

1 **CENTELLA ASIATICA (URB.) L.: THE HERBAL ELIXIR OF EASTERN MEDICINE**

2 **Abstract**

3 *Centella asiatica* (Urb.) L., commonly known as gotukola or mandookparni is a valuable herb of importance known
4 and mentioned in Ayurveda as well as traditional Chinese medicine literature. It belongs to the family Apiaceae with
5 widespread distribution around the world. Tribals or forest dwellers collect it from the wild and sell in the market as
6 vegetable and medicinal herb that bring them a regular income with no investment. But as herb reproduces by
7 vegetative reproduction primarily it could be a threat in the form of overexploitation in absence of awareness among
8 collectors who are usually from rural background. There is a high demand of this herb in national as well as
9 international markets due to its ethnobotanical uses as well as for its products used for pharmaceutical as well as
10 food industry purposes besides other commercial uses of wider applications. Active biochemicals present in this
11 herb are triterpenoids as Asiaticoside, Madecassoside, Asiatic acid, flavanoids, alkaloids and essential oils etc that
12 provide it dermatological, neuroprotective, antioxidant, anti-inflammatory, anti-cancer, antimicrobial and
13 hepatoprotective etc. properties. Many advancements are going to develop techniques for enhancement and
14 extraction of these secondary metabolites *in vitro* culture with the help of various biotechnological tools that are
15 required for conservation of this valuable herb.

16 **Systematic position**

17 Kingdom: Plantae
18 Phylum: Tracheophyta
19 Class: Magnoliopsida
20 Order: Apiales
21 Family: Apiaceae
22 Genus: *Centella* L.
23 Species: *Centella asiatica* (L.) Urb
24



25

26 **Fig. 1 Morphological features of *C. asiatica***

27 Keywords: Asiaticoside, Gotukola, Madecassoside, Mandookaparni, Medhya Rasayana, Neuroprotective, Wound
28 healing

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33 **Common/Vernacular Name:**

34 Gotu kola (Hindi), Asiatic pennywort, Indian pennywort (English),
35 *Chokiora* (Bihar), *Thankuni* (Bengali), *Barmi* (Gujarati), *Mandookaparni* (Hindi), *Supriya*, *Mutthil* (Sanskrit), *B*
36 *rahmi* (Urdu), *Babassa*, *Saraswataku* (Telugu), *Batsyiar/Khliengsyiar* (Khasi, Meghalaya), *Peruk* (Manipuri), *La*
37 *mbak* (Mizoram) and *Manimum* (Assam)

38

39 **Geographic Distribution and Habitat:**

40 This herb is widespread across Eastern Asia, including India, China, Japan, and Australia. *Centella asiatica*
41 thrives in a wide range of ecological environments but is especially common in moist, shaded, and fertile
42 regions such as stream banks and damp pathways in temperate areas across India.

43

44 **Morphological features**

45 A small slender creeping herb. Stems are with stolons, green to reddish green in colour. Leaves are long-stalked,
46 reniform with rounded apex. Flowers are pinkish to red in colour, borne in small, rounded bunches (umbels).
47 Each flower is partly enclosed in two green bracts. It is a creeper that roots at the nodes.

48

49 **Propagation Methods:**

50 The plant primarily reproduces vegetatively via stem cuttings, with sexual reproduction being less frequent.
51 Recent advances in tissue culture have enabled more efficient propagation, germplasm preservation, and cost-
52 effective cultivation (Sharma & Vimala, 2010).

53 **Soil and Climate Requirements:**

54 It can grow in sandy, loamy, or clay soils and tolerates acidic, neutral, or alkaline conditions. The plant is
55 adaptable to partial or full sunlight and prefers moist or wet soil for optimal growth.

56 **Nutritional Profile**

57 Proximate analysis of nutritional analysis by Ogunka-Nnoka *et al.*, 2020 and Chandrika *et al.*, 2015 for leaves of
58 *C asiatica* showed it contained moisture ($13.10 \pm 1.07\%$), ash ($16.5 \pm 0.45\%$), protein ($8.35 \pm 1.28\%$), lipid (1.20
59 $\pm 0.10\%$), fiber ($17.00 \pm 1.87\%$), and carbohydrate ($43.81 \pm 0.70\%$). Fatty acid profiling indicated a
60 predominance of palmitic acid (55.70%) among the saturated fatty acids and linoleic acid (17.50%) among the
61 unsaturated fatty acids. Amino acid analysis revealed high levels of glutamate (13.389 g/100 g) as the major
62 nonessential amino acid and histidine (11.64 g/100 g) as the major essential amino acid. Phytochemical
63 screening identified several bioactive compounds, including proanthocyanin (11.964 $\mu\text{g/g}$), rutin (11.8883 $\mu\text{g/g}$),
64 naringenin (3.0122 $\mu\text{g/g}$), quinine (10.4490 $\mu\text{g/g}$), flav-3-ol (2.5900 $\mu\text{g/g}$), sparteine (3.0122 $\mu\text{g/g}$), phenol
65 (18.8713 $\mu\text{g/g}$), flavonones (2.1836 $\mu\text{g/g}$), steroids (18.8974 $\mu\text{g/g}$), kaempferol (0.7273 $\mu\text{g/g}$), phytate (1.6851
66 $\mu\text{g/g}$), naringenin (2.7523 $\mu\text{g/g}$), resveratrol (10.8596 $\mu\text{g/g}$), tannin (4.4377 $\mu\text{g/g}$), and ribalinidine (3.0500
67 $\mu\text{g/g}$).

68 The abundance of these nutrients and bioactive phytochemicals in *Centella asiatica* leaves highlights their
69 potential applications in the pharmaceutical and food industries.

70 **Edible and Culinary Applications**

71 **Edibility:**

72 The leaves are consumed raw or cooked and are considered a leafy vegetable in countries like India, China, Sri
73 Lanka, and Indonesia. Due to its strong aromatic flavor, it is typically used in moderation when added to salads.

74 **Traditional**

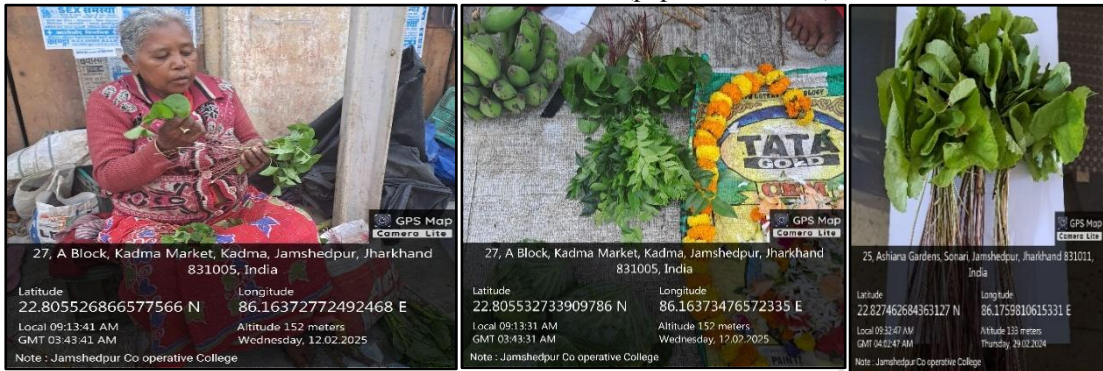
Culinary

Uses:

75 The plant, commonly referred to as **Beng Saag**, is not consumed raw due to its bitterness. Instead, it is
76 thoroughly washed, boiled or sautéed to reduce bitterness, cooked with digestive and aromatic spices

77 **Popular Dishes Include:**

- 78 • Beng Saag Bhaji – Stir-fried with garlic, onions, mustard oil, and chilies
- 79 • Mixed Saag Curry – Cooked with other leafy greens
- 80 • Beng Saag Dal – Lentil-based curry
- 81 • Side Dish for Pakhala – Served with fermented rice water (popular in Odisha)



82 **Fig. 2 Tribals usually women collect gotukola from wild and selling in local haats**

84 Gotu Kola/Thankuni plays a significant role in the economy of tribal forest dwellers, particularly in regions like
 85 India (Odisha, Tripura, Jharkhand) and Madagascar, by serving as both a cash-generating Non-Timber Forest
 86 Product (NTFP) and a source of nutrition. It is collected manually from the wild mostly by rural women and
 87 sold in local haats so playing crucial socio- economy roleguaranteeing future income. Due to its traditional
 88 medicinal usage in Ayurvedic and Chinese medicine it is in high demand in both local and international markets
 89 so that makes it a valuable herb with almost zero investment and contributes to the poverty alleviation of rural
 90 and forest-dwelling populations.

91 **Significance in Ayurveda:**

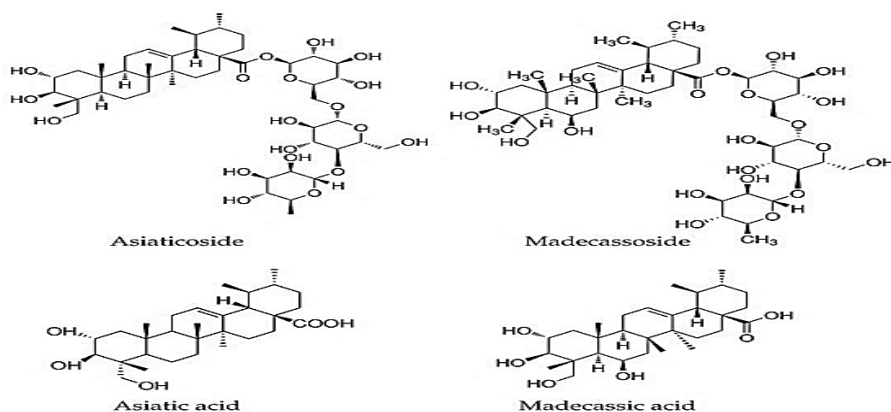
92 *Centella asiatica*, known in Ayurveda as *Mandukparni* or *Brahmi*, and referred to internationally as *Gotu Kola*,
 93 is a highly valued medicinal plant in both traditional and modern healthcare systems (Chopra *et al.*, 1969,
 94 Chopra and Nayar 1956. It holds an important position in the Indian Pharmacopoeia and is recognized for its
 95 therapeutic benefits particularly in enhancing memory and healing wounds. Its medicinal properties are also
 96 documented in international pharmacopoeias, including the European Pharmacopoeia, German Commission E,
 97 and the World Health Organization (Howes & Houghton, 2003).

98 **Phytochemical Composition**

99 Phytochemicals of *Centella asiatica* can be extracted by conventional or novel extraction techniques.

- 100 1. Conventional extraction methods like Soxhlet and cold methods can be used for extraction purposes
 101 but due to prolonged extraction time, less efficiency and requirement of excess amounts of solvent and
 102 compound degradation these are not considered as efficient ones in comparison to novel
 103 techniques. (Savic *et al.*, 2023 and Belwalet *et al.*, 2020).
- 104 2. Novel extraction methods like supercritical fluid extraction ultrasonic extraction, radio frequency and
 105 enzyme-assisted extraction techniques microwave, advance metabolomics techniques, ultra-
 106 performance liquid chromatography (UPLC), NMR, and MS, e.g., liquid chromatography-MS (LC-
 107 MS/MS) and gas chromatography-MS (GC-MS) provides efficient yields with lesser extraction
 108 time (Songvutet *et al.*, 2021, Yuan *et al.*, 2023, Zakaria *et al.*, 2023 and Ondekoet *et al.*, 2020).
- 109 3. Similarly cold plasma as a water pretreatment also extract efficiently by retaining the herb's bioactive
 110 properties with lesser requirement of energy and extraction time (Tripathy *et al.*, 2023).
- 111 4. Likewise, usage of green solvents like triethylammonium sulfate effectively extract asiaticoside
 112 from *Centella asiatica* in comparison to other solvents (Othman *et al.*, 2023).

113 Presence of broad spectrum of **bioactive compounds** by scientific investigations done by (Tan *et al.*,
 114 2021 Schaneberget *et al.*, 2003 Srivastava *et al.*, 1997) validate the claims of traditional Indian medicine
 115 regarding the plant's medicinal potential.



116

117 **Figure 3** Chemical structures of triterpenoid compounds isolated from *Gotukola* (Heinet *et al.*, 2025)

118

Table 1 Metabolite spectrum of *c asiatica* and its therapeutic properties

S No	Phytochemical	Biological activity	Application	Reference no
I.	Triterpenoids <i>Signature</i> <i>Phytochemicals</i>	Research suggests that formulations containing a full spectrum of triterpenoids are more effective than single isolated compounds, indicating a synergistic influence on inflammatory responses and oxidative stress pathways (Shukla <i>et al.</i> , 1999).		
a)	Asiaticoside	promotes collagen production by increasing the expression of transforming growth factor beta-1 (TGF- β 1) and fibronectin. It also supports angiogenesis, facilitating the development of new blood vessels, which plays a crucial role in tissue regeneration.	These properties make it beneficial in the management of skin injuries such as wounds, burns, and scar formation.	(Hashim <i>et al.</i> , 2011, Lokanathan <i>et al.</i> , 2016 Rashid <i>et al.</i> , 2023)
		supports in promoting neurogenesis via by increasing the expression of brain-derived neurotrophic factor (BDNF), brain-derived neurotrophic factor	possible therapeutic value for neurodegenerative disorders.	He <i>et al.</i> , 2023
b)	Madecassoside	Madecassoside plays an important role in suppressing the activation of nuclear factor kappa B (NF- κ B), thereby decreasing inflammation triggered by pro-inflammatory cytokines	It also provides protection against ultraviolet (UV)-induced skin aging and contributes to the wound healing process.	(Won <i>et al.</i> , 2010, Jung <i>et al.</i> , 2013)
		Its anti-inflammatory effects, combined with its capacity to enhance collagen synthesis, make it valuable for skin repair.	Furthermore, incorporating madecassoside into nanoparticle-based delivery systems has been shown to	Lu, <i>et al.</i> , 2023s

			improve its stability and enhance transdermal penetration for dermatological use.	
c)	Asiatic acid (C₃₀H₄₈O₅) and Madecassic acid (C₃₀H₄₈O₆)	demonstrate a wide range of biological activities, including anti-inflammatory, anticancer, hepatoprotective, and neuroprotective effects. These actions are mediated through the regulation of key signaling pathways such as mitogen-activated protein kinases (MAPK), nuclear factor erythroid 2-related factor 2 (Nrf2), and phosphatidylinositol 3-kinase/protein kinase B (PI3K/Akt) signaling pathways	Derivatives of Asiatic acid have attracted attention as potential frameworks for anticancer drug design because of their cytotoxic effects on glioblastoma and melanoma cells. Similarly, Madecassic acid helps in wound healing due to its anti-inflammatory and antioxidant properties.	Tan <i>et al.</i> , 2021
d)	Brahmoside	potential anxiolytic and adaptogenic effects		Ganie <i>et al.</i> , 2022
II.	Flavonoids and other Polyphenols	exhibit antioxidant, anti-inflammatory, neuroprotective, and vasoprotective properties. These are also reported to reduce β -amyloid plaque formation and improve synaptic plasticity	indicating potential benefits in addressing neurodegenerative disorders, cognitive impairment, and overall brain health.	Gray <i>et al.</i> , 2018
a)	Quercetin	Quercetin is known for its strong antioxidant capacity, neutralizing reactive oxygen species (ROS) and by regulating important inflammatory mediators such as cyclooxygenase-2 (COX-2) and tumor necrosis factor-alpha (TNF- α).		Sul, 2021 Domitrović 2012
		It has also shown neuroprotective effects in Alzheimer's disease	highlighting its potential significance in the prevention and management of cognitive decline and age-associated neurological disorders.	Khan <i>et al.</i> , 2020
b)	Kaempferol	demonstrates strong		Chatterjee <i>et</i>

		anti-inflammatory activity, largely through suppression of the NF- κ B signaling pathway and stimulation of Nrf2-driven antioxidant defense mechanisms.		<i>al.</i> , 2024
c)	Rutin	enhancing capillary strength and stability, while also contributing to antioxidant and protection against UV-induced damage.	These properties make it especially beneficial in skin care and vascular health applications.	Michalak <i>et al.</i> , 2022, Peres <i>et al.</i> , 2016
d)	Apigenin, Luteolin, and Hesperidin	further expand the therapeutic profile of the herb, providing additional benefits such as anti-aging effects, anxiety-reducing potential, and antimicrobial activity.		Ullah <i>et al.</i> , 2020
II.	Alkaloids			
a)	Hydrocotyline and Thankunside	associated with neuroprotective and cognitive-enhancing properties. These compounds may also function as cholinesterase inhibitors	potentially supporting memory and overall cognitive performance.	
III.	Essential Oils and Volatile Compounds (0.1% to 0.2% w/w)	produce gentle calming, stress-reducing, and skin-soothing effects, consistent with the traditional use of <i>C. asiatica</i> in holistic healing systems. In addition to improving the sensory (organoleptic) qualities of formulations containing <i>C. asiatica</i> , these volatile compounds may act synergistically to enhance the overall therapeutic effect	Although their pharmacological mechanisms have not been studied as extensively as the plant's other metabolites the possible anxiolytic and mood-balancing properties of these volatile components are supported by observations from aromatherapy practices and traditional medicinal preparations.	Kunjumon <i>et al.</i> , 2022 Murray <i>et al.</i> , 2013
IV.	Monoterpenes and Sesquiterpenes	(Caryophyllene, α-Humulene, and β-Pinene)	antimicrobial, anxiolytic, and dermal applications	Joshi <i>et al.</i> , 2007
V.	Phytosterols	[Phytosterols as β-sitosterol, campesterol, and stigmasterol are predominantly concentrated in the leaves than other parts ³⁵ .		
		These plant sterols are known to influence lipid metabolism by competing with dietary	Through this competitive interaction, they help reduce	Miszczuk <i>et al.</i> , 2024

		cholesterol for absorption within the gastrointestinal tract.	circulating cholesterol levels, thereby supporting cardiovascular health	
		possess anti-inflammatory and antioxidant activities	beneficial in the management of metabolic syndrome and other inflammation-related metabolic disorders.	Feng <i>et al.</i> , 2018
		Emerging evidence also suggests that these compounds can modulate immune responses by affecting immune cell activity and cytokine production; however, further mechanistic research is needed, particularly in relation to formulations derived from <i>C. asiatica</i>	Overall, phytosterols may contribute to broader systemic benefits, including positive effects on cardiovascular, metabolic, and immune function, especially in products designed to address dyslipidemia and chronic inflammatory states.	Shen, <i>et al.</i> , 2024 Hu, <i>et al.</i> , 2017
VI.	Polyacetylenes	mainly concentrated in the roots, with moderate amounts also found in the leaves.		
a)	Cadiyenol and Cadiyenone,	can promote fibroblast migration, stimulate angiogenesis, and enhance skin regeneration properties,	consistent with applications in wound and skin care preparations	Scherer <i>et al.</i> , 2009 Li <i>et al.</i> , 2022
		These constituents may enhance the wound-healing process by influencing extracellular matrix remodeling and regulating cellular signaling pathways involved in tissue repair.		
		act as internal signaling molecules, particularly under environmental stress conditions.	This adaptive function may also have pharmacological significance in humans, as their role in stress-response modulation could contribute to the plant's defense-related and adaptogenic properties	Divekar <i>et al.</i> , 2022 Minto <i>et al.</i> , 2008
VII.	Tannins, Sugars, Resins, Inorganic acids, and amino acids such as Aspartic acid, Glycine, Glutamic acid, α-Alanine, and Phenylalanine.			

VIII.	Rich in vitamins B, C, and G , the total ash includes chlorides, sulfates, phosphates, iron, calcium, magnesium, sodium, and potassium.
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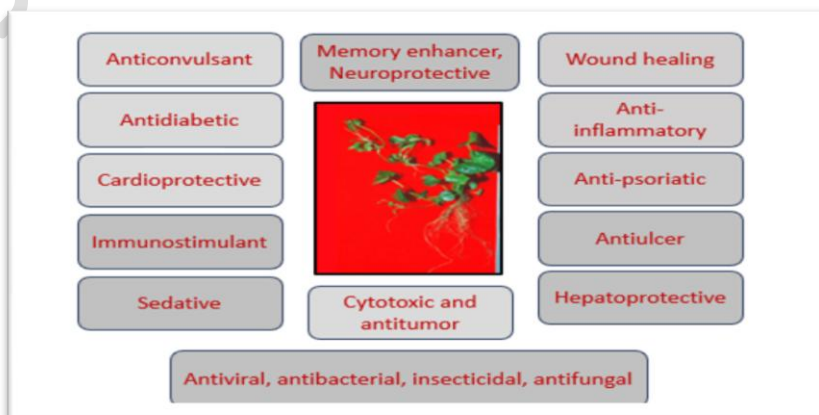
120 **Traditional and Modern Medicinal Uses**

121 *Centella asiatica* has a long-standing presence in the traditional medicine systems across various continents^{46, 47}.
 122 It has been used by multiple indigenous communities for centuries to treat a wide range of health issues,
 123 particularly related to skin, inflammation, and cognition. Its cross-cultural therapeutic use is a testament to its
 124 versatility and effectiveness.

125 In Ayurvedic medicine, *Centella asiatica* is considered a *Rasayanaa* category of rejuvenating herbs.
 126 Traditionally, it is believed to improve cognitive function, enhance skin texture, and support wound healing. Its
 127 extracts are used in modern Indian pharmacology for treating skin disorders such as **leprosy, eczema, and**
 128 **psoriasis**(Feng *et al.*, 2021 Gupta *et al.*, 2000) and for anti-acne application (Kuo *et al.*, 2021).

129 The plant has demonstrated:

- 130 • **Neuroprotective and cognitive-enhancing properties:**
 131 *Centella asiatica* has been reported to exert multiple beneficial effects on the nervous system, including
 132 neuroprotective activity, improvement of memory and cognitive function, as well as antidepressant and
 133 anxiolytic effects(Hengjumrut *et al.*, 2018; Khemawoot *et al.*, 2018). Clinical evidence suggests that CA
 134 supplementation reduced stress levels, alleviated symptoms of anxiety and depression, and improved
 135 attention and adaptive functioning in individuals diagnosed with generalized anxiety disorder, without
 136 producing significant adverse effects(Tan *et al.*, 2021).
- 137 • **Therapeutic applications in vascular and inflammatory disorders:**
 138 CA has been traditionally and clinically utilized in the management of anxiety, inflammatory
 139 conditions, venous insufficiency, and certain types of cancer(Brinkhaus *et al.*, 2000). It has shown
 140 relevance in treating vascular-related disorders such as venous hypertensive microangiopathy, diabetic
 141 microangiopathy, and chronic venous insufficiency. Additionally, clinical investigations have explored
 142 its potential benefits in other conditions, including leprosy, chronic liver diseases, and gastric ulcers
 143 (Tan *et al.*, 2021).
- 144 • Used in **wound care, antiviral, antibacterial, diuretic, antipyretic, and rheumatic** treatments
- 145 • *Centella asiatica* is also actively explored in **clinical research** due to its broad therapeutic potential as
 146 it is widely used for its antimicrobial, antioxidant, neuroprotective and wound healing properties
 147 (Prakash *et al.*, 2017 Tan *et al.*, 2021 Hafiz, *et al.*, 2020).
- 148 • It is used for the treatment of a variety of dermatological conditions as lupus, eczema, psoriasis, leprosy
 149 and varicose ulcers. Its extracts exhibited positive influence on wound healing(Ahmed *et al.*, 2019 and
 150 Arribas-López *et al.*, 2022) by improving collagen synthesis and microcirculatory function (Brinkhaus
 151 *et al.*, 2000 Cesarone *et al.*, 2001 Incandela *et al.*, 2001.)
- 152 • The effects of a standardized extracts of *Centella asiatica*were studied by Damkerngsuntorn, *et al.*,
 153 2020 on postlaser resurfacing wound healing on the face: A Split-Face, Double-Blind, Randomized,
 154 Placebo-Controlled Trial and concluded this treatment regimen might be an option for post-laser
 155 resurfacing.



156

157




Fig. 4 Therapeutic Properties of *C. asiatica*(Brinkhaus *et al.*, 2000, Orhan *et al.*, 2012)

158 **Commercial Applications**


159 Due to the presence of its secondary metabolites *C asiatica* is mostly used in

- 160 • Cosmetics and Skincare (Cosmeceuticals):
- 161 • Pharmaceuticals & Medicine:
- 162 ○ Wound Healing: Used in topical creams and dressings for surgical lesions, bedsores, and skin
163 ulcers as Madecassol® and Centelase®
- 164 ○ Therapeutic Use: Used to treat inflammatory diseases, chronic hepatic disorders, and for
165 promoting cognitive function.
- 166 • Functional Foods and Nutraceuticals:
- 167 ○ Beverages/Food: Used in teas, juices, and herbal products to promote overall health.
- 168 ○ Health Supplements: Capsules, tablets, and powders for enhancing cognitive function and
169 circulation.
- 170 • In the food industry, *C. asiatica* extract represents a promising natural alternative for enhancing
171 product preservation. Extracts of *Centella asiatica* demonstrate strong free-radical scavenging capacity
172 along with notable antimicrobial effects, indicating their suitability as functional additives in food
173 packaging materials. The bioactive compounds present in the plant, particularly terpenoids and
174 phenolic constituents, contribute to its effectiveness against food-borne pathogens by damaging
175 microbial cell membranes and suppressing a broad range of microorganisms. Consequently,
176 incorporating *C. asiatica* extracts into packaging films may provide an innovative approach for **active**
177 **food packaging applications**. Its antioxidant and antibacterial properties can support food quality
178 maintenance, extend shelf life, and reduce spoilage, ultimately helping to minimize food waste (Pillai *et*
179 *al.*, 2024).

180 **Table 2 Some commercial products of *C asiatica***

1.	Centellaeasiaticae herba	commercial formulations such as skin creams, wound healing agents, and moisturizers (Zainol <i>et al.</i> , 2008).	global marketplace, particularly in Germany
2.	SBL Hydrocotyle Asiatica Dilution 30 CH	It helps with skin-related comfort, may support overall vitality and may assist during phases of general weakness sensations as well.	
3.	Himalaya Gotu Kola / <i>Centella asiatica</i> (Capsules)	Improving Memory and Concentration, helps to maintain an excellent cognitive performance It helps to Improve memory and concentration in stressful periods.	
4.	Himalaya Organic Gotu Kola	Nootropic Supplement with Adaptogens for Energy, Focus, Memory, and Calming Relaxation - Vegan, Non-GMO, Gluten Free	
5.	<i>Centella asiatica</i> Extract	The product addresses broad spectrum dermatological concerns.	Alchem International Pvt. Ltd.
6.	<i>Centella asiatica</i> Powder Extract EMPEROR	Strong anti-inflammatory, help comfort skin, Boosts Blood Circulation, Regulates Blood Sugar Level, Stress Buster, Wound Healing. Relieves	

	HERBS®	Joint Pain. Skin Treatment	
7.	Merlion Naturals Gotu kola	Help with memory and mood relaxant	
8.	Dr Willmar Schwabe Germany Hydrocotyle Asiatica (<i>Centella asiatica</i>)	anti-ageing properties. It has a mild laxative effect. It is used to relieve constipation. It works as a diuretic	
9.	Dr. Reckeweg Hydrocotyle Asiatica	Helps manage leprosy, gonorrhoea and leucorrhoea. Helps deal with skin related issues. Assists in improving memory	
10.	<i>Centella asiatica</i> Gotu Kola Cream, Madecassol	Topical cream with 1% titrated <i>Centella asiatica</i> extracts for skin repair. Supports soothing and healing of dry, irritated, or damaged skin	
11.	Natier Centella Asiática y Jengibre (Natural Asian Centella juice with ginger)	Dietary Supplement Nutritional Benefits	
12.	Organic Gotu Kola Powder- <i>Centella asiatica</i> Superfood Powder by IyasaHolistics	for Focus, Calm & Skin Support – Raw Ayurvedic Herb for Drinks & Smoothies – Gotu Kola Supplement	
13.	Gotu kola, Herbal Tea	Memory Boost, Brain Nourishment, Cognitive Function	
14.	One Herb - Gotu Kola Tea Pure <i>Centella asiatica</i> Leaves Single Ingredient Ayurvedic Tea Loose Leaf Infusion	BRAIN TONIC: Improves memory retention and overall cognitive function. Supports mental clarity & stress relief. anti-ageing properties. Supports deep & restful sleep: Herb for longevity healthy hair and skin	
15.	<i>Centella asiatica</i> Serum	Mary & May is a Korean brand of natural cosmetics. It is a serum to instantly soothe irritated, sensitive skin without leaving a sticky feeling	

16.	Ecoberry <i>Centella asiatica</i> Soothing Gel Cream	Safe and effective agent for Anti-ageing, soothing, moisturizing and healing	
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181 Dose dependent cytotoxicity

182 Natural medicinal products derived from the food chain have gained increasing attention for their potential role
 183 in managing chronic diseases. To ensure their safe application, comprehensive toxicity evaluations are essential.
 184 International regulatory authorities emphasize this requirement in their safety guidelines for botanical products,
 185 dietary supplements, and herbal medicines (Speijers *et al.*, 2010).

186 The phytochemical composition of plant extracts, particularly terpenoid content in leaves, can vary depending
 187 on several factors, including plant maturity and nutrient availability (Mangas *et al.*, 2008; Sharma *et al.*, 2014;
 188 Müller *et al.*, 2013). Such variability highlights the importance of standardized extracts when assessing
 189 biological activity and safety.

190 Standardized extracts of *Centella asiatica* (L.) Urban (INDCA) have shown potential in addressing stress-related
 191 behavioural disorders. Because therapeutic management of stress-associated conditions typically requires
 192 prolonged administration over days or weeks to achieve clinical benefits, evaluating long-term safety becomes
 193 crucial. Preclinical toxicological assessments conducted according to established regulatory protocols are
 194 therefore necessary. Previous investigations have reported that leaves extracts were well tolerated in acute oral
 195 toxicity, sub-chronic toxicity, and mutagenicity studies in rats (Deshpande *et al.*, 2015).

196 In a more recent clinical context, Wright *et al.*, (2023) examined a standardized *Centella asiatica* water extract
 197 product was examined. The analytical method used for its evaluation met the United States Food and Drug
 198 Administration criteria for linearity, precision, accuracy, and recovery in plasma and urine samples. The extract
 199 was further assessed in a Phase I randomized, double-blind, crossover study involving eight healthy older adults
 200 who received single doses of 2 g or 4 g. Both doses were reported to be safe and well tolerated, supporting
 201 further clinical development of this formulation.

202 Additionally, under OECD GLP 408 guidelines for a 90-day repeated-dose oral toxicity study (Junsai *et al.*, 2024)
 203 in 100 Wistar rats, Centell-S demonstrated a no-observed-adverse-effect level (NOAEL) of 800 mg/kg/day.
 204 These findings provide supportive evidence for the continued development of Centell-S as a
 205 phytopharmaceutical candidate for clinical use.

206 Biotechnological Advances

207 This herb is in high demand in national as well as international market as it is part of Ayurveda and traditional
 208 Chinese medicine system, there is a concern regarding overexploitation of this herb. IUCN has listed it in least
 209 concern species but wild scale uncontrolled harvesting due to lack of awareness of good agricultural and
 210 collection practices cannot be denied at local rural or in forests areas (Singh *et al.*, 2010). As a prevention
 211 measure many *in vivo* and *in vitro* studies have been done on *Centella asiatica* in past years for sustainable
 212 cultivation, micropropagation and *in vitro* production of medicinal principles of this plant (Sharma *et al.*, 2014,
 213 Sharma and Vimala 2010, Sharma and Vimala 2015, Tiwari *et al.*, 2000, Singh *et al.*, 2023). The biosynthesis of
 214 triterpenoids in *Centella asiatica* occurs mainly via the mevalonate pathway, a fundamental plant metabolic
 215 pathway producing isoprenoid building blocks by key enzymes such as β -amyrin synthase that forms the core
 216 triterpenoid skeleton, which is subsequently modified by cytochrome P450 monooxygenases to generate diverse
 217 bioactive compounds. Recent transcriptomic and metabolomic investigations have identified specific gene
 218 clusters that regulate the formation of major constituents, including Asiaticoside and Madecassoside (Wan *et al.*,
 219 2024). These findings have opened new opportunities for metabolic engineering approaches enhancing both the
 220 production levels and consistency of valuable metabolites.

221 Further *in vitro* culture systems such as cell suspension cultures and hairy root cultures are developed to enable
 222 controlled and large-scale production of triterpenoids (Baek *et al.*, 2022). These systems offer advantages
 223 including improved yield stability, standardized quality, and minimized environmental variability. Moreover,
 224 advanced molecular techniques, including CRISPR-Cas9-based genome editing and transgenic strategies, are
 225 being used to upregulate essential biosynthetic genes and redirect metabolic flux towards production of required
 226 secondary metabolites (Arribas-López *et al.*, 2021).

227 In addition, the neuroprotective potential of Asiaticoside, Madecassoside, and Asiatic acid present in *C.*
 228 *asiatica* were examined as natural inhibitors of acetylcholinesterase (AChE), an enzyme closely associated with

229 neurodegenerative conditions like Alzheimer's disease. *in silico* study incorporated multiple ligand mapping,
230 molecular docking, molecular dynamics (MD) simulations, and MM-PBSA binding free energy analyses. The
231 docking results demonstrated strong binding affinities of Asiaticoside and madecassoside toward the AChE
232 active site, surpassing the reference drug donepezil (−133.51 kJ/mol). Molecular dynamics simulations further
233 supported the stability of these ligand–enzyme complexes, reinforcing their potential as effective AChE
234 inhibitors Khairinisa *et al.*, (2025).

235 Overall, these scientific and technological advances highlight the considerable promise of *Centella asiatica* as a
236 sustainable source of bioactive compounds and support ongoing efforts to develop novel therapeutic agents for
237 cognitive impairment and other neurodegenerative disorders.

238 Conclusion

239 *Centella asiatica*, known traditionally as Mandukparni or Brahmi and globally as Gotu Kola, is a plant of
240 immense medicinal, nutritional, and cultural significance. Its widespread use in both traditional systems like
241 Ayurveda and in modern pharmacology highlights its therapeutic versatility. The plant is rich in bioactive
242 compounds such as triterpenoids, flavonoids, alkaloids, glycosides, and essential fatty acids, which contribute to
243 its wide range of pharmacological activities including neuroprotection, wound healing, anti-inflammatory,
244 antibacterial, and cognitive enhancement effects.

245 With applications spanning from herbal medicines and skincare products to edible leafy greens in various
246 cuisines, *Centella asiatica* holds a unique place in both health and nutrition sectors. Advances in propagation
247 methods, including tissue culture, further enable sustainable cultivation and conservation of this valuable plant
248 species.

249 As global interest in natural and plant-based remedies continues to grow, *Centella asiatica* remains a key
250 botanical resource for developing safe, effective, and affordable therapeutic agents across traditional and
251 modern healthcare systems.

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