contractions caused by spasmogens, Stimulates normalization of the deranged water transport across the mucosal cells, Inhibits GI release of acetylcholine
	Antimicrobial	Binds to adhesions, enzyme inhibition, substrate deprivation, complex with cell wall, membrane disruption, metal ion complexation
Polyphenols and Tannins	Antidiarrhoeal	Makes intestinal mucosa more resistant and reduces secretion, stimulates normalization of deranged water transport across the mucosal cells and reduction of the intestinal transit, blocks the binding of B subunit of heatlabile enterotoxin to GM1, resulting in the suppression of heat-labile enterotoxin-induced diarrhea, astringent action
	Anthelmintic	Increases supply of digestible proteins by animals by forming protein complexes in rumen, interferes with energy generation by uncoupling oxidative phosphorylation, causes a decrease in G.I. metabolism
Coumarins	Antiviral	Interaction with eukaryotic DNA
Terpenoids and	Antimicrobial	Membrane disruption
essential oils	Antidiarrhoeal	Inhibits release of autocoids and prostaglandins
	Antimicrobial	Intercalates into cell wall and DNA of parasites
	Antidiarrhoeal	Inhibits release of autocoids and prostaglandins
Alkaloids	Anthelmintic	Possess anti-oxidating effects, thus reduces nitrate generation which is useful for protein synthesis, suppresses transfer of sucrose from stomach to small intestine, diminishing the support of glucose to the helminthes, acts on CNS causing paralysis
Lectins and Polypeptides	Antiviral	Blocks viral fusion or adsorption, forms disulfide bridges
Glycosides	Antidiarrhoeal	Inhibits release of autocoids and prostaglandins
	Antidiarrhoeal	Inhibits histamine release in vitro
Saponins	Anticancer	Possesses membrane permeabilizing properties
	Anthelmintic	Leads to vacuolization and disintegration of teguments
Steroids	Antidiarrhoeal	Enhance intestinal absorption of Na+ and water

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