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ABSTRACT

Gymnema sylvestre is a plant that has been used in traditional medicine for thousands of years. The plant's medicinal value include bitter, acrid, thermogenic, digestive, liver-tonic, anodyne, and anti-inflammatory properties. The phenolic compounds of this plant are known as stigmasterol and triterpenoid saponins. These compounds have a wide range of pharmacological significance as a naturopathic medication since ancient times. In this review, we will focus on the pharmacodynamics of this medicinal plant, as well as its therapeutic potential for diabetes, obesity, cardiovascular disorders, and dietary supplements in the food industry.

KEYWORDS: Gymnema sylvestrae, Anti-oxidant, Gymnemic acids, Anti-Diabetic.

INTRODUCTION

Plants are a key source of our contemporary medicine and a major trumpcard for drug discovery research. Only 5 to 15% of plans have been researched for their potential as medicines, despite the fact that about 25% of contemporary medications are derived from plant sources.^[1] By using chromatographic methods, it was discovered that Gymnema sylvestre stems contain a number of chemical substances that have medicinal value, including stigmasterol and triterpenoid saponin. Stigmasterol which is responsible for hypoglycemic, anticancer activity and anti-tumor activity is due to triterpenoids.^[2-5] The leaves of the plant contain saponins of the oleanane and dammarane types^[6] Additionally, it also includes cardiac glycosides, anthraquinones, etc^[7] The ethanolic extract of plant demonstrated considerable 1,1-diphenyl-2-picrylhydrazyl radical scavenger action and superior antioxidant activity.^[8] Methanolic extract of plant showed anti-inflammatory effects in rats when carrageenan-induced inflammation was given to Wistar rats^[9] This plant was proved to have high antioxidative potential and hydroxyl free radical scavenging activity against DPPH, with hydroxyl free radical inhibition at a level of 59%^[10] It has a radical-scavenging action against ferric superoxide and hydrogen peroxide.^[11] Clinical studies have demonstrated that this plant can reduce blood sugar, triglycerides, total cholesterol and thereby reduces body weight, in addition to raising the blood's availability of insulin and C-peptide.^[12] In the Ayurvedic medical system and Indian Pharmacopoeia, Gymnema sylvestre is listed as one of the main botanicals used to

treat diabetes.^[13] Although this plant has various beneficial characteristics that demonstrate its value, there isn't much research, particularly clinical trials on it.

BOTANICAL DESCRIPTION AND TAXONOMY OF Gymnema sylvestre

Gymnema sylvestre is a delicate species with a sluggish rate of growth. In the arid woods of central and Southern India and other parts of Asia, it emerges as a densely branched, woody plant that may climb to the top of trees.^[14-15]

Taxonomical Classification



Fig. 1: Gymnema sylvestrae.

This shrub is pubescent in nature and has just sprung stems and branches.^[16] It has a tap root system.^[17] Cylindrical, branching, rigid, twining, terete internodes, 0.7–17.2 cm long, and 2–10 mm in diameter are the characteristics of the stems ^[17-18]. The leaves are 2.5–6 cm long, often oval or elliptic, and feature a distichous phyllotactic opposite arrangement pattern ^[16]. The petioles of the leaves are 1-2 cm long, smooth above, with a rounded base and thickly velvety pubescent beneath. The leaves are acute or briefly acuminate.^[18-19]

TRADITIONAL USES

Gymnema sylvestre is described as the treatment for urinary disorders and glycosuria in the ancient medical text of Shushruta.^[20] According to the traditional medicine, Ayurveda, and homoeopathic schools of medicine, it is a medicinal plant with a variety of possible uses.^[16] The Ayurvedic medical system has traditionally utilised it to treat ailments brought on by phlegm and piles, as well as diabetes, malaria, and snake bites^{[13][19]}. Asthma, bronchitis, amenorrhea, leukoderma, haemorrhoids, jaundice, renal and vesicle calculi, cardiopathy and dyspepsia are among conditions for which the herb is given in Ayurveda.^[20-22] Additionally, many plant components, including the roots, stem, and leaves, have been utilised in traditional medical systems as cardiotonic, digestive, diuretic, laxative, stimulant, stomachic, and uterine tonics.^[22]

PHYTOCHEMISTRY OF Gymnema sylvestre

Utilising chromatographic methods, it was discovered that *Gymnema sylvestre* stems contain a number of chemical substances that have medicinal value, including stigmasterol and triterpenoid saponin. The therapeutic potential of stigmasterol compounds is extensive and includes anti-diabetic, hypoglycemia, antioxidant, and anti-cancer properties. In several investigations, triterpenoid saponins also demonstrated anti-tumor, antifungal, hepatoprotective, and antidiabetic potential.^[2-5] Gymnemic acids and gymnema saponins, which are categorised as oleanane saponins, are the main chemical

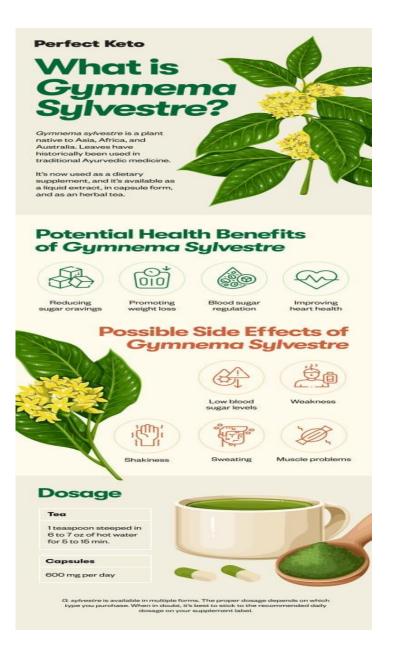
Kingdom	Plantae
Sub-kingdom	Tracheobionta
Super division	Spermatophyta
Division	Magnoliophyta
Class	Magnoliopsida
Subclass	Asteridae
Order	Gentianales
Family	Asclepiadaceae
Genus	Gymnema R. Br.
Species	Sylvestre

Table 1: Taxonomy.

components of this plant. *Gymnema sylvestre* leaves contain saponins of the oleanane and dammarane types^[6] Additionally, this plant's leaves include cardiac glycosides, anthraquinones, saponins, tannin, quinones, flavonoids, and phenols.^[7,23]

ANTIDIABETIC

One of the most significant medicinal herbs for the treatment of diabetes in India is Gymnema sylvestre. An active component obtained from this plant that has antidiabetic properties is gymnemic acid. Numerous studies have been conducted worldwide on this plant's antidiabetic properties. On diabetic rabbits treated with alloxan, Shanmugasundaram et al. (1983) investigated the anti-diabetic efficacy of Gymnema.^[4-5] They demonstrated how Gymnema dried leaf powder controls blood sugar levels and preserved its homeostasis. Additionally, it controlled the amounts of sorbitol dehydrogenase, phosphorylase, and gluconeogenic enzymes. Data indicated that using Gymnema leaf extracts may help type 2 diabetes individuals renew or repair their pancreatic beta cells. After the subjects serum samples were taken, higher levels of insulin were detected.



ANTIOXIDANT

Compared to A. bilimbi and C. frutescens, this plant's ethanol extract demonstrated considerable 1,1-diphenyl-2-picrylhydrazyl (DPPH) radical scavenging action and superior antioxidant capacity.^[8] In a study, *Gymnema sylvestre's* antioxidant efficacy against DPPH was also noted. This plant was shown to be more effective in scavenging DPPH radicals, while in another, it was discovered to be able to lower LDL oxidation.^[24-25] This plant was shown to have high antioxidative potential and hydroxyl free radical scavenging activity against DPPH, with hydroxyl free radical inhibition at a level of 59.8% and DPPH inhibition at 87.3%, respectively.^[10]

Additionally, it was discovered to have considerable radical scavenging action against ferric super oxide and hydrogen peroxide, as well as antioxidant activity in a variety of circumstances, including those involving high fat diets, hydrogen peroxide, nitric oxide, and superoxide radicals induced oxidative stress in rats.^[26-28]

ANTI INFLAMMATORY

When carrageenan-induced inflammation was administered to Wistar rats, *Gymnema sylvestre's* methanoic extract demonstrated anti-inflammatory action in the rats^[9] This plant's methanolic extract considerably (p 0.05) decreased the oedema in the rat paws caused by carrageenan. In a different investigation, this plant's aqueous extract showed inhibitory efficacy against carrageenan-induced rat paw oedema and mouse peritoneal ascites.^[29] A 50% inhibitory dosage of 50–555 nmol/ear of this plant's ethanolic extract was also discovered to have inhibitory effects against TPA-induced inflammation.^[30]

Since time immemorial, the leaves of *Gsylvestre* are said to be anodyne, bitter, acrid, thermogenic, digestive, livertonic, and anti-inflammatory.^[31] in nature. The anti-inflammatory activities of *G. sylvestre* are due to the

plant's bioactive tannins and saponins.^[32] In the study on rats with granuloma and paw oedema brought on by cotton pellets, the aqueous extract of *G sylvestre* leaf was examined for its anti-inflammatory activity at doses of 200, 300, and 500 mg/kg using the drug phenylbutazone as the reference. At a dose of 300 mg/kg, *Gymnema* aqueous extract was shown to significantly reduce paw oedema volume by 48.5% after 4 hours, whereas phenylbutazone significantly reduced paw oedema volume by 57.6%. Furthermore, in comparison to the control group, the aqueous extract at concentrations of 200 and 300 mg/kg showed a reduction in granuloma.^[33]

LIPID LOWERING ACTIVITY

It was discovered that *Gymnema sylvestre* leaf extract has extremely powerful hypolipidaemic action. The leaf extract was given to Wister female rats as part of a study. A high-fat diet was used to cause hyperlipidemia in these rats. This extract was found to drastically reduce cholesterol, Low-Density Lipoprotein (LDL) and triglyceride levels while efficiently raising High-Density Lipoprotein (HDL) levels.^[34] Gymnema sylvestre hydroalcoholic leaf extract also shown to have lipid-lowering capability. In this experiment, rats were administered high cholesterol for seven days, and it was shown that these rats had greater levels of cholesterol, triglycerides, and LDL while having lower levels of HDL. Gymnema sylvestre extract was administered to these rats after seven days, and it was found to reduce the elevated levels of cholesterol, triglycerides, and LDL while increasing HDL levels.^[35-36].Due to the presence of acidic components such flavonoids, saponins, tannins, etc., it has been hypothesised that this plant has the ability to decrease cholesterol levels It was claimed in a number of additional studies to lower triglycerides, cholesterol, Very Low-Density Lipoprotein (VLDL), and Low-Density Lipoprotein (LDL) in diabetic rats.^[36-37]

DENTAL CARIES

Dental caries are infections of the teeth caused by a variety of gram-positive cariogenic bacteria, such as *S. aureus, S. mitis and S. mutans* as well as the fungus-like *Candida albicans*, which attaches to the tooth surface by releasing extracellular polysaccharides from sucrose and metabolises sugar into organic acids, primarily lactic acid, leading to demineralization of the tooth enamel.^{[37-}

^{38]} The *Gymnema sylvestre* leaf extracts in chloroform, petroleum ether and methanolic form were tested against microbial dental infections at concentrations of 25, 50 and 100 mg/mL and were found to be significantly effective against these carcinogenic bacteria, with the methanolic extract possesing the highest activity at the lowest concentration. Due to the plant's hydroalcoholic extract's high potential, "Gurmar Herbal tooth paste" is developed and manufactured.^[39]

ETHNOBOTANICAL USES

Gurmar is referred to as a destroyer of madhumeha (glycosuria) and other urinary ailments in the ancient Indian medical text Sushruta. It contains antiinflammatory, anodyne, digestive, emetic, diuretic, stomachic, stimulant, anthelmintic, laxative, cardiotonic, expectorant, antipyretic, and uterine tonic are all claimed properties of G. sylvestre extract.^[43] Jaundice, constipation, cardiopathy, asthma, bronchitis. amenorrhea, conjunctivitis, renal and vesical calculi, dyspepsia leukoderma and Parkinsonism could be treated with this herb^[44]. The root bark works as an expectorant, emetic, and analgesic for bodily aches, while the root liquid is effective for treating snakebite. In addition it treats respiratory and cardiac conditions, piles and colic discomfort.^[45]



CLINICAL STUDY REPORTS

In addition to numerous studies on animal models, various extracts of this plant have also been examined to investigate its therapeutic potential on human body. 58 people with type 2 diabetes participated in a 90 day trial where the levels of fasting and postbrandial blood glucose as well as lipids were lowered^[12] Another study used *Gymnema* leaf extract to treat 64 persons with type 1 diabetes for 6 to 30 months, which resulted in reduced plasma glucose levels and need for exogenous insulin.^[31]

BIOAVAILABILITY AND TOXICITY

Gymnemic acid's complicated structure, poor lipid solubility, and difficulty passing through biomembrane make it difficult for the body to absorb it through the circulatory system. A phytosome has a greater ability to pass the lipid bio membrane and enter the systemic circulation, which results in higher absorption and utilisation in the body. In doxorubicin-induced cardiotoxicity in rats, the complex has antiapoptotic potential and cardioprotective effects.^[46] Gymnema sylvestre extract is safe when consumed at the prescribed doses, according to toxicology tests. High dosages may cause adverse reactions such hypoglycemia, muscular dystrophy, weakness, and profuse sweating^[47].No harmful consequences were seen after Wistar rats were given 1.00% baseline powder (GSE) in their food for 52 weeks, and no animals perished as a result of the experiment. Gymnema sylvestre therapy for diabetic individuals has been linked to toxic hepatitis or druginduced liver damage (DILI)^[48]

CONCLUSION AND FUTURE PROSPECTUS

The use of herbal remedies as molecular targets in drug research and in pharmacological applications is on the rise at the moment. Humanity is seriously threatened by the new trend of increasing illness incidence and the difficulties that come along with using prescription drugs. Natural remedies are an alternative to pricey pharmaceuticals since they are relatively safe and have fewer negative effects. Nearly 80% of the population is said to rely on natural therapies for their medical needs. Many bioactive substances, including alkaloids, quinine, paclitaxel, opium alkaloids, quinine, atropine, and cardiac glycosides can be found in plants.

Metformin, the first anti-diabetic medication, was a herbal preparation that was extracted from *Galega officinalis*. The extinction of medicinal plants due to urbanization and industrialization poses a significant threat to their commercial importance. *G sylvestre*, a valuable medicinal plant and source of bioactive substances, is at risk of extinction due to continuous deforestation and lack of established lines or varieties. *In vitro* propagation and cell culture technologies for large-scale production of secondary metabolites with medicinal significance are promising approaches.

Various polyherbal formulations, such as Dihar and D-400 contain *G sylvestre* extract, which has been used for

the treatment of Diabetes mellitus. The herb also has good prospects in dietary applications, with products like GNC Herbal Plus Standardized *G. sylvestre* (herbal supplement), Vitamin Shoppe *G. sylvestre* (sugar destroyer), *Gymnema* gold (Nutrigold), and Gurmar capsules (stimulates the heart and circulatory system and activates the uterus). In conclusion, *G. sylvestre* holds immense potential as a potent medication for diabetes, obesity, cardiovascular disorder and dietary supplements in the food industry.

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