



## INDIGENOUS REMEDIES FOR DIABETES MELLITUS

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## ABSTRACT

Diabetes mellitus (DM), a global public health problem, is now emerging as an epidemic world over. According to a widely accepted estimation, the number of diabetic patients would reach 366 million by the year 2030. India now has the world's largest diabetic population, encompassing an estimated 35 million people out of an overall population of 1 billion. In just over 20 years (i.e. 2025) the country will have almost 200 million people (approximately 15% of the population) affected by diabetes or its precursor. In India, indigenous remedies have been used in the treatment of DM since the time of Charaka and Sushruta (6th century BC). The ethnobotanical information reports about 800 plants that may possess anti-diabetic potential. Several such herbs have shown anti-diabetic activity when assessed using presently available experimental techniques. The present review, deals with some selective Indian medicinal plants having pharmacologically established hypoglycemic potential.

**Keywords:** Diabetes mellitus, indigenous remedies

## INTRODUCTION

Diabetes mellitus (DM), a global public health problem, is now emerging as an epidemic world over. According to a widely accepted estimation, the number of diabetic patients would reach 366 million by the year 2030. India now has the world's largest diabetic population, encompassing an estimated 35 million people out of an overall population of 1 billion. In just over 20 years (i.e. 2025) the country will have almost 200 million people (approximately 15% of the population) affected by diabetes or its precursor<sup>1</sup>. Diabetes mellitus is group of syndrome characterized by hyperglycemia altered metabolism of lipids, carbohydrates and proteins with an increase risk complications from vascular disease<sup>2</sup>. It was reported that there is a higher incidence of retinopathy, neuropathy, nephropathy etc. together with diabetes. A wide spread pathological change is thickening of capillary membrane, increase in vessel wall matrix and cellular proliferation resulting in vascular complication like lumen narrowing, early atherosclerosis, sclerosis of glomerular capillaries, retinopathy, neuropathy and vascular insufficiency.

It may affect the disruption of carbohydrate and fat metabolism<sup>3</sup>. Diabetes is a metabolic disorder where in human body does not produce or properly uses insulin, a hormone that is required to convert sugar, starches, and other food into energy<sup>4</sup>. Human body has to maintain the blood glucose level at a very narrow range, which is done with insulin and glucagon. The function of glucagon is causing the liver to release glucose from its cells into the blood, for the production of energy. The condition may be multifactorial origin in which heredity, age, sex, pregnancy, obesity, autoimmune, infections and emotional disturbances may be important. It may precipitated by pancreatic disorders, hormonal disorders (e.g. acromegaly and cushing syndrome), or by administration of drugs (corticosteroids or diuretic, especially thiazides)<sup>5</sup>. There are mainly two types of diabetes—Type 1 and Type 2. In Type 1 diabetes, in the absence of pancreatic b-cells the hormone insulin is not produced while Type 2 diabetes mellitus (T2DM), is characterized by a progressive impairment of insulin secretion by pancreatic b-cells

and by a relative decreased sensitivity of target tissues to the action of this hormone<sup>6</sup>. Gestation diabetes mellitus (GDM) is glucose intolerance being recognized during pregnancy. It can complicate pregnancy leading to prenatal morbidity and mortality, so clinical detection important. In India, indigenous remedies have been used in the treatment of DM since the time of Charaka and Sushruta (6th century BC). Plants have always been an exemplary source of drugs and many of the currently available drugs have been derived directly or indirectly from them. The ethnobotanical information reports about 800 plants that may possess anti-diabetic potential<sup>7</sup>. Several such herbs have shown anti-diabetic activity when assessed using presently available experimental techniques<sup>8</sup>. Although, oral hypoglycemic agents/insulin is the mainstay of treatment of diabetes and is effective in controlling hyperglycemia, they have prominent side effects and fail to significantly alter the course of diabetic complications<sup>9</sup>. As the knowledge of heterogeneity of this disorder increases, there is needed to look for more efficacious agents with lesser side effects. Though development of modern medicine resulted in the advent of modern pharmacotherapeutics including insulin, biguanides, sulfonylureas and thiazolidinediones, there is still a need to look for new drugs as no drug (except strict glycemic control with insulin) has been shown to modify the course of diabetic complications<sup>10</sup>. Apart from currently available therapeutic options, many herbal medicines have been recommended for the treatment of diabetes. Traditional plant medicines are used throughout the world for a range of diabetic presentations. Herbal drugs are prescribed widely because of their effectiveness, less side effects and relatively low cost. Therefore, investigation on such agents from traditional medicinal plants has become more important. India has a rich history of using various potent herbs and herbal components for treating diabetes. Many Indian plants have been investigated for their beneficial use in different types of diabetes and reported in numerous scientific journals. The present review, deals with some selective Indian medicinal plants having pharmacologically established hypoglycemic potential<sup>11</sup>.

Table 1: Selected Indian Medicinal Plants With Hypoglycemic Potential<sup>10,11</sup>

Sr. No	Name Of Plant	Part Used	Model Used	Reported Mechanism of Action	Reference
1	<i>Acacia arabica</i> (Lam.) Muhl. ex Willd. <b>Common name:</b> Indian Gum Arabic tree [Family: Leguminosae]	Seed, powdered seed (2, 3 and 4 mg/kg)	Normal rats, alloxan rats, rabbits	Acts through release of insulin from pancreatic beta cells.	12,13
2	<i>Aegle marmelos</i> (L.) Correa ex Roxb. <b>Common name:</b> Holy Fruit Tree [Family: Rutaceae]	root bark (1 ml/100mg), aqueous leaf extract	Normal fasted rats,	Increases utilization of glucose	14-16

3	<i>Allium cepa L.</i> <b>Common name:</b> Onion [Family:Liliaceae]	ether soluble fraction of onion (0.25 mg/kg p.o.),	STZ rats	Lowers blood glucose level	17-19
4	<i>Allium sativum L.</i> <b>Commonname:</b> Garlic [Family: Alliaceae]	ethanol, petroleum ether and ethyl acetate extract (0.25 mg/kg)	Alloxan rabbits	Has strong antioxidant activity and rapid reactivity with thiol containing proteins responsible for the hypoglycemic property.	20-21
5	<i>Aloe vera (L.) Burm.f.</i> <b>Common name:</b> Aloe [Family: Aloaceae]	leaf pulp extracts	STZ rats	Maintains glucose homeostasis by controlling the carbohydrate metabolizing enzymes and stimulates insulin release from pancreatic beta cells.	22-24
6	<i>Artemisia pallens Wall</i> <b>Common Name:</b> Davana [Family:Compositae]	aerial parts (100 mg/kg, orally)	Alloxan rats	Inhibits glucose re-absorption or increase in peripheral glucose utilization	25
7	<i>Annona squamosa L.</i> <b>Common name:</b> Sugar apple [Family:Annonaceae]	ethanolic leaf-extract (350 mg/kg, orally)	STZ rats and alloxan rabbits	Lowers blood glucose level	26-27
8	<i>Andrographis paniculata</i> <b>Nees</b> <b>Common name:</b> King of Bitter [Family:Acanthaceae]	Andrographis paniculata extract	STZ rats	Prevents glucose absorption from gut.	28-30
9	<i>Azadirachta indica A.Juss.</i> <b>Common name:</b> Neem [Family: Meliaceae]	Hydro alcoholic plant extract, crude ethanolic extract of the plant	STZ rats, alloxan albino rats	Inhibits action of epinephrine on glucose metabolism, resulting in increased utilization of peripheral glucose	31-34
10	<i>Amaranthus esculents</i> [Family: Amaranthaceae]	whole plant , oil fraction	STZ rats	☒Glucose; ☒insulin	35
11	<i>Biophytum sensitivum (L.)</i> <b>Common name:</b> Life Plant [Family: Oxalidaceae]	plant leaf extract	Alloxan rabbits	Stimulates pancreatic beta cells to release insulin	36
12	<i>Boerhavia diffusa L.</i> <b>Common name:</b> Tar vine [Family:Nyctaginaceae]	aqueous leaf extract (100, 200 and 400 mg/kg )	Alloxan rats	Increases plasma insulin levels and improves glucose tolerance	37-38
13	<i>Cassia auriculata L.</i> <b>Common name:</b> Tanner's Cassia [Family:Leguminosae]	aqueous flower extract	STZ rats	Suppresses enhanced gluconeogenesis during diabetes and enhance utilization of glucose through increased glycolysis. in addition to pronounced alpha-glucosidase inhibitory actions resulting in a significant and potent lowering of blood glycemc response	39
14	<i>Caesalpinia bonducella (L.)Roxb.</i> <b>Common name:</b> Chinese Cinnamon [Family: Caesalpinaceae]	aqueous and 50% ethanolic seed extracts	STZ rats	Increases the release of insulin from pancreatic cells	40
15	<i>Citrullus colocynthis (L.)Schrud.</i> <b>Common name:</b> Bitter apple [Family: Cucurbitaceae]	aqueous extract (300 mg/kg)	Normal rabbits, STZ rats	Exerts an insulinotropic effect	41-42
16	<i>Coccoloba indica Wight &amp; Arn.</i> <b>Common name:</b> Ivy gourd [Family: Cucurbitaceae]	alcoholic leaf extract	Guinea pig, Alloxan dogs	Suppresses glucose synthesis, through depression of the key gluconeogenic enzymes glucose-6-phosphatase and fructose-1,6-bisphosphatase and enhances glucose oxidation by shunt pathway through activation of its principal enzyme glucose-6-phosphate dehydrogenase. Also has an insulin secretagogue effect and acts like insulin by correcting elevated enzymes in glycolytic pathway and restoring LPL activity in lypolytic pathway with control of hyperglycemia in diabetes	43-48
17	<i>Casearia esculenta Roxb.</i> <b>Common name:</b> Carilla Fruit [Family:Flacourtiaceae]	root extracts (300 mg/kg p.o.)	STZ rats	Exhibits significant reduction in blood glucose level, a decrease in the activities of glucose-6-phosphatase and fructose-1,6-bisphosphatase and an increase in the activity of liver hexokinase, resulting in potent hypoglycemic activity	49
18	<i>Catharanthus roseus (L.) G.</i>	ethanolic leaf	STZ rats	Increases metabolism of glucose and	50-52

	<i>Don</i> <b>Common name:</b> Madagascarperiwinkle [Family: Apocynaceae]	extract (500mg/kg), dichloromethane: methanol extract of leaves and twigs((500 mg/kg p.o., for 7 and 15 days)		enhances secretion of insulin either from the beta cells of Langerhans or through extrapancreatic mechanism	
19	<i>Camellia sinensis Kuntze</i> <b>Common name:</b> Green tea [Family: Theaceae]	hot water extract of green tea	STZ rats	increases insulin activity	53-54
20	<i>Chamaemelum nobile</i> [Family: Asteraceae]	aqueous extract of leaves	STZ rats	☑Glucose	55
21	<i>Coscinium fenestratum</i> [Family: Menispermaceae]	alcoholic extract/steam barks	STZ rats	☑Glucose, ☑glycosylated hemoglobin, ☑glycogen, ☑lipids, ☑oxidative stress	56
22	<i>Encostemma littorale Blume</i> [Family: Gentianeaceae]	aqueous extract of plant	Alloxan rats	Enhances glucose-induced insulin release from isolated rat pancreatic islets, mediated through K (+)-ATP channel-dependent pathway	57-58
23	<i>Eugenia jambolana Lam.</i> <b>Common name:</b> Indian black berry [Family: Myrtaceae]	pulp extract of the fruits, alcoholic extract (100 mg/kg p.o.)	STZ rats, alloxan rats	May be mediated through an insulin release mechanism or due to alteration in hepatic and skeletal muscle glycogen content and hepatic glucokinase, hexokinase, glucose-6-phosphate and phosphofructokinase levels in diabetic mice. It also enhances serum insulin act and exhibits normoglycemia and better glucose tolerance	59-64
24	<i>Egyptian Morus alba</i> [Family: Moraceae ]	alcoholic extract	STZ rat	☑Glucose, ☑lipid peroxidation, ☑insulin	65
25	<i>Ficus bengalensis L.</i> <b>Common name:</b> Banyan tree [Family: Moraceae]	bark extract	STZ rats, alloxan rats	Stimulates insulin secretion from beta cells of islets of Langerhans and inhibits insulin degradative process.	66-70
26	<i>Hibiscus rosa sinensis L.</i> <b>Common name:</b> China Rose [Family: Malvaceae]	ethanol extract of the plant, alcoholic leaf extract (250 mg/kg p.o. for seven days consecutive)	STZ rats	Stimulates insulin secretion from pancreatic beta cells and increases utilization of glucose, either by direct stimulation of glucose uptake or via the mediation of enhanced insulin secretion	71-72
27	<i>Helicteres isora L.</i> <b>Common name:</b> Screw tree [Family: Sterculiaceae]	ethanolic root extract (300 mg/kg, after 9 days of administration	Mice	Acts through insulin-sensitizing activity	73-74
28	<i>Hintonia standleyan</i> [Family: Rubiaceae]	methanolic extract	STZ rat	☑Glucose	75-76
29	<i>Hypoxis hemerocallidea</i> [Family: Hypoxodaceae]	Aqueous extract/fruits	STZ mice , rat	☑Glucose	77
30	<i>Leonotis leonurus</i> [Family: Lamiaceae]	Aqueous extract/leaves	STZ mice , rat	☑Glucose	78
31	<i>Lepidium sativum</i> [Family: Brassiaceae]	Aqueous extract/leaves	STZ rat	☑Glucose	79
32	<i>Lycium barbarum</i> [Family: Solanaceae]	Isolated compounds /fruits	STZ rat	☑Glucose, ☑oxidative stress, ☑GLUT4, ☑insulin	80-81
33	<i>Mangifera indica L.</i> <b>Common name:</b> Mango [Family:Anacardiaceae]	Aqueous leaf extracts (1 g/kg p.o.)	STZ rats	Possibly acts through intestinal reduction of the absorption of glucose as well as pancreatic and extrapancreatic mechanisms	82-83
34	<i>Momordica cymbalaria</i> Fenzl [Family: Cucurbitaceae]	fruit powder, aqueous fruit extract (0.5 g/kg dose for 6 weeks)	Alloxan rats	May act by increasing hepatic glycogen	84-85
35	<i>Mucuna pruriens (L.)</i> <b>Common name:</b> Velvet bean [Family: Leguminosae]	powdered seeds (0.5, 1 and 2 g/kg), plant extract (200 mg/kg), alcohol extract of the plant (100, 200 and 400	Alloxan rabbits, STZ mice	Possibly acts through stimulation of the release of insulin and/or by a direct insulin-like action due to the presence of trace elements like manganese, zinc, etc.	86-88

36	<i>Morus alba L.</i> <b>Common name:</b> White mulberry [Family: Moraceae]	mg/kg/day) aqueous extract of leaves(200mg/kg)	Mice	Acts by increasing glucose uptake	89
37	<i>Murraya koenigii (L.) Spreng.</i> <b>Common name:</b> curry-leaf tree [Family: Rutaceae]	leaf-powder	Normal rats	Increases glycogenesis and decreases glycogenolysis and gluconeogenesis	90-91
38	<i>Malmea depressa</i> [Family: Annonaceae]	aqueous extract, ethanolic extract , n-butenol fraction/roots	STZ rats	☑Glucose	92
39	<i>Mangifera indica</i> [Family: Anacardiaceae]	aqueous extract/stem barks	STZ rats	☑Glucose	93
40	<i>Momordica charnita</i> [Family: Cucurbitaceae]	methanolic extract, isolated compounds /gourd Aqueous extract/leaves	STZ rats, STZ mice Alloxan-rat	☑Glucose	94-95
41	<i>Ocimum sanctum L.</i> <b>Common name:</b> Holy Basil [Family: Lamiaceae]	ethanolic leaf extract, plant extract (200 mg/kg for 30 days)	STZ rat	Acts by cortisol inhibiting potency	96-98
42	<i>Punica granatum L.</i> <b>Common name:</b> Pomegranate [Family: Punicaceae]	ethanolic flower extract, plant extract (200 mg/kg for 30 days)	STZ rat	Inhibits intestinal alpha-glucosidase activity, leading to antihyperglycemic property	99-100
43	<i>Piper betle</i> [Family: Piperaceae]	aqueous extract, ethanolic extract/leaves	STZ rat	☑Glucose, ☑glycosylated hemoglobin	101-102
44	<i>Psidium guajava Linn.</i> [Family: Myrtaceae]	aqueous extract/whole plant	STZ rat	☑Glucose, ☑lipids, ☑insulin	103
45	<i>Raphanus sativus</i> (Brassicaceae)	aqueous extract /whole plant	STZ rat, mice	☑Glucose, ☑lipids, ☑insulin	104
46	<i>Retama raetam</i> [Family: Fabaceae]	aqueous extract/whole plant	STZ rat	☑Glucose	105
47	<i>Strobilanthes crispus</i> [Family: Acanthaceae]	aqueous extract/leaves	STZ rat	☑Glucose	106
48	<i>Syzygium cordatum</i> [Family: Myrtaceae]	aqueous extract/leaves	STZ rat	☑Glucose, ☑hepatic glycogen	107
49	<i>Salacia reticulata Wight.</i> <b>Common name:</b> Salacia [Family: Celastaceae]	aqueous decoction	STZ rat	Inhibits alpha-glucosidase activity	108
50	<i>Salacia Oblonga Wall.</i> [Family: Celastaceae]	aqueous methanolic extract, aqueous methanolic extract	Zucker rat (OZR)	Acts through inhibition of alpha-glucosidase activity	109
51	<i>Sida cordifolia L.</i> [Family: Malvaceae]	methanolic extract/root	STZ rat	☑Glucose	110
52	<i>Swertia chirayita (Roxb. Ex Fleming) H. Karst.</i> <b>Common name:</b> Indian Gentian [Family: Gentianaceae]	ethanol extract (250 mg/kg)	STZ rat	Stimulates insulin release from islets of Langerhans by depleting aldehyde-fuchsin stained beta-granules and immunostained insulin	111-112
53	<i>Syzygium alternifolium</i> Walp. [Family: Myrtaceae]	Aqueous Extract/seed	Alloxan rats	☑Glucose	113
54	<i>Scoparia dulcis L.</i> <b>Common name:</b> Sweet Broomweed [Family: Scrophulariaceae]	aqueous leaf extract (0.15, 0.30 and 0.45 g/kg body weight for 45	STZ rat , mice	Suppresses glucose influx into the polyol pathway leading to increased activities of antioxidant enzymes and plasma insulin and decreases activity of sorbitol dehydrogenase	114-115

		days p.o.), plant extracts		Also potentiates insulin release from Blood glucose, sorbitol dehydrogenase, glycosylated pancreatic islets.	
55	<i>Taxus yunnanensis</i> [Family: Taxaceae]	aqueous extract, methenolic extract, isolated compounds/ woods	STZ rat	☒Glucose	116
56	<i>Terminalia catappa L.</i> [Family: Combretaceae]	Pet-ether, Methanol, aqueous extract/fruit	Alloxan rats	☒Glucose	117
57	<i>Terminalia chebula</i> [Family: Combretaceae]	chloroform exreacr /seeds	STZ rat	☒Glucose	118
58	<i>Terminalia superb</i> [Family: Combretaceae]	methenolic extract, methylene chloride extract/steam barks	STZ rat	☒Glucose	119
59	<i>Trema orientalis</i> [Family: Ulmaceae]	aqueous extract/steam bark	STZ rat	☒Glucose	120
60	<i>Tremella mesenteric</i> [Family: Combretaceae]	isolated compounds/fruits	STZ rat	☒Glucose	121
61	<i>Triticum repens</i> P.Beauv. [Family: Gramineae]	aqueous extract/rhizomes	STZ rat	☒Glucose	122
62	<i>Viscum album L.</i> [Family: Llorenthaceae]	aqueous extract, ethanolic extract/whole plant	STZ rat	☒Glucose, ☒lipidic Peroxidation	123
63	<i>Zizyphus spina-christi</i> [Family: Rhamnaceae]	n-butenol fraction, isolated compounds/l eaves	STZ rat	☒Glucose, ☒insulin	124
64	<i>Smilax chinensis (L.)</i> [Family:Liliaceae]	ethanolic extract, dried rhizomes	Alloxan rats	Reduces blood glucose level	125
65	<i>Abelmoschus esculentus</i> Linn [Family: Malvaceae]	aqueous extract, fruit	Alloxan mice	Reduces blood glucose level	126

## CONCLUSION

The incidences of modern lifestyle diseases like Type 2 diabetes widely prevalent in industrialized countries are on the rise in developing countries. The burden of T2DM is enormous when the costs of diagnosis and treatment are considered. Due to economic constraints, providing modern medical healthcare in developing countries such as India is still a far-reaching goal. Out of an estimated 250000 higher plants, less than 1% has been screened pharmacologically and very few in regard to DM. Therefore, it is prudent to look for options in herbal medicine for diabetes as well. The goals of medicine no matter to which group it belongs, are the same i.e. the welfare of the patient. One can look towards a future of integrated medicine and hope that research in alternative medicine will help identify what is safe and effective rather than marginalizing, unorthodox medical claims and findings.

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