

Review on Pharmacological and Therapeutic Potential of *Cuscuta* species

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Received: October 16, 2021; **Published:** November 19, 2021

Abstract

The plant genus *Cuscuta*, belongs to the family Cuscutaceae, has been traditionally used as therapeutic treasure against various metabolic and neurological diseases such as mental abstraction, headache, spleen disease, jaundice, diabetes mellitus and hypertension. There are numerous secondary metabolites have been identified that are considered as phytochemical constituents, such as alkaloids, flavonoids, lignins, steroids, phenolic acids, fatty acids, glycosides and polysaccharides. Due to the presence of these bioactive ingredients, various ethnopharmacological actions including anticancer, antiviral, antispasmodic, antihypertensive, antibacterial, antioxidant, diuretic, antihypertensive were explored. This review brings eyesight for the explorations of novel therapeutic agents for the various ailments of mankind.

Keywords: Therapeutic treasure; Phytoconstituents; Pharmacological activity

Introduction

Dodder (*Cuscuta*) is a parasitic plant; having attraction on wide range of plants like agricultural and horticultural crops such as alfalfa, lespedeza, flax, potatoes, chrysanthemum, vine, ivy and petunias [1]. *Cuscuta* is a group of 100-70sp., they are yellow, orange, red or green parasitic plants. *Cuscuta reflexa* commonly known as dodder is also called as devil's hair, witch's hair, love vine, amarber or akashabeta. *C.reflexa* is a homoparasite and also an extensive climber. It contains low level of chlorophyll and photosynthetic activity; completely depends on the host for its energy. Dodder has the capacity to recognize its host plant and also choose an appropriate host between other plants by some volatile compounds that are released by the host plants [2]. *C.reflexa* imbibes nutrients through the vascular tissue of the host plant and grows on it [3]. According to WHO, 80 percent of the world population now using herbal medicines. In ancient Chinese and Japanese medicine, dodder (*Cuscuta planiflora*) seeds are grained in to fine powder and converted to a tablets or encapsulated form to treat osteoporosis, osteoarthritis and general muscular pains [4]. Dodder was now used in Ayurveda to treat jaundice: Some researchers found that dodder possessed antidepressant, anticonvulsant, antibacterial, cytotoxic and hepatoprotective effect. Dodder has some phytochemical substances like phenols, aromatic substances. *C.reflexa* is a rootless, leafless plant and it twinned into other plants for gaining their nutrients. It was commonly called as akashvalli or dodder. Even though wide spread occurrence of Dodder, more than 6 species are found in India. It will grow only on thorny or shrubs and sometimes [5]. It will spread one host to another host by haustorium a special branched organ. Haustorium will penetrate the host cell, which will absorb the water and other nutrients from the host plant. It doesn't attach to the ground and growing with the help of seeds with hard coating and even the seeds can withstand for more than 10 years [6]. This *Cuscuta reflexa* species is investigated for antitumor, anticonvulsant, antioxidant, and induced alopecia activities along with it poses some chemical constituents caucutinin, amarbelin, beta-sterol, stigmaterol, bergenin, etc. The genus *Cuscuta* L. is the one of the herbal constituents in pharma, foods and curative tonics that is used to nourish various body parts [7] (Table 1).

Botanical Identification [9]	Synonyms	Common names
Kingdom : Plantae		
Subkingdom : Tracheobionta	Tamil : verillakothan	Arabic : Hamoul, Shubbak, Dabaie
Superdivision : Spermatophyta	English : Dodder plant	French : Cuscutae fleurs planes, cuscutea petites fleurs.
Division : Angiosperms	Hindi : Amarabela	Germany : klee-seide; ouendel-seide; Thymin-seide.
Class : Eudicots	Sanskrit : Akashvalli, Amaravalli, Khavalli.	Italy : Cuscuta Del trifoglio; cuscutea piccolo.
Subclass : Asterids	Punjabi : Zarbut	Portuguese : (ipo-de-chumbo, Espaguete
Order : Solanales	Malayalam : Moodillathal	Spanish : Cuscuta Del trebol, Epitimo
Family : Cuscutaceae	Urdu : Akashbel	
Genus : <i>Cuscuta</i>	Bengali : Akashbel	
Species : <i>Reflexa</i> Roxb.		

Table 1: Botanical classification, Synonyms and Common names of *Cuscuta* species [8].

Worldwide distribution of *Cuscuta*

Cuscuta (dodder) is commonly found throughout the temperate and tropical regions of the world. This is rarely found in cool temperature; in this condition only four species occur in northern Europe. *Cuscuta approximata* commonly known as alfalfa dodder native to Eurasia and Africa and also found in North America. *C. australis* occurs in large parts of tropical and subtropical Africa as well as in Southern Europe, Asia and Australia. It also occurs in Madagascar. *C. campestris* (golden dodder, field dodder) occurs in America, East and Southern Africa, Northern Africa, Asia and the Indian Ocean islands. *C. reflexa* Roxb. (Giant dodder) originate from tropical and subtropical parts of Asia and America. *C. australis* occurs in wattle localities e.g. along water courses, and occurs from sea level up to 2000m altitude [10-12]. *C. campestris* has a wide tolerance of climatic conditions from warm temperature to sub-tropical and tropical conditions. *C. californica* native to Western North America and commonly called as California dodders. *C. gronovii* was found in United States, Canada, France, Germany, Luxembourg, Netherland and Italy. It grows on temperate forest habitats. *C. denticulata* commonly known as desert dodder or small toothed dodder, inhabitant to the deserts of the South-Western United States and Northern Mexico. *C. salina* (salt marsh dodder) occurs as resident plant of Western North America. *C. compacta* (compact dodder) grows on woody plants and distributed across the Eastern and Midwestern USA, Eastern Canada and Mexico. *C. japonica* commonly known as Japanese dodder found in Asia. *C. coryli* known as hazel dodder is a perennial plant and native to North America [13, 14].

Morphological characteristics

Dodder stem has a long, twinning, branched, glabrous, pale greenish yellow or dotted with red stem. The flower established as glabrous, curved, long ovate, oblong, obtuse, fleshy, pedicels short, 1.5 mm long, white (or) pink in color. Their seeds are large, black and glabrous having 2-4 seeds. Some specifications to be mentioned here, like which is the most common species of plant, it exhibits parasitic interaction [15].



Figure 1: Morphological appearance of *Cuscuta* sp.,

Phytochemical evaluation of *Cuscuta* sp

Based on the type of characteristic pattern, each type of *Cuscuta* sp., have different type of phytochemical activity. Diverse types of phytocomponents have been isolated based on the host and plant nutrition, which includes flavonoids [16], Dulcitol β , Mannitol, Sitosterol, Lycopene, Apigenin-7- β -rutinoside, 6-7 dimethoxy coumarin, quercetin, hyperoside [17,18], Propenamide, Reflexin, lutein, carotene, amarbellin, palmitic, oleic, stearic, linolenic acids, leuteolin, cuscutin, cuscotalin, kaempferol, kaempferol-3-Oglucoside, astragalin, benzopyrones, glucopyranosides, quercetin-3-O-glucoside, bergenin [19, 20]. One more important phytocomponent is LUPEOL extracted and isolated from *Cuscuta reflexa*, which exhibits pharmacological activities such as an antimicrobial, anti-inflammatory, antitumor, antiprotozoal and chemoprotective properties [21]. When act as an anti-inflammatory agents, lupenol leads to decrease interleukin-4 production by using T-helper type 2- cells. In preliminary level of screening, different type of phytoconstituents have been separated [22] includes flavonoids, glycoside, alkaloids, phytosterol, tri-terpenoids and steroids [23]. *Cuscuta reflexa* plant also contained polyphenols and flavonoids [24]. *Cuscuta reflexa* has a specific enzyme that is choline kinase which acts as mitochondrial enzyme [25]. Carboxymethylcellulose (CMC ase) also recover from *Cuscuta reflexa*. These enzymes have been free of β -glucosidase and other enzyme activities [26, 27]. The specific phytochemical, neoxanthin was identified which involved in light driven deep oxidation like a xanthophyll cycles and also helpful for protection against photodamage of the plant. Some other components were isolated from *Cuscuta* along with five known components, through the spectral analysis sequence determination method. And these components pose strong inhibitory activities against α -glucosidase. *C. reflexa* contain highly unusual carotenoid composition [28], it does not contain neoxanthin. Combined analysis of HPLC and MASS spectrometry are used to detect tissues of *C. reflexa* and find out two types of Xanthophyll; one is lutein 5-6 - epoxide and the other one is 9-cis-violaxanthina [29]. These components are used to for potential photo activity of reflexin [30] presented in Table 2.

Plant Name	Plant part	Solvent system used	Extraction method	Name of Separation technique	Phytochemicals	References
<i>C.reflexa</i>	Whole plant	MeOH	Maceration	CC	7'-(3',4'-dihydroxyphenyl)-N-[(4-methoxyphenyl)ethyl] propenamide 7'-(4'-hydroxy,3'-methoxyphenyl)-N-[(4-butylphenyl)ethyl] propenamide 6,7-dimethoxy-2H-1-benzopyran-2-one 2-(3-hydroxy-4-methoxyphenyl)-3,5-dihydroxy-7-O- β -D-glucopyranoside-4H-1-benzopyrane-4-one 3-(3,4-dihydroxyphenyl)-2-propen-1-ethanoate	[31]

					6,7,8-trimethoxy-2H-1-benzopyran-2-one 3-(4-O-β-D-glucopyranoside-3, dimethoxyphenyl)-2-propen-1-olKaempferol Quercetin Lupeol β-sitosterol Galic acid	[32]
		Aq.	Soxhlet	TLC	Quarcetin	[33]
		EtOH	-----	VLC	Odoroside H	[34]
		EtOH			21-hydroxyodoroside H	[34]
					Neritaloside	
					Stropeside	
					N-trans and cis feruloyl tyramines	
					Ethyl caffeate	
					Coumarins	
					Ursolic acid- sitosterol	
					Glucoside	
					4-O-p-coumaroyl- D-glucoside	
					Heneicosanoic acid	[35]
		n-hex	Soxhlet	GC-MS	Pentadecanoic acid	
					Hexadecanoic acid -330.50 gmol-1	
					Heptadecanoic acid	
					Octadecanoic acid -280.45 gmol-1	
					1,2,3 Propanetriol, 1-acetate,	
					Benzofiran 2,3,dihydro	
					Glycerol 1,2-diacetate	
					1 H- 1,2,4-triazol-5-amine 1-ethyl-	[36]
					2-methoxy-4-vinylphenol	
					Triacetin	
					D- glucitol, 4-O-hexyl	
					3,4,5-trimethoxy cinnamic acid	
					Hexadecanoic acid, ethyl ester	
					3,6- di methoxy phenanthrene	
					3,5- di- tert- Butyl-4-hydroxyanisol	
					Vanillin	
					3- aminopyrrolidine	
					Cetene	
					Sarcosine, N- isobutyryl, tetradecyl ester	
					4-((1E)-3-hydroxyl-1-propenyl)-2-methoxy phenol	
					1-octadecene	
					Scoparone	

		Pet Eth	Soxhlet	CC	Hexadecanoic acid, ethyl ester Isorhamnetin Isorhamnetin -3-O-glucoside Isorhamnetin -3-O-robinobioside 2-Methoxy-4-vinyl phenol Benzofuran-2,3-dihydro 3,5-di-tert-Butyl-4-hydroxyanisole	[37]
		MeOH	Maceration	GC-MS	Hexatriacontane - 506.98 gmol-1 n-Hexadecanoic acid - 330.50 gmol-1 Scoparone Hexadecanoic acid methy ester 1,3-Benzenediamine, N,N,N',N'-tetramethyl-Phenol, 4(3-hydroxyprop-1-en-1-yl), 2-methoxy Phenol, 2,4 bis (1,1dimethylethyl); 2,3,5,6-Tetramethyl para phenylene diamine Pregn-4-ene-18-oic acid Swarmalin Cis-swarmelin	[38]
	AP	MeOH	Maceration	Reverse phase HPLC	Coumarin 5,6,7-trimethoxycoumarin An antiviral protein with molecular weight about 14,000---18,000 Dalton	[39] [38]

Fil: filament; Aq: aqueous; MeOH: methanol; EtOH: ethanol; Pet. eth: petroleum ether; n-hex: n-hexan; EA: ethyl acetate; DMC: dichloro methane; CC: column chromatography; HPLC: high performance column chromatography; TLC: thin layer chromatography; VLC: vacuum liquid chromatography; GC-MS: gas chromatography-mass spectrometry.

Table 2: Phytochemical profile of various *Cuscuta* species.

Pharmacological activities of *Cuscuta* species

Plants therapeutic potential varied ranges from direct administration of the leaves, seeds, barks, roots and stems to the extracts and decoctions from different parts of the plants. Many *Cuscuta sp.*, being rich sources of diverse phytochemicals are popular components of various folk medicinal systems. *Cuscuta sp.*, are used in traditional medicine as a diaphoretic, diuretic, purgative, antihelminthic and tonic as well as a treatment for itching and bilious disorders. Seeds, stem and whole plant are utilized as prescription treat different types of ailments. *Cuscuta reflexa* is a treasured medicinal herb and widely used in conventional medicinal system of various Asian countries for treating multiple disorders. It is called a miracle therapeutic plant in the ethanobotany, and a wide array of chemical compounds has been isolated with diverse medicinal properties enlisted in table-3.

Species	Plant part	Preparation method	Traditional uses	References
<i>C.reflexa</i>	Whole plant	Paste	Treatment of swollen testicles, gout and joint pain, causes abortion, anti rheumatic, analgesic.	[40,41, 42,43, 44, 45, 46]
		Maceration	Infection treatment	[47]
		Infusion	Anti-poisonous	[48]
		Juice	Anti septic, useful in itching skin and jaundice	[49, 50]
		Powder	Anti-fertility agent, astringent, diaphoretic.	
	Stem	Pills	Anti tuberculosis.	[51]
		Decoction	Useful in skin disease, used for jaundice, cough, blood purification, bronchitis, fever, sex stimulation, anti diarrheal, anti inflammatory, anti ulcer, anti dandruff., fracture joining.	[52]
		Decoction	Hepatoprotective, anti diarrheal, useful in constipation, stomach disorders, urinary infections, jaundice, epilepsy, cholera, asthma	[50, 52, 54,55,56,57,58,59]
		Paste	Anti-hair fall, anti rheumatic, useful in skin diseases	[56, 15]
		Juice	Jaundice treatment	[13, 56]
	Seeds	Crushed	Blood purifier, purgative, good for brain, fever, anthrax in cattle	[60,56]
		Decoction	Effective in bilious disorders and fever	[61, 62]
		Poultice	Cause abortion	[40,63]
<i>C.chinensis</i>	Leaves	Extract	Carminative, antihelmintic, sedative, diuretic, useful in ulcer, liver disorders	[56]
		Juice	Pain reliever	[16,64]
	Fruits	Extract	Cold treatment	[65]
		Juice	Anti hypertensive, anti diarrheal, useful in jaundice	[66]
		Juice	Effective in scabies, eczema, inducing sterility	[67]
<i>C.japonica</i>	Whole plant	Juice	Antipyretic, cough reliever	[68,69]
		Juice		[70]
	Seeds	Juice	Anti ulcer, anti inflammatory, wound healer, jaundice treatment	[71]
		Dressing	Useful in painful inflammations	[72]
		Paste	Anti ulcer and wound healer	[72]
<i>C.reflexa</i>	Stem	Paste	Carminative, tonic, diuretic, sedative, diaphoretic	[73]
		Paste	Joining fractures	[72]
<i>C.japonica</i>	Leaves	Paste	Expectorant, carminative, tonic, anthelmintic, purgative, diaphoretic, anti-inflammatory, analgesic	[73]
		Paste	Antihypertensive	[74]

<i>C.australis</i>	Seeds	Decoction	Brain tonic Laxative, antihelminthic, astringent, emollient, sedative, sudorific, liver and kidney tonic, useful in sores and measles	[75]
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Table 3: Traditional therapeutic potential of *Cuscuta* species.

Hepatoprotective activity

Cuscuta epithymum is traditionally used as a liver tonic. Hepatoprotective activity with alcoholic extracts of stem of *Cuscuta reflexa* and aqueous extracts of stem of *Cuscuta reflexa* was studied against paracetamol induced hepatotoxicity in rats [76]. The histopathological changes (steatosis, necrosis etc.) were partly or fully prevented in animals treated with two extracts. Ethanolic extracts of *C.australis* also appeared as liver protector against acetaminophen intoxication in an animal model. The methanolic extracts of stem of *C.reflexa* were evaluated for hepatoprotective activity by observing its effects on carbon tetrachloride induced hepatotoxicity in liver histoarchitecture and alteration in certain biological parameters. Seeds of *C.chinensis* are commonly employed to nourish and improve hepatic disorders. Oxidative stress can stimulate the development of acetaminophen induced hepatotoxicity.

Anti-tumor activity

Some species of the genus *Cuscuta* afford alkaloids with indolic nuclei that are considered potential antitumor substance. *C.chinensis* is a popular antitumor prescription in the Unani medicine system. Oral administration of the plant extract at a dose of 1g/kg delayed the appearance and growth of skin papilloma and reduced the chances of carcinoma [77]. *Invivo* anticancer potential of *C.relexa* was determined by using murine model. Alcoholic extract and its chloroform fraction were found more potent. It showed highest toxicity against human breast cancer cell lines. The seed extract of *C.kotschyna* induced apoptosis in breast cancer cell line [78]. The major active phytoconstituents of *C.kotschyna* flavonols, quercetin has been found to reduce cell viability of quite a cost of cancer cell lines. Administration of chloroform and ethanol extracts of *C.relexa* showed antitumor activity against Ehrlich ascites carcinoma tumor in mice at doses of 200 and 400mg/kg body weight orally. It results in a significant decrease in tumor volume and viable cell count but increased non-viable cell count and mean several time, thereby increasing the life span of the tumor-bearing mice. *C.campestris* also has anticancer agents [79, 80].

Anti-oxidant activity

Plants can play a key role to fulfill prerequisite for exploration of biocompatible, effective and economic antioxidants. Many investigators have employed different qualitative and quantitative approaches to detect antioxidants in various *Cuscuta species*. Stem collected from different host and extracted with various solvents were analyzed for quantity of phenolics and flavonoids content. Their antioxidant capacity was measured by using a variety of assays including reducing power, DPPH scavenging activity, percent inhibition of linoleic acid peroxidation. *Invitro* antioxidant activity of *C.reflexa* stems were investigated by estimating the degree of non-enzymatic hemoglobin glycosylation. Ethyl acetate and ethanol extract showed higher activity than other fractions, and very close and identical in the magnitude and comparable to the standard antioxidant agents [81].

Anti-bacterial activity

Crude ethanolic extract of *C.reflexa* showed antimicrobial activity against *E.coli* and *S.sonnei*. *C.relexa* collected from different sear-obial activity against *Staphylococcus aureus*, *Staphylococcus epidermidis*, *E.coli*, *Micrococcus luteus*, *Pseudomonas aeruginosa* [82]. The antimicrobial efficacy was concentratedely dependent against all the tested strains. The methanol extract of *C.reflexa* exhibited antibacterial and free radical scavenging activities. *Xanthomonas campestris* (XC) is a widely spread infectious agent causing a huge loss in food crops with viable symptoms and leave shedding. Aqueous decoction and infusion extract of *C.pedicellata* were evaluated for

antibacterial activity against diverse pathogens of XC using invitro well diffusion method. The Methanolic extract also showed promising high antimicrobial activity.

Antipileptic & anticonvulsive activity

C.reflexa showed significant reduction in the duration of convulsion in tonic seizure induced by pentylenetetrazole in mice. It also reduces the tonic extension convulsion induced by maximum electroshock-induced convulsions [83]. The ethanolic extract has anticonvulsant property and may probably affect both the Gamma Amino Butyric Acid (GABA) aminergic- and glycine-inhibitory mechanism. The main active chemical constituent is flavonoid which is responsible for depressant activity. The processed extracts showed a significant anticonvulsive property by altering the levels of catecholamine and brain amino acids in mice.

Hypoglycemic activity

The hypoglycemic activity of methanol and chloroform extracts of whole plants of *C.reflexa*, investigated in oral glucose tolerance tests in Long Evans rats. Both methanol and chloroform extracts of *C.reflexa* whole plant demonstrated significant oral hypoglycemic activity in glucose-loaded rats. It was further reported that these two compounds act at multiple targets to ameliorate hyperglycemia [84].

Anti-HIV activity

The crude water extracts of *C.reflexa* exhibited anti-HIV activity which could be due to combinatory effects with compounds of different modes of action. The methanol extract of *C.reflexa* exhibited anti-bacterial and free radical scavenging activity [85]. The methanolic extract of *C.campestris* showed weak anti-HIV activity.

Effect on blood pressure

Alcoholic extract of *C.reflexa* have positive inotropic and cardiotonic activities on the perfused frog heart. In a series of experiments on dog blood pressure, it caused a fall in blood pressure [23]. Ethanolic extract of the stem of *C.reflexa* caused a dose dependent decrease in arterial blood pressure and heart rate in pentothal-anaesthetized rats. Hypotensive and brady cardiac effects of *C.reflexa* were found to be independent of cholinergic receptor stimulation or adrenergic blockade [86].

Relaxant & spasmolytic action

Aqueous and alcoholic extracts of *C.reflexa* stems showed relaxant and spasmolytic action on small intestine of guinea pig and rabbit. Extracts also exhibited acetyl choline like action [87].

Cholinergic action

The effects of the stem extract of *C.reflexa* resembled acetylcholine when tested on isolated rabbit ileum and frog rectus abdominals and heart. These effects were blocked by atropine. Effect of the extract on isolated frog rectus abdominal muscle was blocked by pancuronium and potentiated by neostigmine [88].

Diuretic activity

Aqueous and alcoholic extracts of *C.reflexa* showed diuretic activity in Wistar rat [89]. *C.epithymum* has mild diuretic activity and it is used to treat sciatica and scurvy. *C.europaea* seeds are used to treat psoriasis and it also has a diuretic activity. *C.reflexa* seeds are carminative [89]. *C.racemosa* has small diuretic activity and used for treatment of wounds [90].

Anti-diabetic activity

The methanol and aqueous extracts showed significant reduction in blood glucose during OGTT in diabetes rats. The treatment also resulted in an improvement in body weights, decreased Hb1c and restored lipid profile. Methanolic extracts of *C.reflexa* has significant

antidiabetic effects and improves metabolic alterations. Antidiabetic activity of *C.chinensis* was evaluated in dexamethasone-induced insulin-resistant human liver carcinoma (HepG2) cells. *C.chinensis* polysaccharides can reduce blood sugar level in Type-2 Diabetes. The efficacy was tested on alloxan-induced diabetes in a mice model [91].

Hair growth activity

Plants as hair growth promoters have found their use in almost all traditional medicinal systems. The petroleum ether and ethanolic extract of *C.reflexa* were given in male swiss albino rats. *C.reflexa* extract is useful in the treatment of alopecia. This study was shown to be capable of promoting follicular proliferation or preventing hair loss in cyclophosphamide-induced hair fall [92].

Anti-inflammatory effect

In different phases of pathogenesis of cancer, inflammatory reactions play a decisive role. Invitro and invivo tests, aqueous and alcoholic extracts of stem of *C.reflexa* and its ethyl acetate fraction showed remarkable anti-inflammatory activity. *C.reflexa* significantly suppressed inflammation by reducing edema volume up to 80 percent in rats. *C.campestris* markedly inhibited carrageenan-induced edema in rats. *C.chinensis*, showed the potential for treatment of brain inflammation by suppressing the inflammatory responses [93, 94].

Immunological activity

Ethanolic extract of *C.chinensis* showed considerable adjuvant potentials towards cellular and humoral immune responses in mice models and can be used as vaccine adjuvant. Extract enhanced specific antibodies (IgG, IgG1 and IgG2b) to a noticeably high level by affecting Th1 and Th2 cell functions. Kaempferol was identified as the main flavonoid of methanol fraction. Based on several research findings, Kaempferol has potential to treat chronic inflammatory and autoimmune diseases [95]. *C.australis* may act as an immunopotentiator for mammals by increasing the percentage of Phagocytosis.

Anti-obesity activity

C.pedicellata is widely used for management of obesity. Ethanolic extract of *C.pedicellata* has significantly reduced the body weight along with serum lipid profile in high-fat diet fed rats [96]. Recently, the results proved that polyphenols are reported to possess anti-obesity activity.

Effect on melanin production

C.chinensis can promote melanogenesis of amelanotic melanocytes and improved the tyrosinase activities. Furthermore, it significantly enhanced skin melanin and tyrosinase production. It has been reported that in invitro and invivo, the seed aqueous and ethanolic extracts of *C.chinensis* have melanogenesis effect. Consumption of *C.chinensis* extract with milk reduced the melatonin synthesis and thus ameliorated the elimination of melisma [97].

Effect on the reproductive system

C.reflexa has an anti-fertility effect. Methanolic extract arrested the normal estrus cycle and decreased ovarian and uterus weight in adult female mice. *C.reflexa* is rich in flavonoids, as the result reported that flavonoids act as an antifertility agents *C.chinensis* extract, and its isolations can improve reproductive systems of both males and females. Ethanolic extract of *C.chinensis* may improve erectile dysfunction conditions. An herbal formula, KH-204 containing *C.chinensis*, ameliorates erectile dysfunction by its antioxidant and lipid profile improving property [98].

Anti-aging activity

In Chinese herbal medicinal system, *C.chinensis* is an important antichanging prescription. Polysaccharides of *C.chinensis* can exhibit

anti-aging effects by scavenging free radicals and opposing lipid peroxidation. Ethanolic extract of *C.chinensis* significantly supposed the non-enzymatic glycosylation of D-galactose induced rat aging model [99]. Various experimental researches showed that it can regulate immune responses, prolong cell cycle, positively affect body metabolism, improve physiology of internal body organs, and stress management, which proves its anti-aging effects.

Anti-hypertensive & anti-osteoporotic activity

In Pentothal anesthetized rats, ethanolic extract of *C.reflexa* decreased arterial blood pressure and heart beat rate. Four caffeoylquinic acid derivatives were isolated from the active fraction having inhibitory effects on Angiotensin Converting Enzyme (ACE) activity. Presence of these metabolites at least in part is responsible for the anti-hypertensive activity extract. *C.chinensis* effectively boasted tissue regeneration of damaged bones by promoting the formation of osteoblasts from their precursor cells. Five flavonoids were isolated from which Kaempferol and hyperoxide were found osteogenic in nature [100].

Reno protective effects

In Wister rats, aqueous and alcoholic extract of *C.reflexa* exhibited substantial diuretic activity. *C.chinensis* has been used as a kidney tonic since ancient time. Research suggests that *C.chinensis* extract ameliorates renal functions and regulates urine concentration [101].

Conclusion

Cuscuta genus has enormous therapeutic potential since ancient civilizations. The phytoconstituents such as flavonoids, alkaloids, lignans, polysaccharides, steroids, volatile oils and resins are encountered as important bioactive ingredients of the plant. This plant considered as a miracle genus having broad spectrum of pharmacological activities. Doctations, extracts, paste, powder, juice and infusions from various parts of plant impart therapeutic nature against numerous ailments of human beings. Unraveled explorations of these plant medicinal properties, only few species are identified. With the advent of modern scientific technologies, yet to be found out, the other functions of rest of the species. Limitations of this study might include that, only fewer animal studies are employed for the detection of its efficacy of their pharmacological activities. Hence, this reviews, pave a path for the exploration and novel therapeutic measure for the various ailments of human welfare with safe margin.

References

1. Machado MA and Zetsche K. "A structural, functional and molecular analysis of plastids of the holoparasites *cuscuta reflexa* and *cuscuta europaea*". *Planta* 181.1 (1990): 91-96.
2. Kapoor, et al. "Host range of *Cuscuta reflexa* Roxb. In Jammu province of Jammu and Kashmir State, India", *Indian J. Weed Sci* 40.1 (2008): 98-100.
3. Dawson JH., et al. "Biology and control of *Cuscuta*". *Rev Weed Sci* 6 (1994): 265-317.
4. Davidson-Hunt I. "Ecological ethnobotany: stumbling toward new practices and paradigms". *MASA J* 16 (2000): 1-13.
5. Tripathi I. Raj Nighantu. Varanasi: Chaukhambha Krishna Academy (2006): 38-39.
6. Pandey GS. Bhavprakash Nighantu (*Indian Materia Medica*). Varanasi: Chaukhambha Bharti Academy (2004): 447-448.
7. Bapalal G. Nighantu Adarsh. Varanasi: Chaukhambha Bharti Academy 2 (2005): 97-99.
8. Nandkarni KM. *Indian Materia Medica Bombay: Popular Prakashan Pvt Ltd* 1 (1976): 419-420.
9. Vijikumar S. "*Cuscuta reflexa* Roxb. A wonderful Miracle Plant in Ethnomedicine". *Indian J of Natural Sci* 11 (2011): 677.
10. Stefanovic S and Olmstead RG. "Testing the phylogenetic position of a parasitic plant (*Cuscuta*, *Convolvulaceae*, *Asteridae*): Bayesian inference and the parametric bootstrap on data drawn from three genomes". *Syst Bio* 53.3 (2004): 384-399.
11. "Plants Profile for *Cuscuta gronovii* gronovii (scaldweed)". *Plants. usda.gov* (2017).
12. Costea M., et al. "Untangling the systematics of salt marsh dodders: *Cuscuta pacifica*, a new segregate species from *Cuscuta salina* (*Convolvulaceae*)". *Systematic Botany* 34.4 (2009): 787-795.

13. "Cuscuta compacta". Natural Resources Conservation Service PLANTS Database. USDA (2016).
14. "Cuscuta japonica (Japanese dodder)". www.cabi.org (2020).
15. "Plants Profile for Cuscuta coryli (hazel dodder)". plants.usda.gov (2018).
16. Subramanian SS Nair AGR. Chemical components of Cuscuta reflexa (Roxb.) Indian J Chem 1 (1963): 501.
17. Ramachandran Nair AG and Thirupurasundari G. "Coumarins and flavonoids from Cuscuta reflexa parasitic on Bougainvillea spectabilis". Fitoterapia 63.4 (1992): 381-382.
18. Dandapani M and Nagarajan S. Isorhamnetin 3-O-neohesperidoside from Cuscuta reflexa. Indian J Chem 28 (1989): 606-607.
19. Kelker SL, et al. Isolation of compound from Cuscuta reflexa. Indian J Chem Sect 23 (1984): 458-4
20. Anis E et al. Phytochemical studies on Cuscuta reflexa [1999]. J Nat Prod 5 (1999): 124-126.
21. Gallo MBC, et al. "Biological activities of lupeol". Int J Biomed Pharm Sci 3.1 (2009): 46-66.
22. Sharma S, et al. "Antimicrobial Study of Cuscuta reflexa Collected In Different Seasons". Int J Pharm Bio Sci 4.3 (2013): 1393-1397.
23. Singh GS and Garg KN. "Some pharmacological studies on Cuscuta reflexa plant (Akash bel)". Indian journal of Pharmacol 5.2 (1973): 344-345.
24. Siwakoti M and Siwakoti S. Ethanobotanical studies of Satars of Jhapa district, Nepal: A case study of Haldibari VDC. A report submitted to University Grants Commission, Kathmandu, Nepal (1996).
25. Setty PN and Krishnan PS. "Choline kinase in Cuscuta reflexa". Biochem J 126.2 (1972): 313-324.
26. Chatterjee U, et al. "Physico-chemical and functional characterization of a high molecular weight carboxy methyl cellulase from Cuscuta reflexa". Indian journal of Biochem Biophys 34.4 (1997): 354-64.
27. Bungard RA, et al. "Unusual carotenoid composition and a new type of xanthophyll cycle in plants". Proc Natl Acad Sci 96.3 (1999): 1135-1139.
28. Anis E, et al. "Alpha-glucosidase inhibitory constituents from Cuscuta reflexa". Chem Pharm Bull 50.1 (2002): 112-114.
29. Snyder AM, et al. "Carotenoid specificity of light-harvesting complex II binding sites occurrence of 9-cis-violaxanthin in the neoxanthin-binding site in the parasitic angiosperm Cuscuta reflexa". J Biol Chem 279.7 (2004): 5162-8.
30. Tripathi VJ, et al. "A new flavanone, reflexin, from Cuscuta reflexa and its selective sensing of nitric oxide". Appl Biochem Biotechnol 127.1 (2005): 63-7.
31. Anis E, et al. α -glucosidase inhibitory constituents from Cuscuta reflexa. Chem Pharm Bull 50.1 (2002): 112-114.
32. Shailajan S and Joshi H. "Optimized separation and quantification of pharmacologically active markers quercetin, kaempferol, β -sitosterol and lupeol from Cuscuta reflexa Roxb". J Pharm Res 4 (2011): 1851-1853.
33. Teware K. "Pytochemical extraction and TLC estimation of extract of Cuscuta reflexa". World J Pharm Pharm Sci 5 (2016): 378-384.
34. Versiani MA, et al. "Cytotoxic cardiac glycoside from the parasitic plant Cuscuta reflexa". Chem Nat Compd 53 (2017): 915-922.
35. Jahan IA, et al. "Elemental and fatty acid content of four medicinal plants: Kaiempferia rotunda, Cuscuta reflexa, Centella asiatica and Asparagus racemosus". European J Med Plants (2015): 1.
36. Bais N and Kakkar A. "Comparative phytochemical analysis of Cuscuta reflexa parasite grown on Cassia fistula and Ficus benghalensis by GC-MS". Int J Pharm Pharm Sci 5 (2013): 350-355.
37. Dokuparthi SK, et al. "Phytochemical investigation and evaluation of antimutagenic activity of the extract of Cuscuta reflexa Roxb by Ames Test". Int J Pharm Sci Res 5 (2014): 3430-3434.
38. Rath D, et al. "Identification of bioactive constituents from different fractions of stems of Cuscuta reflexa Roxb". Using GC-MS. Nat Prod Res 32 (2017): 1977-1981.
39. Uddin SJ, et al. "Swarnalin and cis-swarnalin, two new tetrahydrofuran derivatives with free radical scavenging activity, from the aerial parts of Cuscuta reflexa". Nat Prod Res 21 (2007): 663-668.
40. Patel JN and Patel NK. "Study of parasite hosts of the genus Cuscuta and its traditional uses in Planpur Taluka, Gujarat, India". Ethnobot Leaf 14 (2010): 126-131.
41. Basak S, et al. "Role of some ethno medicines used by the Santal tribal people, of the district Bankura, WB, India, for abortifacient

- purposes". J Med Plants Stud 4 (2016): 125-129.
42. Singh RS, Shahi SK. "Diversity of medicinal plants of Ratanpur region of Bilaspur district (Chhattisgarh)". J Med Plants 5 (2017): 276-281.
 43. Mohapatra SS., et al. "Ethnomedicinal plants used in balasore district of Odisha: a comprehensive report". Int J Cur Microbiol App Sci 7 (2018): 1959-1963.
 44. Rai Y, Kumar D. "Survey on medicinal climbers in meerut district, Uttar Pradesh, India". Imperial J Interdisciplinary Res 2 (2016): 603-610.
 45. Nita RD and Haresh DL. "Ethno-botanical survey of some medicinal plants in jatasankar region of girnar forest, Gujarat, india". Glob J Res Med Plants India Med 2 (2013): 830-841.
 46. Paudel N., et al. "Some medicinal plants use in ethnical group from batnagar, eastern, Nepal". Am Sci Res J Eng Tech Sci 41 (2018): 233-239.
 47. Rahmatullah M., et al. "Survey and scientific evaluation of medicinal plants used by the Pahan and Teli tribal communities of Natore district, Bangladesh". Afr J Tradit Complementary Altern Med 9 (2012): 366-373.
 48. Singh EA., et al. "Medicinal plants used by the Thakar tribes of Raigad district, Maharashtra for the treatment of snake-bite and scorpion-bite". Int J Phytother Res 2 (2012): 26-35.
 49. Singh S. "Ethnobotanical study of some climbers of Parsa district forest of Nepal". J Med Plants 4 (2016): 6-10.
 50. Ahirwar RK. "Diversity of ethnomedicinal plants in Boridand forest of district Korea, Chhattisgarh, India". Am J Plant Sci 6 (2015): 413-425.
 51. Khattak NS., et al. "Ethno veterinary uses of medicinal plants of district Karak, Pakistan". J ethnopharmacol 171 (2015): 273-279.
 52. Akter MH., et al. "Synergistic antihyperglycemic activity of *Coccinia grandis* leaves and *Cuscuta reflexa* stems". J Pharm Pharm Sci 5 (2016): 236-243.
 53. Yaseen G., et al. "Ethnobotany of medicinal plants for livelihood and community health in deserts of Sindh-Pakistan". In Plant and Human Health, Springer, Cham 1 (2018): 767-792.
 54. Singh S and Sharma A. "Studies on ethnomedicinal Plant of Baghicha Jashpur Chattisgarh". J Sci Lett 2 (2017): 48-55.
 55. Chowdhury M. "Folk medicines used by the Rabha tribe in Coochbehar district of West Bengal: a preliminary report". Adv Ethnobot (2007): 289-296.
 56. Patel H and Patel N. "Sacred and medicinal plant diversity of patan sacres grove of Patan District (NG)". Life Sci Leaf 92 (2017): 50-60.
 57. Seliya AR and Patel NK. "Ethnomedicinal uses of climbers from Saraswati river region of Patan district, North Gujarat". Ethnobot leaf 13 (2009): 865-872.
 58. Kala CP. "Ethnomedicinal botany of the Apatani in the Eastern Himalayan region of India". J Ethnobiol Ethnomed 1 (2005): 1-8.
 59. Khatun MM and Rahma M. "Medicinal plants used by the village Pania under Baghmara District, Bangladesh". Discovery 54 (2018): 60-71.
 60. Azam MN., et al. "Ethnomedicines used by the Oraon and Gor tribes of Sylhet district, Bangladesh". Am-Eurasian J Sustain Agric 7 (2013): 391- 402.
 61. Khalid M., et al. "Characterization of ethno-medicinal plant resources of karamar valley Swabi, Pakistan". J Radiat Res Appl Sci 10 (2017): 152-163.
 62. Azam MN., et al. "Medicinal plants used by the traditional medical practitioners of Barendra and Shamatat (Rajshahi & Khulna Division) region in Bangladesh for treatment of cardiovascular disorders". J Med Plants. 2 (2014): 9-14.
 63. Dutta ML. "Plants used as ethnomedicine by the Thengal Kacharies of Assam, India". Asian J Plant Sci Res 7 (2017): 7-8.
 64. Kirtikar KR and Basum BD. Indian medicinal plants. Delhi: Periodical Experts Book Agency 1 (1984).
 65. Verma N and Yadav RK. *Cuscuta reflexa*: a parasitic medicinal plant. Plant Arch 18 (2018): 1938-1942.
 66. Kumar S., et al. "Ethnobotanical study of some common plants from district hamirpur of Himachal Pradesh (India)". Int J Adv Res 3 (2015): 492-496.

67. Shipa A., et al. "Phytotherapeutic practices of a folk medicinal practitioner in Kishoreganj district, Bangladesh". J Med Plants 6 (2018): 240-242.
68. Begum HA., et al. "Phytochemical evaluation of ethanobotanically selected medicinal plants of mardan, Pakistan". J Adv Bot Zool 3 (2015): 1-5.
69. Khan W., et al. "Ethno-ecology, Human Health and Plants of the Thandiani Sub Forest Division, Abbottabad, KP, Pakistan". In Plant and Human Health, Springer, Cham 1 (2018): 547-567.
70. Patel S., et al. "Dixit VK. An updated review on the parasitic herb of *Cuscuta reflexa* Roxb". Jo Chin Integr Med. 10 (2012): 249-255.
71. Qureshi R and Bhatti GR. "Ethnobotany of plants used by the thari people of nara desert, Pakistan". Fitoterapia. 79 (2008): 468-473.
72. Qureshi R., et al. "Ethnomedicinal uses of herbs from northern part of Nara desert, Pakistan". Pak J Bot 42 (2010): 839-851.
73. Fahmy GM. Qatar biodiversity newsletter. Ostrich 2 (2008): 1-5.
74. Talha J., et al. Hypertension and herbal plants. Int Res J Pharm 2 (2011): 26-30.
75. Chen GT., et al. "Medicinal uses, pharmacology, and phytochemistry of convolvulaceae plants with central nervous system efficacies: a systematic review". Phytother Res 32 (2018): 823-864.
76. Versiani MA. "Studies in the chemical constituents of *Bombax ceiba* and *Cuscuta reflexa*". Karachi: University of Karachi/H.E.J Research Institute of Chemistry (2004).
77. Bais N and Kakkar A. "Phytochemical Analysis of Methanolic Extract of *Cuscuta reflexa* Grown on *Cassia fistula* and *Ficus benghalensis*". Int J of Pharm Sci 25.2 (2014): 33-36.
78. Jha U and Shelke T. "Hepatoprotective activity of hydroalcoholic extracts of *Cuscuta reflexa* roxb in paracetamol intoxicated albino rats". IJRAP 2.4 (2011): 1290-1293.
79. Nisa M., et al. "Effect of *Cuscuta chinensis* water extract on 7, 12-dimethylbenz [a] anthracene-induced skin papillomas and carcinomas in mice". J Ethnopharmacol 18 (1986): 21-31.
80. Sepehr MF., et al. "The *Cuscuta kotschyana* effects on breast cancer cells line MCF7". J Med Plants Res 5 (2011): 6344-6351.
81. Selvi EK., et al. "Phytochemical profiling and evaluation of the hepatoprotective effect of *Cuscuta campestris* by high-performance liquid chromatography with diode array detection". Anal Lett 51 (2018): 1464-1478.
82. Riaz M., et al. "Natural products from *Cuscuta reflexa* Roxb. With antiproliferation activities in HCT 116 colorectal cell lines". Nat. Prod. Res 31.5 (2017): 583-587.
83. Solat P., et al. "Antimicrobial, Antioxidant And Minerals Evaluation of *Cuscuta europea* and *Cuscuta reflexa* Collected From Different Hosts And Exploring Their Role As Functional Attribute". Int. Res J Pharm. App Sci., 3.5 (2013): 43-49.
84. Sharma S., et al. "Antimicrobial Study of *Cuscuta reflexa* Collected In Different Seasons". Int J Pharm Bio Sci 4.3 (2013): 1393-1397.
85. Borole SP., et al. "Evaluation of anti-epileptic activity of *Cuscuta reflexa* Roxb". Res J Pharm Biol Chem Sci 2.1 (2011): 657-663.
86. Fadia H Al-Sultany., et al. "Studying Hypoglycaemic Activity of *Cuscuta chinensis* Lam. on Type 1 Diabetes Mellitus in White Male Rats". IOP Conf. Series: Journal of Physics: Conf. Series 1294 (2019): 062020.
87. Mahmood N., et al. Constituents of *Cuscuta reflexa* are anti-HIV agents. Antivir Chem Chemother 8.1 (1997): 70-74.
88. Singh GS and Garg KN. "Some pharmacological studies on *Cuscuta reflexa* plant (Akash bel)". Indian journal of Pharmacol 5.2 (1973): 344-345.
89. Gilani AUH and Aftab K. Pharmacological actions of *Cuscuta reflexa*. Int J Pharmacogn 30.4 (1992): 296-302.
90. Prasad DN. "Preliminary pharmacological investigation on *Cuscuta reflexa* Roxb". Indian J Med Res 53 (1965): 465-470.
91. Kayath HP and Goel NK. "Effects of *Cuscuta* stem extract on various animal tissues". Indian J Pharmacol 27.4 (1995): 227-229.
92. Sharma S., et al. "Comparative study of *Cuscuta reflexa* and *Cassytha filiformis* for diuretic activity". Phamacognosy Res 1.5 (2009): 327-330.
93. Sandeep S and Abilasha Mittal. "Antidiabetic activity of *C. reflexa*. International Journal of Pharma and Chemical Research 3.3 (2017).

94. Ghayoumi A and Mashayekhi A. "Scleroderma treatment in Iranian traditional medicine: a case report". *Adv Herb Med* 2 (2016): 1-4.
95. Lin MK, et al. "Kaempferol from Semen cuscuteae attenuates the immune function of dendritic cells". *Immunobiology* 216.10 (2011): 1103-9.
96. Zekry SH, et al. "Effect of metabolites isolated from *Cuscuta pedicellata* on high fat diet-fed rats". *Med Chem Res* 24 (2015): 1964-1973.
97. Mojtabae M, et al. "The effect of the traditional medicine product" Milk-Cuscuta" on skin hyper pigmentation in patients with melasma". *Middle East J Family Med* 7 (2018): 204-211.
98. Ferraz HO, et al. "Antiulcer and antioxidant activities and acute toxicity of extracts of *Cuscuta racemosa* Mart (Convolvulaceae)". *Lat Am Jo Pharm* 30 (2011): 1090-1097.
99. Li CS, et al. "Advances and challenges in screening traditional Chinese anti-aging materia medica". *Chin J Integr Med* 19.4 (2013): 243-52.
100. Yang L, et al. "Antiosteoporotic compounds from seeds of *Cuscuta chinensis*". *J Ethnopharmacol* 135.2 (2011): 553-60.
101. Patel S, et al. "A study on the extracts of *Cuscuta reflexa* Roxb. In treatment of cyclophosphamide induced alopecia". *DARU J of Pharm Sciences* 22.1 (2014): 7.

Volume 1 Issue 5 December 2021

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