"HERBAL FORMULATIONS FOR MANAGEMENT OF DIABETES MELLITUS"

Herbal products: Herbal products comprise of herbal formulations made from one or more herbs.

Herbal medications: Ayurvedic herbal medications are defined as extracts from combination of herbs, single herbs, ayurvedic proprietary medicines or a compound of herbs in any form (tablets, capsules, powder, crude extract or decoction). The various part of herbs like roots, leaves, stems, barks are utilised for the preparation of the herbal medications.

Herbal formulations are the basis for finished herbal products and may include milled or pulvarised herbal materials, or extracts, tinctures and fatty oils of herbal materials.² Herbal preparations are produced by extraction, fractionation, purification or concentration or by other physical or biological processes.³

Herbal materials include, in addition to herbs, fresh juices, gums, fixed oils, essential oils, resins and dry powders of herbs.⁴

Herbs

Herbs include crude plant material such as leaves, flowers, fruit, seed, stems, wood, bark, roots, rhizomes or other plant parts, which may be entire, fragmented or powdered.⁵

Diabetes mellitus

Diabetes is a chronic, metabolic disease indicated by elevated levels of blood glucose (or blood sugar), which leads over time to serious damage to the heart, blood vessels, eyes, kidneys and nerves.

Diabetes is an illness where the insulin processing functions aberrantly due to pancreas not being able to secrete the hormone in adequate amounts. This in turn leads to uncontrollable levels of glucose or sugars in blood circulation and affects energy metabolism and kidney function in the affected individual.⁶

Types of diabetes

Type 1 diabetes mellitus (T1D) is an autoimmune disease that leads to the destruction of insulin-producing pancreatic beta cells. ⁷

Type 2 Diabetes Mellitus (T2DM) is one of the most common metabolic illness worldwide and its development is primarily caused by a combination of two main factors: faulty insulin release by pancreatic β -cells and the incapability of insulin-sensitive tissues to respond to insulin ⁸

Gestational diabetes

Gestational diabetes (GD) is a type of diabetes that develops in gestation when blood sugar magnitude gets too high. Gestational diabetes usually appears during the middle of pregnancy, between 24 and 28 weeks. Developing Gestational diabetes doesn't mean you already had diabetes before you got pregnant. This condition in fact appears because of the high blood sugar elevation during pregnancy.

Prediabetes

A condition in which blood sugar level has spiked, but is not sufficiently high enough to be type 2 diabetes. Without intervention, it's likely to lead to type 2 diabetes.

Diabetes mellitus can be managed by maintaining a healthy diet, avoiding junk food, following good sleep patterns, regular exercises etc.

Nowadays, diabetes mellitus can be managed by including nutraceutical substances in the diet.

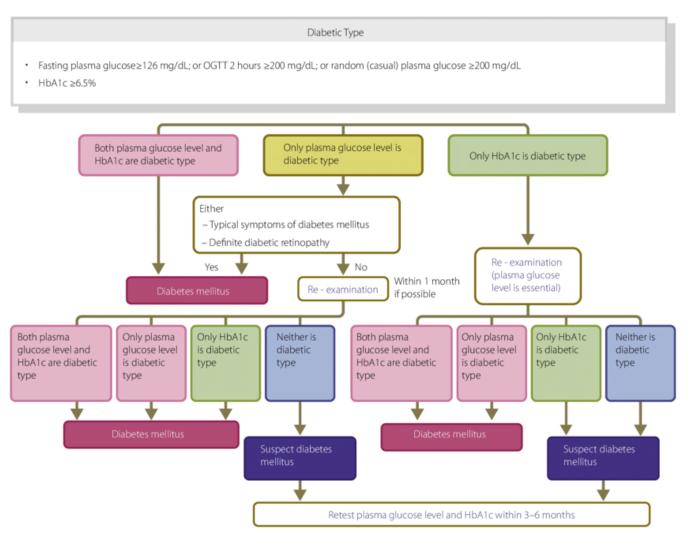
Some of these nutraceutical substances are sugar apple, curry leaves, amla, banana stem, bitter gourd etc.

How is diabetes diagnosed?¹⁴

• The detection of diabetes mellitus should be as extensive as possible. It is verified by the presence of chronic hyperactive glycemia, and by the presence of other factors, such as associated symptoms, clinical laboratory findings, a family history of diabetes, and body weight history of the patient. ⁹⁻¹³

For the diagnosis of diabetes, either of the following approaches shall be followed:

- Two assessments of the diabetic type in each patient (one blood glucose test is obligatory).
- One assessment of the diabetic type (with obligatory blood glucose testing) along with the presence of chronic hyperactive glycemic symptoms*. (*typical symptoms of chronic hyperglycemia (e.g., dry mouth, polyposis, polyuria, body weight loss, or diabetic retinopathy).
- Substantiation of a prior diagnosis of 'diabetes'.



Flow chart outlining the steps in the clinical diagnosis of diabetes mellitus. OGTT, Oral glucose tolerance test. 14

How is hyper-glycemia assessed?

- The cases are to be classified into the normal type, borderline type, or diabetic type, according to the combination of fasting and 2-h post-75 g oral glucose tolerance test (OGTT) glucose values.
- The cases where fasting glucose values are 100–109 mg/dL (5.6–6.1 mmol/L) are classified into the 'high normal' category as part of the normal type¹⁵.
- The OGTT is to be proactively considered in high-risk individuals (i.e., those who are suspected to have diabetes or the borderline type, those whose fasting glucose values are indicated to be 'high-normal', those with HbA1c values of \geq 5.6%, cases with obesity or dyslipidemia, and those with strong family history of diabetes¹³).
- At present, HbA1c values are calculated by point-of-care testing (POCT) and devices are not to be used for the diagnosis ¹³.

GOALS AND STRATEGIES FOR DIABETES MANAGEMENT

What are the objectives of diabetes management?

• The objectives of diabetes management are to ameliorate metabolic dysfunctions rising from hyperglycemia, to arrest the evolution or advancement of diabetic problems and conditions associated with diabetes, and to enable affected persons to maintain their quality of life (QOL) and life probability similar to those of healthy individuals.

How is a basic treatment strategy developed for each diabetic patient?

- The treatment strategy for diabetes may vary depending on the type, complaint condition, age, metabolic abnormality, and status of diabetic complications.
- Insulin remedy is to be given not only to cases who are insulin-dependent but also to pregnant cases, cases witnessing surgery that involves whole- body operation, and cases with severe infection, indeed if they aren't insulin-dependent. In addition, insulin remedy is to be given to those in whom glycemic pretensions aren't attainable with oral hypo- glycemic agents (OHAs) or glucagon- such as peptide 1 (GLP-1) receptor agonists.
- OHA and/ or GLP- 1 agonist remedy is to be given to noninsulin-dependent cases in whom favourable glycemic control isn't attainable with acceptable medical nutrition therapy (MNT) and physical exertion/ exercise continued for 2-3 months. OHA and/ or GLP- 1 agonist remedy or insulin remedy may be given to these cases from the onset depending on the inflexibility of the metabolic complaint involved.
- Continued remedy is essential for cases with diabetes to help the onset or progression of complications. Team care- grounded diabetes education for these cases forms the foundation of their diabetes treatment.

MEDICAL NUTRITION THERAPY (MNT)

What is the part of MNT and the optimal nutritive balance in MNT for patients with diabetes?

- Carbohydrates, proteins and fats should account for 50-60%, $\le 20\%$, and 20-30%, respectively, of the total energy intake in MNT for patients with diabetes.
- The ratios of macronutrients may vary depending on physical activities and the severity of diabetic complications, as well as on the food preferences of each patient with diabetes. 14

· Applicable to most cases of type 2 diabetes · Without acute metabolic disorder Casual plasma glucose levels approximately 250 ~ 300 mg/dL or lower · Negative urinary ketone bodies The target for glycemic control is established for each patient by the physician-in-charge, taking into account the patient's age and the condition. Start of treatment (First consultation) · Education of patients regarding diet therapy, exercise therapy, improvement of lifestyle Achievement of target for Glycemic control glycemic control target not achieved · Education of patients regarding diet therapy, exercise therapy, improvement of lifestyle · Treatment with oral hypoglycemic agents Insulin treatment · Treatment with GLP-1 receptor agonist Achievement of target for Glycemic control glycemic control target not achieved • Education of patients regarding diet therapy, exercise therapy, improvement

of lifestyle

insulin

Treatment of patients in a non-insulin-dependent state. This provides a guide to the management of patients without acute metabolic disorder¹⁴

combination with a GLP-1 receptor agonist

· Intensive insulin treatment

Achievement of target for

glycemic control

Treatment with increased dose of oral hypoglycemic agent, or their combination
Switch to insulin or to an oral hypoglycemic agent in combination treatment with

· Switch to a GLP-1 receptor agonist or an oral hypoglycemic agent or insulin in

Glycemic control

target not achieved

· Education of patients regarding diet therapy, exercise therapy, improvement

Sl no	Scientific name	Common name	Family	Parts utilized	Activity	Ref no
1	Annona squamosa Linn	Custard apple	Annonaceae	Leaves, fruit	Leaf extracts show antidiabetic, cytotoxic, antimalarial, and immunosuppressive activities.	16
2	Murraya koenigii Linn	Curry leaves	Rutaceae	Leaves	Boost insulin activity	17
3	Momordica charantia Linn	Bitter gourd	Cucurbitaceae	Fruit, vegetable	Mimics insulin and aids in glucose utilization	19
4	Phyllanthus emblica Linn	Amla	Phyllanthaceae	Fruits, leaves	stimulates pancreatic insulin secretion, and restores and regenerates β-cells	20
5	Musa sapientum Linn.	Banana	Musaceae	Plant stem, fruit, flower	induced blood sugar reduction may be due to possible inhibition of free radicals and subsequent inhibition of tissue damage induced by alloxan	18
6	Punica granatum Linn.	Pomegranate	Lythraceae	Fruit, leaves	action on β-cells of pancreas that stimulates insulin release and decreases glucose unacceptance.	21
7	Beta vulgaris Linn.	Beetroot	Amaranthaceae	Tap root	It is a good source of fibre which helps to increase satiety and slow down the absorption of carbohydrates preventing sudden spikes in blood sugar levels	

8	Withania somnifera Linn.	Ashwagandha	Solanaceae	Root	Reduce blood sugar to normal levels by boosting insulin synthesis and sensitivity	23
9	Acacia Arabica Linn.	Babul	Fabaceae	Bark, pulp	slow down any glucose absorption and support a keto diet by keeping insulin spikes at bay	24
10	Allium cepa Linn.	Onion	Amaryllidaceae	Bulb	improve insulin sensitivity and regulate blood glucose level	25
11	Allium sativum Linn.	Garlic	Amaryllidaceae	Bulb	Lower blood glucose level, stimulate insulin activity	26
12	Aloe vera (L.) Burm.f.	Aloe vera	Asphodelaceae	leaves	lowers blood glucose levels in diabetic patients. improves the responsiveness of the body tissues towards insulin	27
13	Syzygium cumini Linn	Jamun	Myrtaceae	Leaves, fruits, seeds	slow down the rate of sugar released into the blood and increase the insulin levels in the body	28
14	Butea monosperma (Lam.) Taub.	Palasa	Fabaceae	Flowers, leaves	It might reduce blood sugar levels by improving glucose tolerance and regulation of excessive blood sugar levels	29
15	Bombax diffusa	semul	Malvaceae	Flowers	lower the hyperlipidemic problems associated with diabetes mellitus., enhance the insulin release	30

16	Areca catechu Linn	supari	Arecaceae	Nut	Reduce the blood sugar level	26
17	Artemisia pallens Wall. ex DC.	Davana	Asteraceae	Leaves and flowers	significant blood glucose lowering effect	26
18	Trigonella foenum graceum Linn	Fenugreek	Fabaceae	Seeds and leaves	activating of insulin synthesis and promote its release from the pancreatic β -cells	31
19	Gymnema sylvestre R. Br.	Gurmar	Apocynaceae	Leaves	delaying the glucose absorption in the blood.	32
20	Chamaecostus cuspidatus (Nees & Mart.) C.Specht & D.W.Stev.	Insulin plant	Costaceae	Whole plant	Reduce the blood sugar level	33
21	Abroma agustum Linn	Abroma agusta	Steruliaceae	Root	Hypoglycemic and antilipidemic effect	34
22	Acer saccharum Marshall	Sugar Maple	Sapindaceae	Juice	Hhypoglycemic and antioxidant effect	35
23	Coccinia indica (L.) Voigt	Ivy gourd	Cucurbitaceae	Leaves	Antioxidant, antihyperglycemic and antilipidemic	36

24	Acalypha wilkesiana Müll.Arg.	Copperleaf	Euphorpiaceae	Leaves	Hypoglycemic and antioxidant	37
25	Acorus calamus Linn.	Sweet flag	Acoraceae	Leaves/bark	α-Glucosidase inhibition and hypolipidemic	38
26	Aegle marmelos (L.) Corrêa	Bael	Rutaceae	Leaves/bark/ro ot	Antidiabetic, Antioxidant and anticataract activity	39
27	Afzelia Africana Sm.	African mahogany	Fabaceae	Stem bark	Antidiabetic and hematological effect	40
28	Eriobotrya japonica (Thunb.) Lindl.	Japanese Loquat	Rosaceae	Leaves	Antihyperglycemic	41
29	Prosopis glandulosa Torr.	Honey mesquite	Fabaceae	Leaves	Increase insulin secretion	42
30	Amaranthus spinosus Linn.	Spine amaranth	Amaranthaceae	Leaves	α -Amylase inhibition, antioxidant and anticholesrolemic	43
31	Anogeissus acuminate (Van Heurck & Müll.Arg.)	Button tree	Combretaceae	Whole plant	Antioxidant	44
	Gere & Boatwr (terminalia phillyreifolia)					

32	Artemisia Linn.	Artemisia sphaerocephala krasch	Compositae	Gum/seed powder	Antidiabetic	45
33	Artemisia dracunculus Linn.	Tarragon	Asteraceae	Aerial part	Antidiabetic	46
34	Artemisia herba-alba Asso.	White wormwood	Asteraceae	Whole plant	Antihyperglycemic and antihyperlipidemic	47
35	Lonicera japonica Thunb.	Japanese honeysuckle	Caprifoliaceae	Whole plant	α-Glucosidase inhibition	48
36	Ascophyllum nodosum Linn.	Rockweed	Algae	Young tips	α -Glucosidase inhibition and α -amylase	49
37	Aspidosperma macrocarpon Mart.	Guatambu	Apocynaceae	Steam/Bark Root wood	α -Amylase inhibition	50
38	Atractylodes macrocephala Koidz.	Bai Zhu	Asteraceae	Whole plant	α-Glucosidase inhibition	48
39	Azadirachta indica A.Juss.	Neem	Meliaceae	Leaves, flower and seed	Antidiabetic and antihypertension	51,5 2
40	Barringtonia racemose Linn.	Common Putat	Lecythidaceae	Seed	α -Glucosidase and α -amylase inhibition	53

41	Bauhinia forficate	Brazilian orchid tree	Leguminosae	Leaves	Antidiabetic	26
42	Belamcanda chinensis Linn.	Blackberry lily	Irdaceae	Leaves	Hypoglycemic	54
43	Benincasa hispida (Thunb.)	Wax gourd	Cucurbitaceae	Fruit	Antidiabetic, ameliorate the derangement in lipid metabolism in diabetics.	55
44	Nervilia plicata	Pleated Leaf Nervillia	Orchiaceae	Stem	Antidiabetic	56
45	Berberis vulgaris Linn.	Barberry	Berberidaceae	Leaves	Hypoglycemic	57
46	Psoralea corlifolia Linn.	Purple Fleabane	Fabaceae	Leaves	Antioxidant	39
47	Rauwolfia serpentinum Linn.	Indian snakeroot	Apocyanaceae	Leaves	Hypoglycemic	58
48	Silybum marianum Linn.	milk thistle	Asteraceae	Whole plant	Antihyperglycemic	59
49	Argyreia cuneate Ker Gawl.	purple morning glory	Convolvulaceae	Whole plant	Antihyperglycemic	60
50	Desmodium gangeticum Linn.	salparni	Fabaceae	Whole plant	Antidiabetic	61

These are commonly available from the plants, in the market etc which can be used directly, or can be processed at home, and can be used in form of decoction, powder, etc

1. Annona squamosa Linn. 16



Synonym: Sweetsop, Custard Apple

Biological source: It comprises of dried leaves of *Annona squamosa* Linn. belonging to family Annonaceae

Phytochemical constituents: anonaine, annonacin, squamocin, asimicin, phytochemical assessments have emphasised that numerous active compounds, such as acetogenins and flavonoids.

Description: Annona squamosa Linn. is a small tree, which is well branched belonging to the family Annonaceae that bears edible fruits called sugar-apples or sweetsops. It tolerates a tropical flat climate and it is the most widely cultivated of its species. Annona squamosa Linn. is a small, semi-deciduous, branched shrub or small tree 3 to 8 metres (10 to 26 feet) tall.

Formulation: decoction

- Decoction preparation
- Chop the leaves into small pieces
- boil the leaves in water for about 5-10 min
- consume the freshly prepared decoction
- 2. Murraya koenigii Linn. 17



Synonym: karivepaaku, karibevu kadi patta

Biological source: It comprises of dried leaves of Murraya koenigii Linn belonging to family Rutaceae

Phytochemical constituents: alkaloids, glycosides, steroids, tannins and terpenoids monoterpenes. linalool, elemol, geranyl acetate, myrcene, α-terpinene (E)-β-ocimene.

Description: It's a small tree, growing 4–6 metres (13–20 ft) in height, with the trunk up to 40 cm (16 in) in width. The leaves are pinnate, aromatic, with 11–21 leaflets, each leaflet 2–4 cm ($\frac{3}{4}$ –1+ $\frac{1}{2}$ in) long and 1–2 cm ($\frac{1}{2}$ – $\frac{3}{4}$ in) broad

Consumption:

- the leaves can be directly be consumed orally
- the leaves can be included in diet in the dishes
- the leaves can be dried, crushed into powder and can be consumed with water.
- 3. Momordica charantia Linn. 19



Synonym: bitter apple, bitter gourd, karela,

Biological source: It comprises of the fruits and leaves of *Momordica charantia* Linn. belonging to family Cucurbitaceae

Phytochemical constituents: polysaccharides, saponins, and phenolics cucurbitane-type triterpenoids, cucurbitane-type triterpene glycosides, phenolic acids, flavonoids, essential oils, fatty acids, amino acids, sterols, saponins and protein

Description: The tree is medium in size, reaching 1-8 m (3 ft 3 in -26 ft 3 in) in height. The branches are finely pubescent (not glabrous), 10-20 cm (3.9–7.9 in) long, usually deciduous. The leaves are simple, subsessile and are present close along branchlets, light green in colour, exhibit pinnate leaves. The flowers are greenish-yellow. The fruit is nearly spherical, light greenish-yellow, fairly smooth and hard on appearance.

Juice preparation:

- Take a fresh bitter gourd, chop into pieces
- Combine the bitter gourd and half cup of water in a mixer
- Prepare a blend
- Strain by adding little water
- And consume immediately

4. Phyllanthus emblica Linn.²⁰



Synonym: Indian gooseberry, āmalakī, emblic myrobalan

Biological source: It comprises of the fruits and leaves of *Phyllanthus emblica* Linn. belonging to the family phyllanthaceae.

Phytochemical constituents: ellagic acid, emblicanine A, emblacani B, Gallic acid, phyllantine, quercetin and phyllantidine

Description: the tree is medium in size, reaching 1-8 m (3 ft 3 in -26 ft 3 in) in height. The branches are finely pubescent (not glabrous), 10-20 cm (3.9–7.9 in) long, usually deciduous. The leaves are simple, subsessile and set nearly along the branches, resembling pinnate leaves, and are light green in appearance. The flowers are greenish-yellow. The fruit is nearly globular, light greenish-yellow, quite smooth and hard on appearance, with six vertical stripes or furrows

Consumption

Eat the fruit fresh:

This is the greatest and the efficient method for the consumption of amla

Amla juice:

- first remove the seed
- crush the pulp
- extract the juice by blending and straining with water

Amla powder:

- Take fresh amla fruits
- Cut into pieces
- Sun-dry the amla pulp
- grind it into a fine powder.

5. Musa sapientum Linn. 18



Synonym: Common banana

Biological source: It comprises of the fruit, stem of Musa sapientum Linn. belonging to family

Musaceae

Phytochemical constituents: phenolics, carotenoids, biogenic amines and phytosterols

Description: The banana plant is among the largest blossomy flowering plant. All aerial parts of the plant grow from a structure usually called a "corm". Plants normally have a good height and are fairly sturdy with a treelike appearance, but what appears to be the trunk is actually a "false stem" or pseudo-stem. Bananas grow in a wide variety of soils, as long as the soil is at least 60 centimetres (2.0 ft) deep, has good water supply and isn't compacted. Banana plants are one of the rapidly growing of all plants, with diurnal surface growth rates recorded of 1.4 square metres (15 sq ft) to 1.6 square metres (17 sq ft).

Consumption

- Blend chopped banana stem with water
- Untill it becomes a smooth paste.
- Add some plant-based buttermilk and salt.
- Mix well and drink it immediately.
- 6. Punica granatum Linn.²¹



Synonym: Malum granatum, Punica granatum, anar

Biological source: It comprises of fruits and leaves of *Punica granatum* Linn belonging to family Lythraceae

Phytochemical constituents: Punicalagin, Ellagic acid, Cyanidin. Liteolin, organic acids, phenolic acids, sterols, terpenoids, fatty acids, triglycerides, alkaloids and some other compounds.

Description: Punica granatum Linn. is a shrub growing 5 to 10 m (16 to 33 ft) in height, and has multiple spiny branches and is long-lived, with some species in France have survived for 200 years. Punica granatum Linn. leaves are opposite or subopposite, glossy, narrow oblong, entire, 3–7 cm $(1+\frac{1}{4}-2+\frac{3}{4}$ in) long and 2 cm $(\frac{3}{4}$ in) broad. The flowers are bright red and 3 cm $(1+\frac{1}{4}$ in) in diameter, with three to seven petals. Some fruitless species of Punica granatum Linn. are grown only for obtaining the flowers.

Consumption:

Pomegranate tea:

- Add 3 mugs of pomegranate seeds to a blender.
- Store the juice in a vessel after straining.
- Combine three-fourth mug of piping hot water with one-quarter mug of pomegranate juice.

7. Beta vulgaris Linn.²²



Synonym: Beetroot, common beet, or garden beet

Biological source: It comprises of taproot portion of a *Beta vulgaris* Linn. belonging to family Amaranthaceae

Phytochemical constituents: betalains (e.g., betacyanins and betaxanthins), flavonoids, polyphenols, Saponins and inorganic Nitrate (NO₃)

Description: Beets, *Beta vulgaris* Linn. are blossoming root vegetables in the family Chenopodiaceae grown for their consumable root. The plant leaves are seen growing on stems and is usually upright with a long main root. The leaves are arranged alternately on the stem, ovoid shaped and grow 20–40 cm (7.9–15.7 in) in length. The roots are usually red in colour. The plant produces stalkless green flowers that can grow 1–2 m (3.3–6.6 ft) in height. Beets are usually grown as yearly plants, harvested after one growing season. Beets may also be referred to as beetroot, garden beet or spinach beet and originated from the Mediterranean.

Juice preparation:

- Take a fresh beetroot, chop into pieces
- Combine the beetroot and half cup of water in a mixer
- Prepare a blend
- Strain by adding little water
- And consume immediately

8. Withania somnifera Linn²³





Synonym: Ashwagandha or winter cherry,

Biological source: Ashwagandha is derived from the root of the *Withania somnifera* Linn plants belonging to family Solanaceae

Phytochemical constituents: steroidal lactones, alkaloids, saponin, flavonoids, tannin, starch, phenolic content, carbohydrate, withanolides, sitoindosides, anaferine, anahygrine, β -sitosterol, chlorogenic acid, cysteine, cuscohygrine, pseudotropine, withanine, scopoletin

Description: This species is a 35–75 cm (14–30 in) tall growing shrub. The branches expand spirally from the central stem. Leaves appear to be dull green, elliptic, usually up to 10–12 cm (3.9–4.7 in) long. The flowers are small in size, green coloured and bell-shaped. The fruit becomes orange-red on ripening.

Usage

- Take one glass of milk and upto its half volume water in a pan and bring it to boil.
- Introduce Ashwagandha root powder, upto 1/2 teaspoon and boil for additional 5 minutes.
- To the admixture add around (2 teaspoons) crushed walnut and almonds
- Drink this admixture to maintain blood sugar level.

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