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Website: [www.thepharmaresearch.info](http://www.thepharmaresearch.info)  
E-ISSN 0975-8216 (Online)

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Editor-In-Chief,  
The Pharma Research  
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### An overview of Diabetes, A Review

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#### **Abstract**

Diabetes mellitus (DM) is the commonest endocrine disorder that affects more than 100 million people worldwide (6% of the population). It is caused by the deficiency or ineffective production of insulin by pancreas which results in increase or decrease in concentrations of glucose in the blood. It is found to damage many of the body systems, particularly the blood vessels and nerves. Many herbal plants with hypoglycemic properties are known from across the world. In India, diabetes has been known for a long time, but its incidence is not of the same magnitude across the subcontinent. The wide range of structures of the plant constituents, which appear to be the active hypoglycemic principles, suggests different sites of action within the body. Whether these plants truly possess hypoglycemic properties needs to be investigated for those plants that are commonly used in the management of diabetes. Researches conducted in the last few decades on plants, mentioned in ancient literature or used traditionally for diabetes, have shown anti diabetic property. Among them, 30 plants and their products (active natural principles and crude extracts) that have been mentioned used in the Indian traditional system of medicine have shown experimental or clinical anti diabetic activity. *Trigonella foenumgraecum*, *Momordica charantia*, *Tinospora cordifolia*, *Encostema littorae*, *Gymnema sylvestre*, *Azadiracta indica*, *Syzygium cumini* are some of the most effective and the most commonly studied Indian plants in relation to diabetes.

**Keyword:** *Trigonella foenumgraecum*, *Momordica charantia*, *Tinospora cordifolia*, *Encostema littorae*, *Gymnema sylvestre*, *Azadiracta indica*, *Syzygium cumini*, Diabetic

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#### **Introduction**

Diabetes, the word came from Greek, means to siphon. The most common sign was excessive urination. Mellitus, the word came from Latin, means honey sweet. Diabetes mellitus (DM) is a metabolic disorder characterized by hyperglycemia (high blood glucose) and other signs, as distinct from a single illness or condition (Diagnosis and Classification, 2004). It is a chronic metabolic disease with inability

to maintain blood glucose concentrations within physiological limits. It develops when the pancreas does not produce enough insulin or when the body cannot effectively uses the insulin it produces.

#### **Classification of Diabetes Mellitus** **Type-1: Autoimmune, Insulin Dependent Diabetes Mellitus (IDDM).**

It is a catabolic disorder in which circulating insulin is

virtually absent due to beta cell destruction.

#### **Type-2: Non-autoimmune, Non-Insulin Dependent Diabetes Mellitus (NIDDM)**

Circulating endogenous insulin is sufficient but is often either sub-normal or relatively inadequate because of tissue insensitivity.

#### **Type-3: Secondary Diabetes**

1. Genetic
2. Insulin receptor abnormality
3. Pancreatic disease
4. Hormonal abnormality
5. Drug and Chemical Induced Diabetes mellitus

#### **Type-4: Gestational Diabetes Mellitus**

Some women develop intolerance during pregnancy and it is characterized by excessive hunger, thirst and frequent urge to urinate.

#### **Sign and Symptoms**

The major indications of diabetes include excessive thirst and frequent urination (polyuria) which compels to drink a large volume of water (polydipsia), glycosuria, excessive hunger (polyphagia), and loss of weight and lack of energy (asthenia) in case of Type-1 mostly. But Type-2 is different in diagnosis as the symptoms are very vague and unnoticeable. Symptoms of IDDM rise promptly while in NIDDM take a year or more than a year (Memon Badaruddin A., et al. 2002). Dangerous symptoms include smell of acetone on patient's breathing, rapid deep breathing (kussmaul breathing) and altered state of consciousness or arousal which may lead to diabetic coma (Akhtar J., et al. 2005).

#### **Factors Responsible for Diabetes**

Major causative factors include genetic susceptibility, autoimmunity, environmental factors, mental stress and strain, obesity, excessive sleep, high carbohydrate diet, high blood pressure, low activity

level and poor diet (Memon Badaruddin A., et al. 2002) (Patel D., et al. 2009).

#### **The Risks and Complications of Uncontrolled Diabetes**

If not controlled, diabetes can put you at risk for a host of complications that can affect nearly every organ in the body. They include: the heart and blood vessels, eyes, kidneys, nerves, gums and teeth etc. Heart disease and blood vessel disease are the biggest complications that people with uncontrolled diabetes face. Approximately 65% of death from diabetes is due to heart disease and stroke. Diabetes can also cause poor blood flow in the legs and feet (peripheral artery disease). Diabetes cause a number of eye problems include: Glaucoma, Cataracts and Diabetic retinopathy.

#### **Herbal Treatment for Diabetes**

Although, many drugs and intervention are available to manage diabetes, in most cases they are expensive, produce serious side effects, and in addition they are not considered to be safe for use during pregnancy. The use of herbal medicines for the treatment of diabetes mellitus has gained importance throughout the world. The W.H.O. also recommended and encouraged this practice especially in countries where access to the conventional treatment of diabetes is not adequate (Trivedi, P. C. et al. 2006).

Medicinal plants are being looked up once again for the treatment of diabetes. Many conventional drugs have been derived from prototypic molecules in medicinal plants. Metformin exemplifies an efficacious oral glucose-lowering agent. Its development was based on the use of *Galega officinalis* to treat diabetes. *Galega officinalis* is rich in guanidine, the hypoglycemic component. Because guanidine is too toxic for clinical use, the alkyl biguanides synthalin A and synthalin B were introduced as oral anti-diabetic agents in Europe in the 1920s but were discontinued after insulin became more widely available. However,

experience with guanidine and biguanides prompted the development of metformin. To date, over 400 traditional plant treatments for diabetes have been reported, although only a small number of these have received scientific and medical evaluation to assess their efficacy. Major hindrance in amalgamation of herbal medicine in modern medical practices is lack of scientific and

clinical data proving their efficacy and safety. There is a need for conducting clinical research in herbal drugs, developing simple bioassays for biological standardization, pharmacological and toxicological evaluation, and developing various animal models for toxicity and safety evaluation. It is also important to establish the active component/s from these plant extracts (Devasagayam P. A. et al. 2007).

**Table 1: List of Plants with Putative Anti-Diabetic Effects**

Plant (family)	Part	Model	References
St. John's Wort	leaves	streptozotocin-diabetic rats	Özgür Devrim Can et al.(2011)
<i>Pyrus bioisieriana</i> Buhse	leaves	Alloxan-rat	Mohammad ES et al.(2011)
coconut			
<i>Aegle marmelos</i> (Rutaceae)	protein	Alloxan-rat	G. Salil a,c, K.G. Nevina,b, T. Rajamohana,?(2011)
	Leaves	Alloxan-rat	Ponnachan et al. (1993)
		STZ-rat	Narendhirakannan et al. (2006)
		STZ-rat	Seema et al. (1996)
		STZ-rat	Kesari et al. (2006)
<i>Allium cepa</i> (Liliaceae)		STZ-rat	Kamalakannan et al. (2005)
<i>Allium sativum</i> (Liliaceae)		STZ-rat	Jain et al. (1974)
<i>Amaranthus esculantus</i> (Amaranthaceae)	Seeds	Alloxan-rabbit	Jain et al. (1975)
<i>Annona squamosa</i> (Annonaceae)	Fruits	Alloxan-rabbit	Kim et al. (2006)
	Bulbs	STZ-rat	
	Bulbs		
	Whole plant		Gupta et al. (2005); Kaleem et al. (2006)
		STZ-rat	Gupta et al. (2005)
	Leaves	Alloxan-rat	
<i>Averrhoa bilimbi</i> (Oxalidaceae)	Fruit pulp	STZ-rat, Alloxan-rabbit	Tan et al. (2005)
<i>Baccharis trimera</i> (Myrtaceae)		STZ-rat	
<i>Bryophyllum pinatum</i> (Crassulaceae)	Leaves	STZ-mice	Oliveira et al. (2005)
<i>Canarium schweinfurthii</i> (Burseraceae)	Leaves	STZ-rat	Ojewole (2005)
<i>Catharanthus roseus</i> (Apocynaceae)	Leaves	STZ-rat	Kamtchouing et al. (2005)
<i>Chamaemelum nobile</i> (Asteraceae)	Stem barks	STZ-rat	Singh et al. (2001)
<i>Coscinium fenestratum</i> (Menispermaceae)			Eddouks et al. (2005)
<i>Egyptian Morus alba</i> (Moraceae)	Whole plant	STZ-rat	
<i>Eugenia jambolana</i> (Asteraceae)	Leaves	STZ-rat	Shirwaikar et al. (2005)
<i>Hintonia standleyana</i> (Rubiaceae)	Stem barks	STZ-rat	Singap et al. (2005)
<i>Hypoxis hemerocallidea</i> (Hypoxidaceae)			Sharma et al. (2006); Ravi et al. (2005)
<i>Leonotis leonurus</i> (Lamiaceae)	Stem barks	STZ-rabbit	Guerrero-Analco et al. (2005); Navarette et al. (2005)
<i>Lepidium sativum</i> (Brassicaceae)	Fruit pulp, seeds	STZ-rat	Ojewole (2006)
<i>Lycium barbarum</i> (Solanaceae)	Stem barks		
<i>Malmea depressa</i> (Annonaceae)		STZ-mice, rat	Ojewole (2005)
<i>Mangifera indica</i> (Anacardiaceae)	Fruits	STZ-mice, rat	

<i>Momordica charantia</i> (Cucurbitaceae)	Leaves	STZ-rat	Eddouks et al. (2005)
	Leaves	STZ-rat	Zhao et al. (2005); Wu et al. (2006) Andrade-Cetto et al. (2005)
<i>Murraya koenigii</i> (Rutaceae)	Fruits	STZ-rat	
	Roots	Alloxan-rat STZ-rat	Aderibigbe et al. (1999) Ojewole (2005)
<i>Ocimum sanctum</i> (Labiatae)	Leaves	STZ-rat	Sekar et al. (2005)
	Stem barks	Alloxan-rat	Reyes et al. (2006)
<i>Piper betle</i> (Piperaceae)	Leaves	STZ-mice	Sitasawad et al. (2000)
	Leaves	Alloxan-rat STZ-rat	Yadav et al. (2005) Satishsekar et al. (2005)
<i>Psidium guajava</i> (Myrtaceae)	Fruits	Alloxan-rat and STZ-rat	Yadav et al. (2002)
	Seeds	Alloxan-rabbit STZ-rat	Watal et al. (2005) Watal et al. (2007)
<i>Raphanus sativa</i> (Brassicaceae)	Leaves	STZ-rat	Chattopadhyay et al. (1993)
<i>Retama raetum</i> (Fabaceae)		STZ-rat	Arambewela et al. (2005); Santhakumari et al. (2006)
<i>Salvia officinalis</i> (Lamiaceae)		STZ-rat	Ojewole (2005)
<i>Scoparia dulcis</i> (Scrophulariaceae)	Leaves	STZ-rat	
<i>Strobilanthes crispus</i> (Acanthaceae)	Leaves	STZ-at	Taniguchi et al. (2006)
<i>Syzygium cordatum</i> (Myrtaceae)	Leaves	STZ-rat	Maghrani et al. (2005)
<i>Syzygium cumini</i> (Caesalpiniaceae)	Leaves	STZ-mice, rat	Lima et al. (2006)
<i>Taxus yunnanensis</i> (Taxaceae)	Whole plant	STZ-rat	Pari et al. (2005)
<i>Terminalia chebula</i> (Combretaceae)	Whole plant	STZ-rat	Fadzelly et al. (2006)
<i>Terminalia superba</i> (Combretaceae)	Leaves	STZ-rat	
<i>Trema orientalis</i> (Ulmaceae)	Whole plant	STZ-rat	Musabayane et al. (2005)
<i>Tremella mesenterica</i> (Combretaceae)	Leaves	STZ-mice, rat	Oliveira et al. (2005)
<i>Viscum album</i> (Loranthaceae)	Leaves	STZ-rat	Banskota et al. (2006)
<i>Zizyphus spina-christi</i> (Rhamnaceae)	Leaves	STZ-rat	Rao et al. (2006)
	Leaves, seeds Woods	STZ-rat	Kamtchouing et al. (2005)
	Seeds	STZ-rat	Dimo et al. (2006)
	Stem barks	STZ-rat	
	Stem barks	STZ-rat	Lo et al. (2006)
	Fruits	STZ-rat	
	Whole plant	STZ-rat	Orhan et al. (2005)
	Leaves	STZ-rat	Abdel-Zaher et al. (2005)

**Table: List of Plants with Putative Anti-Diabetic Effects**

Plant (family)	Part	Model	References
<i>Aegle marmelos</i> (Rutaceae)	Leaves	Alloxan-rat STZ-rat	Ponnachan et al. (1993) Narendhirakannan et al. (2006)
<i>Allium cepa</i> (Liliaceae)		STZ-rat	Seema et al. (1996)
<i>Allium sativum</i> (Liliaceae)		STZ-rat	Kesari et al. (2006)
<i>Amaranthus esculantus</i> (Amaranthaceae)	Seeds	Alloxan-rabbit	Kamalakannan et al. (2005)
	Fruits	Alloxan-rabbit	Jain et al. (1974)
	Bulbs	STZ-rat	Jain et al. (1975)
	Bulbs		Kim et al. (2006)

<i>Annona squamosa</i> (Annonaceae)	Whole plant	STZ-rat	Gupta et al. (2005); Kaleem et al. (2006)
	Leaves	Alloxan-rat	Gupta et al. (2005)
	Fruit pulp	STZ-rat, Alloxan-rabbit STZ-rat	Tan et al. (2005)
<i>Averrhoa bilimbi</i> (Oxalidaceae)	Leaves	STZ-mice	Oliveira et al. (2005)
<i>Baccharis trimera</i> (Myrtaceae)	Leaves	STZ-rat	Ojewole (2005)
<i>Bryophyllum pinatum</i> (Crassulaceae)	Leaves	STZ-rat	Kamtchouing et al. (2005)
<i>Canarium schweinfurthii</i> (Burseraceae)	Leaves	STZ-rat	Singh et al. (2001)
<i>Catharanthus roseus</i> (Apocynaceae)	Leaves	STZ-rat	Eddouks et al. (2005)
<i>Chamaemelum nobile</i> (Asteraceae)	Stem barks	STZ-rat	Singh et al. (2001)
<i>Coscinium fenestratum</i> (Menispermaceae)	Whole plant	STZ-rat	Eddouks et al. (2005)
<i>Egyptian Morus alba</i> (Moraceae)	Leaves	STZ-rat	Shirwaikar et al. (2005)
<i>Eugenia jambolana</i> (Myrtaceae)	Stem barks	STZ-rat	Singap et al. (2005)
<i>Hintonia standleyana</i> (Rubiaceae)	Stem barks	STZ-rabbit	Sharma et al. (2006); Ravi et al. (2005)
<i>Hypoxis hemerocallidea</i> (Hypoxidaceae)	Fruit pulp, seeds	STZ-rat	Guerrero-Analco et al. (2005); Navarette et al. (2005)
<i>Leonotis leonurus</i> (Lamiaceae)	Stem barks	STZ-rat	Ojewole (2006)
<i>Lepidium sativum</i> (Brassicaceae)	Fruits	STZ-mice, rat	Ojewole (2005)
<i>Lycium barbarum</i> (Solanaceae)	Leaves	STZ-mice, rat	Ojewole (2005)
<i>Malmea depressa</i> (Annonaceae)	Leaves	STZ-rat	Eddouks et al. (2005)
<i>Mangifera indica</i> (Anacardiaceae)	Leaves	STZ-rat	Zhao et al. (2005); Wu et al. (2006)
<i>Momordica charantia</i> (Cucurbitaceae)	Fruits	STZ-rat	Andrade-Cetto et al. (2005)
<i>Murraya koenigii</i> (Rutaceae)	Stem barks	STZ-rat	Aderibigbe et al. (1999)
	Fruits	STZ-mice, rat	Ojewole (2005)
	Leaves	STZ-mice, rat	Ojewole (2005)
<i>Ocimum sanctum</i> (Labiatae)	Leaves	STZ-rat	Eddouks et al. (2005)
<i>Piper betle</i> (Piperaceae)	Leaves	STZ-rat	Zhao et al. (2005); Wu et al. (2006)
<i>Psidium guajava</i> (Myrtaceae)	Fruits	STZ-rat	Andrade-Cetto et al. (2005)
<i>Raphanus sativa</i> (Brassicaceae)	Fruits	STZ-rat	Andrade-Cetto et al. (2005)
<i>Retama raetum</i> (Fabaceae)	Roots	STZ-rat	Aderibigbe et al. (1999)
<i>Salvia officinalis</i> (Lamiaceae)	Leaves	Alloxan-rat	Ojewole (2005)
<i>Scoparia dulcis</i> (Scrophulariaceae)	Stem barks	STZ-rat	Sekar et al. (2005)
<i>Strobilanthes crispus</i> (Acanthaceae)	Leaves	STZ-rat	Reyes et al. (2006)
<i>Syzygium cordatum</i> (Myrtaceae)	Leaves	STZ-rat	Sitasawad et al. (2000)
<i>Syzygium cumini</i> (Caesalpiniaceae)	Leaves	Alloxan-rat	Yadav et al. (2005)
<i>Taxus yunnanensis</i> (Taxaceae)	Fruits	STZ-mice	Satishsekar et al. (2005)
<i>Terminalia chebula</i> (Combretaceae)	Fruits	Alloxan-rat	Yadav et al. (2002)
<i>Terminalia superba</i> (Combretaceae)	Seeds	STZ-rat	Watal et al. (2005)
<i>Trema orientalis</i> (Ulmaceae)	Leaves	Alloxan-rat and STZ-rat	Watal et al. (2007)
<i>Tremella mesenterica</i> (Combretaceae)	Leaves	Alloxan-rabbit	Chattopadhyay et al. (1993)
<i>Viscum album</i> (Loranthaceae)	Leaves	STZ-rat	Chattopadhyay et al. (1993)
<i>Zizyphus spina-christi</i> (Rhamnaceae)	Leaves	STZ-rat	Arambewela et al. (2005); Santhakumari et al. (2006)
	Leaves	STZ-rat	Ojewole (2005)
	Leaves	STZ-rat	Taniguchi et al. (2006)
	Leaves	STZ-at	Maghrani et al. (2005)
	Whole plant	STZ-rat	Lima et al. (2006)
	Whole plant	STZ-mice, rat	Pari et al. (2005)
	Whole Leaves	STZ-rat	Fadzelly et al. (2006)
	Whole plant	STZ-rat	Musabayane et al. (2005)
	Whole plant	STZ-rat	Musabayane et al. (2005)

	Leaves	STZ-mice, rat	Oliveira et al. (2005)
	Leaves	STZ-rat	Banskota et al. (2006)
	Leaves, seeds	STZ-rat	Rao et al. (2006)
	Woods	STZ-rat	Kamtchouing et al. (2005)
	Seeds	STZ-rat	Dimo et al. (2006)
	Stem barks	STZ-rat	Lo et al. (2006)
	Stem barks	STZ-rat	Orhan et al. (2005)
	Fruits	STZ-rat	Abdel-Zaher et al. (2005) Federica Menichini et al. (2011)
	Whole plant		
	Leaves		

### Advantages of Herbal Medicine

1. Lower costs than conventional medicine.
2. Reduced side effects compared to conventional treatments.
3. Often aim to remove underlying cause rather than treat symptoms.
4. Holistic approach promoting good overall health.
5. Potential to reduce the burden on conventional medicine.
6. Growing research in activities and improvements in production (Mukherjee, P. K., et al. 2002).

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