



Review on therapeutic potential of *Gymnema sylvestre*: From traditional ayurvedic medicine to modern applications

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Abstract

In this article, we will explore the rich history of *Gymnema sylvestre*, also known as Gurmar or the "sugar destroyer," and its significance in both traditional Ayurvedic medicine and modern applications. We will delve into the traditional uses of *Gymnema sylvestre* in managing diabetes, controlling sugar cravings, and promoting digestive health, as documented in ancient Ayurvedic texts. Furthermore, we will highlight the global exploration of *Gymnema sylvestre* by European botanists and scientists during the 18th and 19th centuries, which led to the discovery of its unique properties and sparked scientific curiosity. We will discuss the scientific studies that began in the late 19th century, focusing on the identification of active compounds called gymnemic acids, responsible for the plant's anti-diabetic effects. The article will also cover the expanding research on *Gymnema sylvestre*, which uncovered additional health benefits such as its anti-inflammatory, anti-obesity, and anti-microbial properties. The isolation, synthesis, and extensive study of the plant's active compounds will be emphasized. Overall, this article aims to provide an insightful overview of *Gymnema sylvestre*, its historical significance, modern applications, and the need for further research on its active compounds to unlock its full therapeutic potential.

Keywords: *Gymnema sylvestre*, anti-inflammatory, sugar destroyer, gymnemic acid, anti-obesity

Introduction

Gymnema sylvestre, commonly known as Gurmar, is a woody climbing shrub native to the tropical regions of India, Africa, and Australia. The plant has been traditionally used in Ayurvedic medicine to treat various ailments, including diabetes, obesity, and digestive disorders. *Gymnema sylvestre*, a remarkable medicinal plant native to the Indian subcontinent, has a rich history deeply intertwined with traditional Ayurvedic medicine and global exploration. This plant, commonly known as "gurmar" or "sugar destroyer," possesses unique properties that have fascinated scientists and herbalists for centuries. In this article, we delve into the Indian and global history of *Gymnema sylvestre*, tracing its origins, traditional uses, and its significance in modern times.

Indian history

Gymnema sylvestre has a long-standing history in India, where it has been utilized in Ayurvedic medicine for over 2,000 years. The ancient Ayurvedic texts, such as the Charaka Samhita and Sushruta Samhita, mention the therapeutic properties of *Gymnema sylvestre* for treating various ailments. Its Sanskrit name, "gurmar," translates to "destroyer of sugar," indicating its traditional use in managing diabetes and controlling sugar cravings.

In Ayurveda, the leaves of *Gymnema sylvestre* were commonly chewed to suppress the sweet taste and cravings associated with sugar consumption. This practice was believed to support healthy blood sugar levels and aid in weight management. Ayurvedic practitioners also recognized its potential in promoting digestive health, alleviating coughs, and soothing skin conditions.

Global exploration and scientific discoveries

The global exploration era in the 18th and 19th centuries played a pivotal role in introducing *Gymnema sylvestre* to the Western world. European botanists and scientists came across this intriguing plant during their travels to India.

Its unique properties and traditional uses sparked scientific curiosity, leading to further investigation.

In the late 19th century, scientific studies on *Gymnema sylvestre* began to gain momentum. Researchers discovered the presence of active compounds known as gymnemic acids, which were found to be responsible for the plant's anti-diabetic effects.

These gymnemic acids were observed to inhibit the absorption of glucose in the intestines and increase insulin secretion, making *Gymnema sylvestre* a potential therapeutic agent for diabetes management ^[1].

As research progressed, scientists also uncovered additional health benefits of *Gymnema sylvestre* beyond glycemic control ^[2]. Its anti-inflammatory, anti-obesity, and anti-microbial properties were studied, further expanding its potential applications in modern medicine. The plant's active compounds were isolated, synthesized, and studied extensively for their pharmacological activities.

Modern uses and commercial significance

Today, *Gymnema sylvestre* holds significant commercial value as a natural remedy for diabetes and weight management. It is widely available in various forms, including teas, capsules, extracts, and herbal supplements. The anti-diabetic properties of *Gymnema sylvestre* have made it a popular choice for individuals seeking alternative approaches to manage blood sugar levels.



Fig 1: Gymnema products available in market: (1) Gymnema leaf extract. (2) Gymnema Homeopathic liquid. (3) Gymnema capsules. (4) Gymnema Tea

Moreover, *Gymnema sylvestre* extracts are incorporated into dietary supplements and herbal formulas targeting weight loss and appetite control. Its ability to reduce sugar cravings and enhance glucose metabolism has attracted attention from the weight management industry.

While *Gymnema sylvestre* has gained popularity worldwide, it is important to note that scientific research on its efficacy and safety is ongoing [3]. As with any herbal remedy, it is recommended to consult healthcare professionals before incorporating *Gymnema sylvestre* into a treatment regimen.

Morphology

Leaves

The leaves of *Gymnema sylvestre* are simple, opposite, and elliptic or ovate in shape. They are about 2-6 cm in length and 1.5-3.5 cm in width. The leaves are dark green on the upper surface and pale green on the lower surface. They have a smooth texture and a glossy appearance [4].

Stem

The stem of *Gymnema sylvestre* is woody, cylindrical, and climbs with the help of tendrils. The stem can grow up to 10 meters in length and has a diameter of about 1 cm. The stem is grayish-brown in color and has numerous nodes and internodes.

Flowers

The flowers of *Gymnema sylvestre* are small, yellowish-green in color, and occur in clusters of 10-20 flowers. They are about 5 mm in length and have a bell-shaped corolla with five petals. The flowers bloom during the months of July to October.

Fruit

The fruit of *Gymnema sylvestre* is a pod-like capsule that is about 2 cm in length and contains numerous seeds. The capsule is green when immature and turns brown when ripe. The seeds are flat, oval in shape, and about 3 mm in length.

Roots

The roots of *Gymnema sylvestre* are thick, woody, and have a deep taproot system. The roots contain gymnemic acid, which is the active compound responsible for the plant's medicinal properties.

Propagation

Gymnema sylvestre can be propagated through seeds or vegetative means. The seeds of the plant are sown in nursery beds during the monsoon season. Some of the propagation methods for *Gymnema sylvestre* in India:

1. Seed propagation

The seeds of *Gymnema sylvestre* can be used to propagate new plants. The seeds can be sown in a well-draining soil mix and kept moist. Germination usually takes place within two to four weeks. Once the seedlings have developed two or three sets of leaves, they can be transplanted into individual pots.

2. Stem cutting propagation

This method involves taking stem cuttings from an established *Gymnema sylvestre* plant. The cuttings should be around 4-6 inches long and should have several nodes. The lower leaves should be removed, and the cuttings should be planted in a well-draining soil mix. The cuttings should be kept moist and placed in a warm, bright location. Roots should develop in around 2-3 weeks.

3. Division

Mature *Gymnema sylvestre* plants can be divided into smaller sections to propagate new plants. Carefully dig up the plant and separate the root ball into smaller sections, making sure each section has a few stems and roots. Replant the sections in individual pots or directly into the ground.

4. Air layering

This propagation method involves encouraging roots to grow on a stem that is still attached to the parent plant. Choose a healthy stem and make a small incision on the stem. Place a small amount of rooting hormone on the wound and cover it with moist sphagnum moss. Wrap the moss with plastic wrap and secure it in place with string or twist ties. Roots should develop in around 4-6 weeks, and the rooted stem can be cut from the parent plant and potted up.

5. Plant tissue culture

Also known as micropropagation, is a technique used to propagate plants aseptically in a laboratory setting. This technique involves the growth of plant tissues, cells, or organs in an artificial medium under controlled conditions [4].

Methodology

- To initiate a tissue culture of *Gymnema sylvestre*, the first step is to select a healthy plant with desirable traits, such as high medicinal value or rapid growth rate. The plant material is then sterilized to remove any bacteria or fungi that could contaminate the culture.
- Once sterilized, the plant material is cut into small pieces, such as shoot tips, nodal segments, or leaf explants, and placed onto a nutrient-rich agar medium

- that contains plant growth regulators, such as auxins and cytokinins, to promote growth.
- c. The culture is then placed in a growth chamber with controlled environmental conditions, including temperature, light intensity, and humidity, to encourage plant growth. Over time, the plant tissue will proliferate and form a callus, which can be subcultured onto fresh medium to produce multiple copies of the original plant.
 - d. The tissue culture of *Gymnema sylvestre* can be used to produce large quantities of plants with desirable traits for research or commercial purposes. It can also be used to produce disease-free plants and conserve rare or endangered plant species.

To initiate a tissue culture of *Gymnema sylvestre*, the first step is to select a healthy plant with desirable traits, such as high medicinal value or rapid growth rate. The plant material is then sterilized to remove any bacteria or fungi that could contaminate the culture [5].

Once sterilized, the plant material is cut into small pieces, such as shoot tips, nodal segments, or leaf explants, and placed onto a nutrient-rich agar medium that contains plant growth regulators, such as auxins and cytokinins, to promote growth.

The culture is then placed in a growth chamber with controlled environmental conditions, including temperature, light intensity, and humidity, to encourage plant growth. Over time, the plant tissue will proliferate and form a callus, which can be subcultured onto fresh medium to produce multiple copies of the original plant [6].

The tissue culture of *Gymnema sylvestre* can be used to produce large quantities of plants with desirable traits for research or commercial purposes. It can also be used to produce disease-free plants and conserve rare or endangered plant species.

Need of *Gymnema sylvestre*

The active compounds present in *Gymnema sylvestre* include gymnemic acids, which are a group of triterpene saponins, and other compounds such as flavonoids, anthraquinones, and polypeptides. These compounds are responsible for the various medicinal properties of the plant and have been studied extensively by scientists over the years.

One of the main reasons why there is a need for active compounds of *Gymnema sylvestre* is because of its anti-diabetic properties. Diabetes is a chronic metabolic disorder characterized by high blood glucose levels due to either insulin deficiency or insulin resistance. *Gymnema sylvestre* has been found to be effective in lowering blood glucose levels and improving insulin sensitivity, which makes it a promising natural treatment for diabetes.

The active compounds present in *Gymnema sylvestre*, particularly gymnemic acids, have been shown to inhibit the absorption of glucose in the intestine and increase insulin secretion from the pancreas [7]. This helps to regulate blood glucose levels and improve insulin sensitivity in diabetic patients. In addition, *Gymnema sylvestre* has been found to regenerate pancreatic beta cells, which are responsible for insulin secretion. This means that the plant has the potential to treat both type 1 and type 2 diabetes.

Another reason why there is a need for active compounds of *Gymnema sylvestre* is its anti-inflammatory properties.

Inflammation is a natural response of the body to injury or infection, but chronic inflammation can lead to various diseases such as cancer, cardiovascular disease, and autoimmune disorders. *Gymnema sylvestre* has been found to possess potent anti-inflammatory effects, which have been attributed to the presence of flavonoids and other active compounds.

The anti-inflammatory effects of *Gymnema sylvestre* have been demonstrated in various studies [8]. For example, a study published in the Journal of Ethnopharmacology found that gymnemic acids extracted from *Gymnema sylvestre* were effective in reducing inflammation in rats with carrageenan-induced paw edema. Another study published in the Journal of Medicinal Food found that an extract of *Gymnema sylvestre* reduced inflammation in mice with lipopolysaccharide-induced acute lung injury.

Gymnema sylvestre also possesses anti-cancer properties, which make it a promising natural treatment for cancer. Cancer is a complex disease characterized by the uncontrolled growth and spread of abnormal cells. *Gymnema sylvestre* has been found to possess various anti-cancer properties such as inducing apoptosis (programmed cell death) in cancer cells, inhibiting angiogenesis (the formation of new blood vessels) in tumors, and suppressing the metastasis (spread) of cancer cells.

The active compounds present in *Gymnema sylvestre*, particularly gymnemic acids, have been found to induce apoptosis in various cancer cell lines such as breast cancer, prostate cancer, and leukemia [8]. In addition, *Gymnema sylvestre* has been found to inhibit angiogenesis in tumors, which can help to prevent the growth and spread of cancer cells. A study published in the journal Cancer Letters found that an extract of *Gymnema sylvestre* inhibited the growth and metastasis of melanoma cells in mice.

Gymnema sylvestre also possesses anti-obesity properties, which make it a promising natural treatment for obesity. Obesity is a major public health concern and is associated with various health problems such as diabetes, cardiovascular disease, and cancer. *Gymnema sylvestre* has been found to be effective in reducing body weight, body mass index (BMI), and waist circumference, which are all indicators of obesity.

The active compounds present in *Gymnema sylvestre*, particularly gymnemic acids, have been found to inhibit the absorption of fat in the intestine and increase the metabolism of fats in the liver. This helps to reduce the accumulation of fat in the body and promote weight loss. In addition, *Gymnema sylvestre* has been found to suppress the appetite, which can help to reduce food intake and promote weight loss [9].

Overall, the various therapeutic properties of *Gymnema sylvestre* make it a promising natural treatment for various diseases. However, it is important to identify and isolate the active compounds present in the plant in order to develop more effective treatments. In addition, further research is needed to determine the optimal dosage, safety, and efficacy of these compounds.

One of the challenges in isolating and identifying the active compounds of *Gymnema sylvestre* is the complexity of the plant's chemical composition. The plant contains various phytochemicals such as saponins, flavonoids, anthraquinones, and polypeptides, which have different chemical structures and properties. This makes it difficult to isolate and identify the active compounds responsible for the plant's therapeutic properties [10].

However, advances in analytical chemistry and bioassay techniques have made it possible to identify and isolate the active compounds present in *Gymnema sylvestre*. For example, high-performance liquid chromatography (HPLC) and mass spectrometry (MS) have been used to separate and identify the various compounds present in the plant. In addition, bioassay-guided fractionation has been used to identify the fractions of the plant extract that possess the desired therapeutic properties.

One of the active compounds present in *Gymnema sylvestre* that has been extensively studied is gymnemic acid. Gymnemic acid is a group of triterpene saponins that are responsible for the plant's anti-diabetic and anti-obesity properties [11]. Several studies have demonstrated the effectiveness of gymnemic acid in reducing blood glucose levels, improving insulin sensitivity, and promoting weight loss in diabetic and obese individuals.

Another active compound present in *Gymnema sylvestre* is quercetin, a flavonoid that possesses anti-inflammatory and anti-cancer properties. Quercetin has been found to inhibit the production of pro-inflammatory cytokines such as interleukin-1beta (IL-1 β) and tumor necrosis factor-alpha (TNF- α), which are involved in the development of various inflammatory diseases [12]. In addition, quercetin has been found to induce apoptosis in various cancer cell lines such as prostate cancer, breast cancer, and leukemia.

In conclusion, the various therapeutic properties of *Gymnema sylvestre* make it a promising natural treatment for various diseases such as diabetes, inflammation, cancer, and obesity. The active compounds present in the plant, particularly gymnemic acid and quercetin, have been extensively studied and have been found to possess potent therapeutic properties. However, further research is needed to determine the optimal dosage, safety, and efficacy of these compounds.

Cultivation methods and records

The cultivation of *Gymnema sylvestre* involves a series of steps that are designed to optimize plant growth and ensure a high-quality yield. The cultivation methods vary depending on the location and the farmer's preferences. However, the general cultivation practices involve the following:

Soil preparation

The first step in the cultivation of *Gymnema sylvestre* is soil preparation. The soil must be well-drained, with a pH range of 6.0 to 7.5. The soil should be deep and loose to allow the roots to grow and spread easily. It is also important to add organic matter to the soil to improve its fertility and water-holding capacity.

Planting

Gymnema sylvestre can be propagated through seeds or stem cuttings. Seed propagation is less common, as it takes longer for the plant to mature and produce a viable yield. Stem cuttings are the preferred method of propagation, as they produce a higher yield in a shorter amount of time. The cuttings should be taken from healthy, disease-free plants [13].

Irrigation

Gymnema sylvestre requires regular watering, especially during the growing season. The frequency and amount of irrigation will depend on the climate and soil conditions. Overwatering can lead to root rot, while underwatering can stunt growth and reduce the yield.

Fertilization

To ensure a healthy and robust yield, it is important to provide the plant with adequate nutrients. Organic fertilizers are preferred, as they release nutrients slowly and improve soil fertility over time [13]. The fertilizer should be applied in small amounts and spread evenly around the plant.

Cultivation records

India is the largest producer of *Gymnema sylvestre*, with most of the production taking place in the southern states of Tamil Nadu, Kerala, and Karnataka. The cultivation methods used in India vary depending on the location and the farmer's preferences. However, the general cultivation practices involve the steps mentioned above.

In Tamil Nadu, the cultivation of *Gymnema sylvestre* takes place primarily in the districts of Salem, Namakkal, and Erode [14]. The plant is grown as an intercrop with other crops, such as maize and cotton. The cultivation practices in Tamil Nadu involve planting the stem cuttings in well-prepared soil, providing regular irrigation and organic fertilizers, and using natural pest control methods.

In Kerala, the cultivation of *Gymnema sylvestre* takes place primarily in the districts of Thrissur, Palakkad, and Malappuram. The plant is grown as a monocrop or an intercrop with other crops, such as ginger and turmeric [15]. The cultivation practices in Kerala involve planting the stem cuttings in well-drained soil, providing regular irrigation and organic fertilizers, and using natural pest control methods.

In Karnataka, the cultivation of *Gymnema sylvestre* takes place primarily in the districts of Chamarajanagara, Chikkaballapur, and Kolar. The plant is grown as an intercrop with other crops, such as sunflower and maize. The cultivation practices in Karnataka involve planting the stem cuttings in well-drained soil, providing regular irrigation and organic fertilizers, and using natural pest control methods.

In Malaysia, the plant is grown primarily in the states of Johor and Pahang [15]. The cultivation practices in Malaysia involve planting the stem cuttings in well-drained soil, providing regular irrigation and organic fertilizers, and using natural pest control methods.

In China, the plant is grown primarily in the Yunnan and Sichuan provinces. The cultivation practices in China involve planting the stem cuttings in well-drained soil, providing regular irrigation and organic fertilizers, and using natural pest control methods.

In Brazil, the plant is grown primarily in the state of Minas Gerais. The cultivation practices in Brazil involve planting the stem cuttings in well-drained soil, providing regular irrigation and organic fertilizers, and using natural pest control methods.

Constituents	Classification	Chemical Structure	References										
Triterpene saponins	Gymnemic acids-acylated (tigloyl, methylbutyryl) diastereoisomers of deoxygymnemic acid (DACA) which is a 3-O-β-D-glucuronide of gymnemasaponin (Glc-1Rg, 21R, 22R, 23-O, 26-Hydroxycholestan-12-one).		(Liu et al., 1992)										
		<p>Gymnemic acid types</p> <table border="0"> <tr> <td>R1</td> <td>R2</td> </tr> <tr> <td>Gymnemic acid I Tigloyl</td> <td>Ac</td> </tr> <tr> <td>Gymnemic acid II 2-methylbutyryl</td> <td>Ac</td> </tr> <tr> <td>Gymnemic acid III 2-methylbutyryl</td> <td>H</td> </tr> <tr> <td>Gymnemic acid IV Tigloyl</td> <td>H</td> </tr> </table>	R1	R2	Gymnemic acid I Tigloyl	Ac	Gymnemic acid II 2-methylbutyryl	Ac	Gymnemic acid III 2-methylbutyryl	H	Gymnemic acid IV Tigloyl	H	(Yu et al., 2009)
R1	R2												
Gymnemic acid I Tigloyl	Ac												
Gymnemic acid II 2-methylbutyryl	Ac												
Gymnemic acid III 2-methylbutyryl	H												
Gymnemic acid IV Tigloyl	H												
Cleanane Saponins	Gymnemic acids and gymnemasaponins		(Yu et al., 2009)										
		<p>Gymnemasaponin Types</p> <table border="0"> <tr> <td>R1</td> <td>R2</td> </tr> <tr> <td>Gymnemasaponin II H</td> <td>H</td> </tr> <tr> <td>Gymnemasaponin IV Glc</td> <td>H</td> </tr> <tr> <td>Gymnemasaponin V Glc</td> <td>Glc</td> </tr> </table> <p>Glc-β-D-glucopyranosyl</p>	R1	R2	Gymnemasaponin II H	H	Gymnemasaponin IV Glc	H	Gymnemasaponin V Glc	Glc	(Gohji et al., 1998)		
R1	R2												
Gymnemasaponin II H	H												
Gymnemasaponin IV Glc	H												
Gymnemasaponin V Glc	Glc												
Gymnemanol	3,β, 16,β, 22, 4-23-26-pentahydroxycholestan-12-one		(Yoshikawa et al., 1997; Yoshikawa et al., 1998)										
Zammarone Saponins	Gymnemoside A, B, C,D,E,F		(Yoshikawa et al., 1997; Yoshikawa et al., 1998)										
Gymnesticogenin	Pentahydroxytriterpene		(Yoshikawa et al., 1998)										
Surnarin	A 35-Amino acid peptide with a molecular weight of 4209	$-1(\text{Glu}-\text{Gln}-\text{Cys}-\text{Val}-\text{Arg}-\text{Lys}-\text{Asp}-\text{Glu}-\text{Leu}-10\text{Cys}-\text{Ile}-\text{Pro}-\text{Tyr}-15\text{Leu}-\text{Asp}-\text{Cys}-\text{Cys}-\text{Glu}-20\text{Pro}-\text{Leu}-\text{Glu}-\text{Cys}-\text{Lys}-25\text{Lys}-\text{Val}-\text{Asn}-\text{Tyr}-\text{Tyr}-28\text{Asp}-\text{Ile}-\text{Lys}-\text{Cys}-\text{Ile}-\text{Glu})-\text{NH}_2$ (Glu = pyrrolysine-acid)	(Imoto et al., 1991)										
Triterpenoid saponins Gymnemasin A Gymnemasin B Gymnemasin C Gymnemasin D	3-O-β-D-glucopyranosyl (1-3)-β-D-glucopyranosyl-22-O-tigloyl-gymnemanol 3-O-β-D-glucopyranosyl-(1-3)-β-D-Glucuronopyranosyl-Gymnemanol 3-O-β-D-glucuronopyranosyl-22-O-tigloyl-gymnemanol 3-O-β-D-glucopyranosyl-gymnemanol		(J-GLCBAL-Japan Science and Technology Agency, n.d.a; J-GLCBAL-Japan Science and Technology Agency, n.d.b; J-GLCBAL-Japan Science and Technology Agency, n.d.c; J-GLCBAL-Japan Science and Technology Agency, n.d.d)										
Flavonol glycoside	Kaempferol 3-O-β-D-glucopyranosyl-(1-4)-α-L-rhamnopyranosyl-(1-6)-β-D-glucopyranoside		(Liu et al., 2004)										
Steroids	Stigmasterol		(Potawale et al., 2008)										
Lapoeol			(Bischofberger et al., 1973)										
β-Quercitol			(Potawale et al., 2008)										
Paraben	Quercitol		(Bischofberger et al., 1973)										
	Quercitol		(Thwart et al., 2014)										
	Condufitol A		(Thwart et al., 2014)										

Fig 2: Phytoconstituents of *gymnema sylvestre* plant

Phytochemistry

The stems of *Gymnema sylvestre* have been found to contain therapeutically important chemical compounds such as stigmasterol and triterpenoid saponin. Stigmasterol has been shown to have antidiabetic, hypoglycemic, antioxidant, and anticancer activities, while triterpenoid saponins have been found to exhibit anti-tumor, anti-fungal, hepatoprotective, and antidiabetic potential. The plant also contains gymnemic acids and gymnemasaponins, which are classified as oleanane saponins [16]. The leaves of the plant contain saponins, anthraquinones, cardiac glycosides, and other compounds such as tannin, quinones, flavonoids, and phenols. The detailed list of phytochemical compounds found in the analysis of *Gymnema sylvestre* is provided in the table below.

Gymnema sylvestre contains several bioactive compounds that are responsible for its therapeutic effects [17]. Some of the active compounds of *Gymnema sylvestre* include:

1. **Gymnemic acids:** These are a group of compounds that are found in the leaves of the plant. They are known for their ability to reduce the absorption of sugar in the body.
2. **Saponins:** These are plant compounds that have a soapy texture and are commonly found in various plant species. Saponins have been shown to have antioxidant, anti-inflammatory, and cholesterol-lowering properties.
3. **Flavones:** These are a type of flavonoid that have antioxidant and anti-inflammatory properties. Flavones are commonly found in various plant species and are known to have several health benefits.
4. **Alkaloids:** These are a group of nitrogen-containing compounds that are found in various plant species. Some alkaloids have been shown anti-inflammatory properties.

Other compounds found in *Gymnema sylvestre* include tannins, anthraquinones, and phytosterols [18]. Each of these compounds has its unique health benefits and contributes to the overall therapeutic properties of the plant.

Chemical basis of mode of action of gymnemic acid

Gymnemic acid is a group of triterpene saponins that are primarily responsible for the therapeutic properties of *Gymnema sylvestre*. These compounds have been extensively studied and have been found to possess various biological activities such as anti-diabetic, anti-obesity, and anti-inflammatory properties [19].

The chemical structure of gymnemic acid consists of a triterpene moiety and one or more sugar residues. The triterpene moiety is composed of a cycloartane ring system

with several hydroxyl groups and a carboxyl group. The sugar residues are usually attached to the hydroxyl groups of the triterpene moiety and can vary in their number and position.

The mode of action of gymnemic acid is primarily based on its interaction with various molecular targets in the body [20]. One of the primary targets of gymnemic acid is the taste receptors on the tongue, which are responsible for detecting the sweet taste of carbohydrates.

Gymnemic acid has been found to interact with the sweet taste receptors and block the sensation of sweetness. This can help to reduce the craving for sweet foods and reduce the intake of carbohydrates. In addition, gymnemic acid has been found to reduce the absorption of glucose in the intestine and increase the uptake of glucose in the peripheral tissues such as skeletal muscle and adipose tissue.

Gymnemic acid has also been found to increase insulin secretion from the beta cells of the pancreas. Insulin is a hormone that regulates the uptake and utilization of glucose by the cells in the body. By increasing insulin secretion, gymnemic acid can help to reduce blood glucose levels and improve insulin sensitivity [21].

Furthermore, gymnemic acid has been found to inhibit the activity of alpha-amylase and alpha-glucosidase, which are enzymes involved in the breakdown of carbohydrates in the intestine. By inhibiting these enzymes, gymnemic acid can reduce the absorption of carbohydrates and help to lower blood glucose levels.

In addition to its anti-diabetic properties, gymnemic acid has also been found to possess anti-obesity and anti-inflammatory properties. Gymnemic acid has been found to inhibit the absorption of fat in the intestine and increase the metabolism of fats in the liver. This can help to reduce the accumulation of fat in the body and promote weight loss. In addition, gymnemic acid has been found to inhibit the production of pro-inflammatory cytokines such as interleukin-6 (IL-6) and tumor necrosis factor-alpha (TNF- α). This can help to reduce inflammation in the body and prevent the development of various inflammatory diseases [22].

Overall, the mode of action of gymnemic acid is primarily based on its interaction with various molecular targets in the body such as taste receptors, enzymes, and hormones. The interaction of gymnemic acid with these targets can help to reduce blood glucose levels, improve insulin sensitivity, promote weight loss, and reduce inflammation. Further research is needed to determine the optimal dosage, safety, and efficacy of gymnemic acid for various health conditions [23].

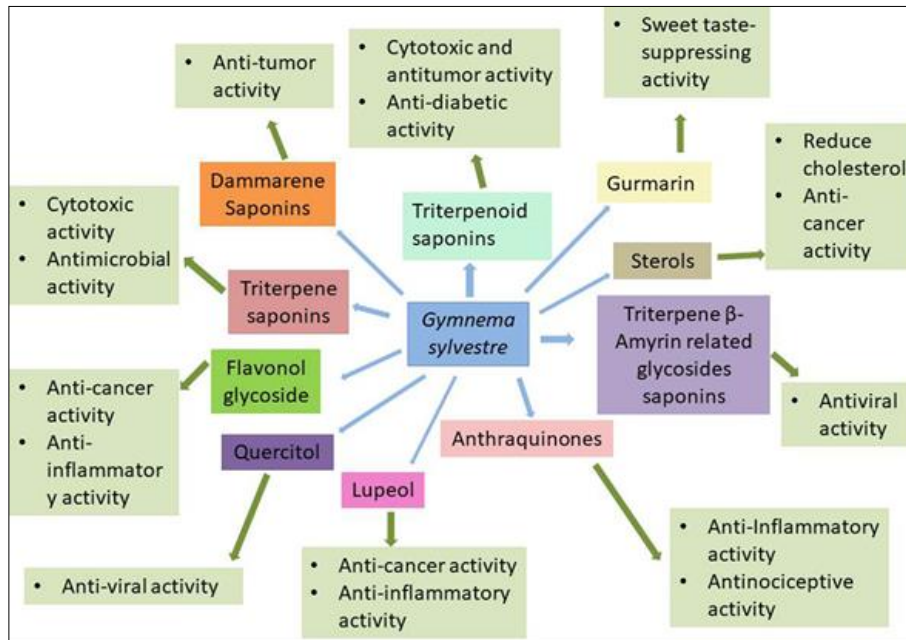


Fig 3: Pharmacological activities of the constituents of *Gymnema sylvestre*

Molecular basis of mode of action

The molecular basis of the mode of action of gymnemic acid involves the interaction of this compound with various molecular targets in the body, leading to its therapeutic effects. The primary molecular targets of gymnemic acid include taste receptors, enzymes, and hormones involved in glucose and lipid metabolism [24].

One of the primary molecular targets of gymnemic acid is the sweet taste receptors on the tongue. Gymnemic acid has been found to interact with these receptors and block the sensation of sweetness. This can help to reduce the craving for sweet foods and lower the intake of carbohydrates. The molecular mechanism of this interaction involves the binding of gymnemic acid to the sweet taste receptors, leading to a conformational change in these receptors that inhibits their function. The exact molecular details of this interaction are not yet fully understood and require further research.

Another important molecular target of gymnemic acid is the enzyme alpha-amylase, which is involved in the breakdown

of complex carbohydrates into simple sugars in the intestine [25]. Gymnemic acid has been found to inhibit the activity of alpha-amylase, leading to a reduction in the absorption of carbohydrates and a lowering of blood glucose levels. The molecular mechanism of this inhibition involves the binding of gymnemic acid to the active site of alpha-amylase, leading to a disruption of the enzyme-substrate complex and inhibition of enzymatic activity.

Gymnemic acid also interacts with another enzyme involved in carbohydrate metabolism, alpha-glucosidase. This enzyme is responsible for the breakdown of disaccharides and oligosaccharides into monosaccharides, which can be absorbed by the intestine. Gymnemic acid has been found to inhibit the activity of alpha-glucosidase, leading to a further reduction in the absorption of carbohydrates and a lowering of blood glucose levels [26]. The molecular mechanism of this inhibition involves the binding of gymnemic acid to the active site of alpha-glucosidase, leading to a disruption of the enzyme-substrate complex and inhibition of enzymatic activity.

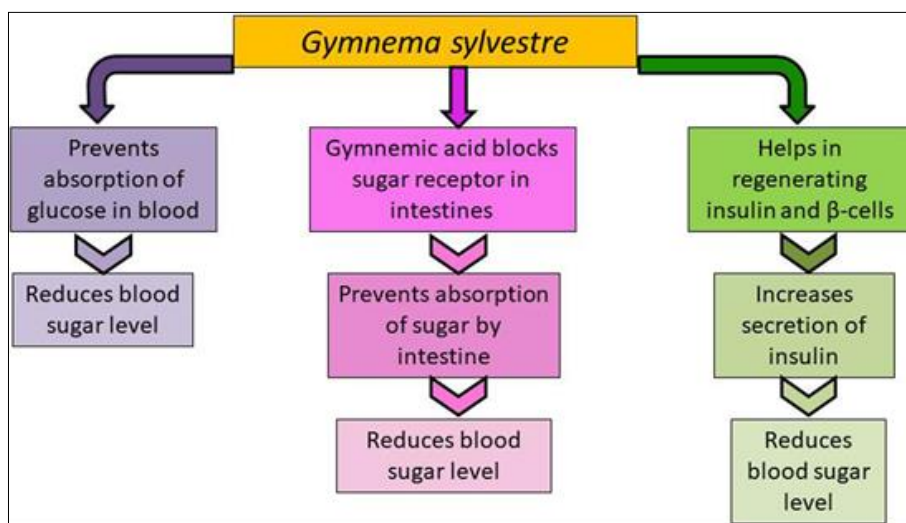


Fig 4: Mechanism of glucose lowering activity by *Gymnema sylvestre*.

Another hormone involved in lipid metabolism that is affected by gymnemic acid is adiponectin. This hormone is secreted by adipose tissue and is involved in the regulation of glucose and lipid metabolism. Gymnemic acid has been found to increase the secretion of adiponectin, leading to an improvement in insulin sensitivity and a reduction in inflammation [27]. The molecular mechanism of this interaction involves the binding of gymnemic acid to specific receptors on adipose tissue, leading to an activation of downstream signaling pathways that promote adiponectin secretion.

Overall, the molecular basis of the mode of action of gymnemic acid involves the interaction of this compound with various molecular targets in the body, leading to its therapeutic effects. These interactions involve the binding of gymnemic acid to specific receptors and enzymes, leading to a disruption of their function and subsequent modulation of glucose and lipid metabolism. Further research is needed to fully understand the molecular details of these interactions and to develop new therapeutic applications of gymnemic acid.

Conclusion

Gymnema sylvestre, commonly known as Gurmar, is a remarkable medicinal plant with a rich history rooted in traditional Ayurvedic medicine. Its traditional uses in managing diabetes, controlling sugar cravings, and promoting digestive health have been recognized for centuries. Through global exploration and scientific discoveries, researchers have uncovered the plant's active compounds, such as gymnemic acids, and their potential therapeutic effects.

The modern uses of *Gymnema sylvestre* have expanded, with commercial products available in various forms, including teas, capsules, extracts, and herbal supplements. It has gained popularity as a natural remedy for diabetes and weight management, offering an alternative approach for individuals seeking to manage blood sugar levels and control cravings. However, ongoing scientific research is necessary to fully understand its efficacy and safety.

The propagation of *Gymnema sylvestre* can be achieved through various methods, such as seed propagation, stem cutting propagation, division, air layering, and plant tissue culture. These techniques enable the production of large quantities of plants with desirable traits, including high medicinal value or rapid growth rate, for research or commercial purposes. Plant tissue culture, in particular, provides a controlled environment for the growth and proliferation of plant tissues, offering opportunities for disease-free plant production and conservation of rare or endangered species.

The active compounds present in *Gymnema sylvestre*, particularly gymnemic acids, hold promise for addressing various health concerns. Their anti-diabetic properties, anti-inflammatory effects, anti-cancer properties, and anti-obesity effects have been studied extensively. However, further research is needed to optimize dosage, ensure safety, and determine efficacy in treating specific diseases.

The complexity of *Gymnema sylvestre*'s chemical composition poses challenges in isolating and identifying its active compounds. However, advancements in analytical chemistry and bioassay techniques have facilitated the identification and isolation of these compounds, such as

gymnemic acid and quercetin. These discoveries contribute to a better understanding of the plant's therapeutic potential. In conclusion, *Gymnema sylvestre* represents a valuable medicinal plant with a long-standing history in traditional medicine. Its global exploration and scientific discoveries have shed light on its active compounds and expanded its applications in modern medicine. Continued research and exploration are crucial for unlocking its full potential and harnessing the benefits of this remarkable plant for human health.

Future perspective

Gymnema sylvestre, with its rich history and proven medicinal properties, holds great potential for the future. As scientific research continues to unravel its complex chemical composition and therapeutic effects, there are several promising areas where *Gymnema sylvestre* can make a significant impact.

1. **Diabetes Management:** *Gymnema sylvestre* has shown promising results in managing diabetes by lowering blood glucose levels and improving insulin sensitivity. With the growing prevalence of diabetes worldwide, there is a need for effective and natural alternatives to traditional medications. Further research on the active compounds of *Gymnema sylvestre* and their mechanisms of action can lead to the development of more targeted and potent treatments for diabetes.
2. **Weight Management:** The anti-obesity properties of *Gymnema sylvestre* make it a valuable candidate for weight management interventions. By suppressing appetite, inhibiting fat absorption, and increasing fat metabolism, *Gymnema sylvestre* has the potential to play a role in combating the global obesity epidemic. Future studies can explore the optimal dosage, long-term effects, and combination therapies involving *Gymnema sylvestre* for sustainable weight loss.
3. **Anti-inflammatory and Anticancer Applications:** *Gymnema sylvestre*'s anti-inflammatory properties and ability to induce apoptosis in cancer cells make it a promising candidate for the development of novel anti-inflammatory drugs and anticancer therapies. Further research can focus on identifying specific compounds within *Gymnema sylvestre* that exhibit potent anti-inflammatory and anticancer activities, and exploring their mechanisms of action for targeted treatments.
4. **Digestive Health:** Traditional Ayurvedic medicine recognizes *Gymnema sylvestre*'s potential in promoting digestive health. Ongoing research can further investigate its effects on gut health, gastrointestinal disorders, and the gut microbiome. Understanding the interactions between *Gymnema sylvestre* and the gut microbiota can open up new avenues for personalized medicine and interventions targeting digestive disorders.
5. **Plant Breeding and Cultivation:** *Gymnema sylvestre* is primarily grown in the tropical regions of India, Africa, and Australia. However, with increasing global demand and interest in its medicinal properties, there is a need to develop improved cultivars and cultivation techniques to ensure a sustainable supply. Plant breeding efforts can focus on selecting varieties with higher concentrations of active compounds, enhanced yield, and improved disease resistance.

6. Formulation Development: *Gymnema sylvestre* extracts are currently available in various forms, including teas, capsules, and herbal supplements. Future research can explore innovative formulation approaches, such as nanoparticles or controlled-release systems, to improve bioavailability and optimize the delivery of active compounds. This can enhance the efficacy and convenience of *Gymnema sylvestre*-based products for consumers.
7. Safety and Standardization: While *Gymnema sylvestre* is generally considered safe, further studies are needed to evaluate its long-term safety profile, potential drug interactions, and optimal dosages. Standardization of *Gymnema sylvestre* extracts and products is crucial to ensure consistent quality and therapeutic efficacy. Establishing standardized protocols for cultivation, extraction, and manufacturing processes can contribute to the development of reliable and regulated *Gymnema sylvestre*-based products.

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