Double-blind, placebo-controlled clinical evaluation of an Ayurvedic formulation (GlucoCare capsules) in non-insulin dependent diabetes mellitus

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ABSTRACT

Diabetes mellitus describes a metabolic disorder of multiple etiologies characterized by insulin resistance, relative insulin deficiency and hyperglycemia with disturbances of carbohydrate, fat and protein metabolism. The goal for treatment of diabetes is to prevent its acute manifestations and long-term microvascular and macrovascular complications. The present study was conducted to evaluate the efficacy and safety of an Ayurvedic formulation (GlucoCare Capsules) in non-insulin dependent diabetes mellitus. Fifty NIDDM patients of pitta-kapha prakriti attending the outpatient department of the Government Ayurvedic Medical College, Guwahati, Assam, India were included in the study, and randomly divided into 2 groups, GlucoCare and placebo. All received either GlucoCare or placebo in a dose of 2 capsules twice daily, before meals for 3 months. All 50 patients completed the study - no drop outs, withdrawals or patients lost to follow up. The GlucoCare group showed significant improvement in symptoms from the 2nd month till the end of the study. GlucoCare was well tolerated by all patients throughout the treatment period with no evidence of adverse effects. The study indicates clinical efficacy of GlucoCare Capsules in the management of NIDDM in those belonging to pitta-kapha prakriti. The formulation is well tolerated and appears safe in the dosage used.

Key words: Ayurvedic formulation, Ayurvedic formulation, non-Insulin dependent diabetes mellitus.

INTRODUCTION

In many countries, Non-Insulin Dependent Diabetes Mellitus (NIDDM) is one of the most prevalent and fastest growing diseases. Family physicians play a central role in its management. Though recognized as a distinct clinical syndrome for centuries, our understanding of the disease, its causation and mechanism of progression, are still evolving. In 2004, India had an estimated 37.76 million diabetics; 21.4 million in urban areas and 16.36 million in rural areas. The Chennai Epidemiological study (CURES) found percentages with diabetic retinopathy, microalbuminuria and peripheral neuropathy to be 17.6%, 26.9% and 26.1% respectively. Although many drugs improve glycemic control, they do not necessarily provide real-world benefits. In the recent ACCORD (Action to Control Cardiovascular Risk in Diabetes) and ADVANCE (Action in Diabetes and Vascular Disease: Preterax and Diamicron MR Controlled Evaluation) trials, intensive glycemic control had minimal effect on clinical cardiovascular outcomes. In fact, in a recent meta-analysis, combination therapy with metformin and glyburide increased the risk of a composite end-point of cardiovascular events and mortality (relative risk 1.43, 95% confidence interval (CI) 1.10 to 1.85). Use of thiazolidinediones has recently been called into question because of increased risk of cardiovascular events and fracture. Compliance with medically advised medication is far from ideal. Cost of therapy and adverse effects of medication have been prime factors for inadequate patient compliance.

More than one-third of the world population uses complementary and alternative medicine (CAM) therapies, often without consulting or even informing their family physicians (FPs). It is important for FPs to ask patients about their CAM use and provide evidence-based information about the safety and efficacy of commonly used CAM therapies. 
For diabetes, oral medication is usually the first line treatment after diet and exercise, which aim at management of acute conditions and prevention of long term complications. Exercise, diet and weight control continue to be essential and effective means of improving glucose homeostasis. However, lifestyle management measures may be insufficient or patient compliance difficult, rendering conventional drug therapies (i.e., oral glucose-lowering agents and insulin injection) necessary in many patients. In addition to adverse effects, drug treatments are not always satisfactory in maintaining euglycemia and avoiding late stage diabetic complications.

In Ayurveda, diabetes mellitus is named Madhumeha, a type of Prameha. Charaka and Sushruta classify 20 varieties of ‘Meha’ under ‘Vata’ ‘Pitta’ and ‘Kapha’ categories. Madhumeha includes three clinical types, Vataja, Pittaja and Kaphaja. Kaphaja Madhumeha may be related to high growth hormone levels, the pittaja type in particular to high levels of glucocorticoids. In the progression of diabetes, the initial stage of Prameha, Kapha dosha is at a high level later changing to Kapha Kshaya. Similarly Pittaja Prameha leads to Pitta Kshaya. However, in both Kaphaja and Pittaja Prameha Vata Vriddhi is a common denominator which in its terminal stage leads to Madhumeha: subtype 1 of Vataja Prameha. Specific treatments prescribed in Ayurvedic texts include herbs, detox therapies and dietetics. In recent times, the herbs have been subjected to scientific scrutiny and analysis.

Medicinal herbs with antihyperglycemic activities are increasingly sought after by diabetic patients and health care professionals. Commonly used herbs and other alternative therapies, are considered less likely to have the side effects of conventional treatments for NIDDM. Alternative therapies for diabetes have been extensively researched, particularly in India. Here, an Ayurvedic formulation GlucoCare capsules (manufactured by The Himalaya Drug Company, Bangalore, India) using a combination of herbs, is evaluated for use in NIDDM. Its composition is given in Table 1.

### Aim of the study
The present study was conducted to evaluate the efficacy and safety of an Ayurvedic formulation (GlucoCare Capsules) in non-insulin dependent diabetes mellitus.

### MATERIALS AND METHODS

#### Study design
Fifty NIDDM outpatients attending the Government Ayurvedic Medical College, Guwahati, Assam, India were randomly divided into GlucoCare and placebo groups. Patients underwent clinical examination on entry and at monthly intervals. They were given diaries to note incidence and severity of symptoms. Adverse effects if any were recorded. The study protocol was approved by the hospital’s and institute’s Ethics Committee. If they so desired, patients were free to withdraw from the study, which was conducted in accordance with the Helsinki Declaration, and the Government of India’s Ministry of Health GCP Guidelines. Informed written consent was obtained from all study participants.

A questionnaire and clinician’s observations and confirmation were used for Prakriti analysis. Primary prakriti analysis was made based on a set of simple questionnaire and the clinician’s observations and confirmation.

#### Inclusion criteria
Patients of Pitta-Kapha prakriti, aged 30 to 65, male or female, diagnosed with NIDDM characterized by plasma glucose concentration >200 mg/dL or fasting plasma glucose (FPG) >126 mg/dL or 2 hr post prandial glucose (PPG) >200 mg, positive urine sugar and having symptoms of polyuria, polyphagia and polydipsia, and willing to give written informed consent.

#### Exclusion criteria
Insulin-dependent diabetes mellitus (IDDM – Type I), with acute complications (nephropathy, neuropathy, retinopathy and gangrene), pregnant or lactating women, malignant

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**Table 1: Composition of GlucoCare capsule**

| Name of the active ingredients | Quantity per capsules (mg) |
|-------------------------------|-----------------------------|
| Exts.                         |                             |
| Commiphora wightii            | 7.5                         |
| Shilajeet (Purified)          | 7.5                         |
| Gymnema sylvestre             | 35                          |
| Pterocarpus marsupium         | 30                          |
| Glycyrrhiza glabra            | 20                          |
| Casearia esculenta            | 25                          |
| Syzygium cumini               | 23                          |
| Asparagus racemosus           | 20                          |
| Boerhaavia diffusa            | 20                          |
| Sphaeranthus indicus          | 10                          |
| Tinospora cordifolia          | 10                          |
| Tribulus terrestris           | 10                          |
| Phyllanthus amarus            | 10                          |
| Gmelina arborea               | 10                          |
| Gossypium herbaceum           | 10                          |
| Aloe vera                     | 5                           |
| Triphala                      | 8                           |
| Momordica charantia           | 10                          |
| Piper nigrum                  | 2.5                         |
| Ocimum sanctum                | 2.5                         |
| Abutilon indicum              | 2.5                         |
| Curcuma longa                 | 2.5                         |
| Rumex maritimus               | 2                           |
| Tribkat                       | 2                           |
hypertension, history of severe unstable angina, myocardial infarction, cardiovascular accidents, renal failure, or allergy to medications, or not willing to give written informed consent.

**Study procedures**
Fifty patients of NIDDM attending the out-patient department of Government Ayurvedic Medical College, Guwahati, Assam, India were included in the study and were divided arbitrarily randomized into two groups, i.e., GlucoCare group and placebo group. All the patients received either GlucoCare or placebo in a dose of two capsules twice daily, before meals for three months in a random manner. Patients underwent clinical examination on entry and at monthly intervals for three months. They were provided with a diary to note down the incidence and severity of symptoms. Adverse effects if any were recorded. The prakriti other than Pitta-Kapha was rejected.

**Follow-up and assessment**
All subjects underwent clinical examination and evaluation of blood sugar levels on entry and at monthly intervals for the 3 month study period. At each monthly visit, subject evaluations were based on symptoms, fasting plasma glucose (FPG) and post-prandial glucose (PPG).

**Primary and secondary outcome measure**
The primary end-point was symptomatic relief, reduction and control of NIDDM symptoms, polydipsia, polyuria, polyphagia, burning sensation in hands and soles, pain/cramps, fasting and postprandial blood sugar levels. GlucoCare safety and toxicity profiles were secondary end points.

**Statistical analysis**
All the values are expressed as Mean ± SD or incidence of symptoms. Statistical analysis was done by using Student’s t-test or ANOVA. The minimum level of significance was fixed at P < 0.05. Statistical analysis was performed using GraphPad Prism software (Version 4.01).

**RESULTS**
The two groups’ baseline characteristics on entry were comparable [Table 2]. Mean ages were 50.40 ± 6.50 years in the GlucoCare group and 53.50 ± 8.60 years in the placebo group. Male : female ratios were 15:10 (GlucoCare) and 16:9 (placebo).

All 50 patients completed the study. Patients on GlucoCare group showed significant symptomatic improvement from 1st month to the end of the study [Table 3].

Results indicate that from 2nd month onwards GlucoCare subjects significantly decreased polydipsia, and continued to improve until the end of the study. Subjects on placebo did not respond significantly.

Polyuria observed in 14 patients on entry decreased to 12 patients after one month, and 1 patient by the end of study. Placebos showed no significant improvement in polyuria.

Similar trends were seen for other symptoms: polyphagia, burning sensation in hands/feet, and pain/cramps. By the 2nd month, most Glucocare subjects showed overall improvement.

The GlucoCare group showed significant reductions in FBS from 180 ± 10.5 mg% to 130 ± 5.5 mg%, and PPBS from 200 ± 22.5 mg% to 140 ± 13.5 mg%. The placebo group showed no significant reductions in either [Table 3].

End of study evaluations assessed overall response to treatment. In the GlucoCare group, all subjects said they had excellent response to treatment, 76% were symptom-free [Table 4].

**Adverse effects**
GlucoCare capsules were well tolerated by all GlucoCare subjects throughout the treatment period. No serious hematological or biochemical abnormalities were experienced by any subject. There was no evidence of adverse side-effects.

**DISCUSSION**
Despite understanding NIDDM etiopathogenesis, rises in insulin-resistant cases and failure of oral hypoglycemic
agents (OHAs) are alarming. The past two decades have seen explosive increases in numbers diagnosed with NIDDM globally. In India, an estimated 19.4 million are affected, a figure likely to increase to 57.2 million by 2025. Even today, India has the largest number of diabetics of any country in the world. In the 1970s, diabetes prevalence among urban Indians was 2.1%, a figure now standing at 12.1%.

Moreover, an equally large pool of individuals with impaired glucose tolerance (IGT) are at risk of developing NIDDM in the near future.

The prakriti of the two groups were matched (Pitta-Kapha) so their drug response was comparable. The present study indicates the efficacy of the Ayurvedic formulation in controlling fasting and postprandial blood sugar of Diabetic patients belonging to Pitta-Kapha prakriti. No significant benefit in glycosylated hemoglobin values was found, however. All subjects tolerated the drug, none withdrew. To evaluate safety and efficacy more accurately, future studies will need larger sample sizes.

Insulin lowers plasma glucose levels by stimulating glucose uptake into muscle and inhibiting hepatic glycogen breakdown. Catecholamines cause hyperglycaemia by stimulating hepatic glycogenolysis and inhibiting insulin stimulated glucose utilisation in muscle. GlucoCare has been found to inhibit catecholamine induced hyperglycaemia and significantly improve liver glycogen storage. It has also been found to increase incorporation of C - 14 glucose in liver slices in alloxan-induced diabetes in rats.\(^9\) Now, a more specific analysis of GlucoCare ingredients: Gymnema sylvestre, (Mesasringi) one of the important ingredients of GlucoCare, has been found to be effective in diabetes by increasing beta cell function possibly by repair/regeneration of the beta cells.\(^{10,11}\) Momordica charantia (Karavallaka) seeds were found to contain molecules with insulin-like bioactivity.\(^{11}\) It is possible that GlucoCare reduced blood sugar levels by improving the plasma insulin, increasing peripheral utilization of glucose, improving liver glycogen storage, and by its intrinsic antidiabetic action.

In both insulin-dependent diabetes (IDDM) and non-insulin dependent diabetes (NIDDM), morbidity and mortality from cardiovascular disease is greatly increased, possibly due to increased serum lipid. Furthermore, there is considerable evidence that control of serum lipids results in the reduction of the incidence of coronary heart disease. It is therefore important to understand the effects of the treatments used in diabetes on serum lipids and lipoproteins.\(^{12}\) In this combination of GlucoCare, Commiphora wightii (Guggulu) plays a complementary role by renormalizing the serum lipids and cholesterol possibly

### Table 3: Effect of drug therapy on symptoms of non-Insulin dependent diabetes mellitus

| Parameter (No. of patients with) | GlucoCare (n = 25) | Placebo (n = 25) |
|----------------------------------|--------------------|-----------------|
|                                  | At entry | 1st month | 2nd month | End of study | At entry | 1st month | 2nd month | End of study |
| Polydipsia                        | 17       | 15        | 9*        | 2*          | 19       | 19        | 15        | 8           |
| Polyuria                          | 14       | 12        | 10*       | 1*          | 16       | 15        | 12        | 10          |
| Polyphagia                        | 16       | 12        | 7*        | 2*          | 18       | 17        | 15        | 9           |
| Burning sensation in hands and soles | 10 | 7        | 4*        | 2*          | 12       | 11        | 10        | 7           |
| Pain/cramps                       | 9        | 6         | 3*        | 1*          | 11       | 10        | 10        | 6           |
| Fasting blood sugar (Mean ± SD)   | 180 ± 10.5 | 150 ± 16.5 | 140 ± 13.5 | 130 ± 15.5 | 180 ± 12.5 | 170 ± 10.5 | 160 ± 19.5 | 150 ± 18.5 |
| Post-prandial blood sugar (Mean ± SD) | 200 ± 22.5 | 190 ± 18.5 | 160 ± 15.5 | 140 ± 13.5 | 210 ± 20.5 | 190 ± 18.5 | 180 ± 15.5 | 160 ± 17.5 |
| Glycosylated hemoglobin (%)       | 6.35 ± 1.42 | -        | -         | 6.0 ± 2.5  | 7.0 ± 1.0 | -         | -         | 6.0 ± 1.5  |

*P < 0.05 as compared to respective ‘At entry’ values

### Table 4: Overall response to treatment at 3rd month

| Parameters                  | GlucoCare (n = 25) | Placebo (n = 25) |
|-----------------------------|--------------------|-----------------|
|                             | No. of patients    | Response (%)    | No. of patients | Response (%) |
| Symptom-free                | 19/25              | 76              | 7/25            | 28           |
| Patient's impression        |                    |                 |                |
| Excellent                   | 19/25              | 76              | 3/25           | 12           |
| Good                        | 4/25               | 16              | 1/25           | 4            |
| No response                 | 2/25               | 8               | 21/25          | 84           |


due to its androgen (AR), glucocorticoid (GR) antagonistic activities that potentially aids the person suffering from diabetes.\textsuperscript{[13]} Shilajit is a pale-brown to blackish-brown exudation, of variable consistency, exuding from layers of rocks in many mountain ranges of the world, especially the Himalayas. It has been used as a rejuvenator and an adaptogen for thousands of years, in one form or another, as part of traditional systems of medicine for treatment of diabetes.\textsuperscript{[14]}

Among crude components extracted from the leaves of \textit{Gymnema sylvestre} is one of the triterpene saponins that suppress sweetness by a reversible effect on the sweet taste receptors. Pharmacological tests also show reduction in blood sugar. The extract suppresses increases in blood glucose by inhibiting its reuptake in the intestines.\textsuperscript{[4]} It also provides effective hyperglycemic control, which is crucially important to prevent the micro- and macrovascular complications of diabetes mellitus\textsuperscript{[15]} and enhances peripheral utilization of glucose.\textsuperscript{[9]}

\textit{Pterocarpus marsupium} (Asana) can control diabetic-related metabolic alterations apart from controlling glucose levels.\textsuperscript{[16]} Its hypoglycemic action may be due to its reduced glucose absorption from the gastrointestinal tract.\textsuperscript{[17]} A clinical trial has shown that oral intake of \textit{P. marsupium} extract has potent hypoglycemic activity (both fasting and postprandial) that can be comparable with tobutamide.\textsuperscript{[18]}

Licorice flavonoid oil (LFO) from \textit{Glycyrrhiza glabra} (Yashti) contains Hydrophobic flavonoids with abdominal fat-lowering and hypoglycemic effects in obese animal diabetic models. Mediation by PPAR-gamma activation has been suggested.\textsuperscript{[19]}

\textit{Casearia esculenta}, (Chilhaka bheda) is an indigenous plant popularly used as an antidiabetic in South India. Oral administration of aqueous extracts lowers blood glucose levels under normal and glucose load conditions, possibly due to inhibition of blood glucose absorption from the gut\textsuperscript{[20]} and the presence of potent anti-hyperglycemic factor(s).\textsuperscript{[21]} It also is a potent antioxidant.

Seeds of \textit{Syzygium cumini} (Jambu) show preferred hypoglycemic activity, establishing its positive pharmacological activity.\textsuperscript{[22]}

Extracts of \textit{Asparagus racemosus} (Satavari) root contain large amounts of flavonoids, polyphenols and vitamin-C, and exhibit extreme antioxidant activity. They scavenge free radicals such as peroxide, hydroxyl radical, hydrogen peroxide and nitric oxide, all of which participate in various pathophysiologies. They can thus play an important role in NIDDM.\textsuperscript{[28]}

Treating normal and diabetic rats with extracts of \textit{Boerhaavia diffusa} (Punarnava) resulted in significant decreases in blood glucose and increases in plasma insulin levels, establishing their hypoglycemic action; also observed were significant reductions in glycosylated hemoglobin and increases in total hemoglobin.\textsuperscript{[24]}

\textit{Sphaeranthus indicus} (Munditika) is a noted hypoglycemic herb.\textsuperscript{[25]} Fasting normal rats treated with its alcoholic extract significantly improved on oral glucose tolerance tests, suggesting an application to diabetes mellitus.\textsuperscript{[26]}

\textit{Tinospora cordifolia} (Guduchi) shows hypoglycemic activity possibly by stimulating endogenous insulin secretion by altering the cell membrane permeability.\textsuperscript{[27,28]} It also has aldose reductase inhibitory actions, thus preventing cataract formation.\textsuperscript{[29]}

A trial has shown that extracts of \textit{Tribulus terrestris} (Gokshura) significantly decrease fasting glucose levels. They also cause significant decrease in glycosylated hemoglobin levels, and total cholesterol, triglycerides and LDL-cholesterol.\textsuperscript{[30]}

\textit{Fruits of Phyllanthus amarus} (Tamalaki) show hypoglycemic activity as does hydro-alcoholic extract of \textit{Gmelina arborea}.\textsuperscript{[34]}

\textit{Gossypium herbaceum} (Karpasa) has therapeutic action against elevated blood sugar, cholesterol and triglyceride, also exhibiting an ability to reduce blood sugar.\textsuperscript{[35]}

\textit{Aloe vera} is noted for hypoglycemic activity and used in diabetes mellitus.\textsuperscript{[38]} Some studies have indicated that long-term \textit{Aloe vera} (Kumari) gel treatment helps diabetic hyperglycemia. Administration of gel with a high fat diet prevents development of insulin resistance and glucose intolerance.\textsuperscript{[36]} Oral intake prevents progression of NIDDM-related symptoms.\textsuperscript{[37]}

'Triphala (equal proportion of \textit{Terminalia chebula}, \textit{Terminalia belerica}, and \textit{Emblica officinalis}) shows hypoglycemic activity.\textsuperscript{[39]}

It has been reported that isolates and extracts of \textit{Momordica charantia} contain a hypoglycaemic principle - an insulin-like peptide called foetidin, momordicin or charantin (polypeptide p-insulin). According to some investigators, this glucoside (polypeptide p-insulin) is useful for management of both IDDM and NIDDM. Furthermore, it is possible that the plant extract mimics or improves insulin action at the cellular level, or even exhibits extra-pancreatic action.\textsuperscript{[40]}

Extracts of \textit{Momordica charantia} enhance insulin secretion by the islets of Langerhans, reducing glyco genesis in liver tissue, enhancing peripheral glucose utilization and increasing serum protein levels.\textsuperscript{[10]}

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Piper nigrum (Marica) seeds have a hypoglycemic effect, and also check antioxidant levels. This is necessary and sufficient for control of complications arising from glycation and glycoxidation of proteins and membranes.[41]

Ocimum sanctum (Tulasi) leaf extracts greatly stimulate insulin secretion from the β-cells via physiological pathways.[42] They prevent cataract in rat lenses by simultaneously restoring antioxidant defenses and inhibiting protein precipitation. Concurrent administration of Vitamin-E helps reverse changes in diabetic retinopathy.[43] Alh Siddu indicum (Aitabla) leaves show hypoglycemic activity.[44] The plant also has good wound healing activity, and is helpful in management of wounds and other superficial dermatological infections in diabetes mellitus.[45]

Curcumin, the yellow phenolic curcuminoid present in turmeric, has been reported to have a wide range of biological activities. Even very small dietary levels (0.002%) delay galactose induced cataract in rats.[46] It exhibits very high lipid-soluble antioxidant action, and may be helpful in diabetes.[47]

Curcuma longa root also contains Rumarin, used for its anti-pruritic activity, and may be helpful in reducing itching sensations in those with hyperglycemia.[48] Its additional neuroprotective activity may be helpful in managing other neurological problems seen in diabetes.[49]

Trikatu is a phyto-combination (Piper longum, Piper nigrum and Zingiber officinale) that increases bioavailability by promoting rapid absorption from the gastrointestinal tract, or preventing metabolism/oxidation during first passage through the liver after being absorbed, or a combination of these mechanisms, helping improve most drugs’ therapeutic activity.[50]

This study demonstrates GlucoCare’s efficacy in a small population of a particular Prakriti. Further studies should include a larger population of patients, including all Prakritis.

**CONCLUSION**

Results of this clinical study of 50 patients of pitta-kapha prakriti suffering from NIDDM, indicate that GlucoCare Capsules have beneficial effects in relieving symptoms and bringing about overall improvement. 19 of the 25 patients on GlucoCare felt that they showed adequate response to treatment. The present clinical study indicates significant clinical efficacy of GlucoCare Capsules in management of NIDDM. The formulation was well tolerated and appeared to be safe in the dosage used.

**DECLARATION**

The authors declare that the WHO Criteria for Medicinal Drug Promotion (1986) were meticulously followed during this clinical study and its publication.

**REFERENCES**

1. Gerstein HC, Miller ME, Byington RP, Goff DC, Jr., Bigger JT, Buse JB, et al. Effects of intensive glucose lowering in type 2 diabetes. N Engl J Med. 2008; 358(24):2545-59.

2. IDF - Diabetes Atlas, 3rd Edition, International Diabetes Federation, Brussels, 2007.

3. IDF - Diabetes Atlas, 3rd Edition, International Diabetes Federation, Brussels, 2007.

4. Rao AD, Kuhadiya N, Reynolds K, Fonseca VA. Is the combination of sulfonfonylureas and metformin associated with an increased cardiovascular disease or all-cause mortality? A meta-analysis of observational studies. Diabetes Care 2008;31:1672-8.

5. Lipscotme LL. Thiazolidinediones: do harms outweigh benefits? CMAJ 2009;180:16-7.

6. Upadhyay OP, Upadhyay D. A few facts of historical interest relating to diabetes mellitus. Indian J History Sci 1987;22:235-9.

7. Shimizu K, Iino A, Nakajima J, Tanaka K, Nakajyo S, Urakawa N, et al. Suppression of glucose absorption by some fractions extracted from Gymnema sylvestre leaves. J Vet Med Sci 1997;59:245-51.

8. Shastry J, Prakash A, Prasad R. Hypoglycaemic activity of a leaf extract from Gymnema sylvestre in non-insulin-dependent diabetes mellitus patients. J Ethnopharmacol 1996;52:207-12.

9. Mitta SK, Gopumadhavan S, Muralidhar TS, Seshadri SJ. Effect of D-400, a herbomineral formulation on liver glycocontent and microscopic structure of pancreas and liver in streptozotocin induced diabetes in rats. Indian J Exp Biol 1996;34:964-7.

10. Vasu V, Grover JK, Rathi SS. Evaluation of anti-hyperglycemic and hypoglycemic effect of Trigonella foenum-graecum Linn, Ocimum sanctum Linn and Pterocarpus marsupium Linn, in normal and alloxanized diabetic rats. J Ethnopharmacol 2002;79:95-100.
Sharma and Patki: GlucoCare capsules in non-insulin dependent diabetes mellitus

18. Harirahan RS, Venkataraman S, Sunitha P, Rajalakshmi S, Samal KC, Routray BM, et al. Efficacy of Vijayasara (Pterocarpus marsupium) in the treatment of newly diagnosed patients with Type-2 diabetes mellitus: a flexible dose double blind multicenter randomized controlled trial. Diabetologia Croatica 2005;34:13-20.

19. Nakagawa K, Kishida H, Arai N, Nishiyama T, Mae T, Licorice Flavonoids Suppress Abdominal Fat Accumulation and Increase in Blood Glucose Level in Obese Diabetic KK-Ay Mice. Biol Pharm Bull 2004;27:1775-8.

20. Prakash A, Sethupathy S, Pugalendi KV. Effect of Casearia esculenta root extract on blood glucose and plasma antioxidants status in streptozotocin diabetic rats. Pol J Pharmacol 2003;55:43-9.

21. Prakash A, Sethupathy S, Pugalendi KV. Antioxidative and antioxidant effects of Casearia esculenta root extract in streptozotocin-induced diabetic rats. Yale J Biol Med 2005;78:15-23.

22. Singh N, Gupta M. Effects of ethanolic extract of Syzygium cumini (Linn) seed powder on pancreatic islets of alloxan diabetic rats. Indian J Exp Biol 2007;45:861-7.

23. Velavan S, Nagulendran K, Mahesh R, Hazeena Begum V. In vitro antioxidant activity of Asparagus racemosus root. Phcog Mag 2007;3:26-33.

24. Pari L, Amarnath Satheesh M. Antidiabetic activity of Boerhaavia diffusa L.: effect on hepatic key enzymes in experimental diabetes. J Ethnopharmacol 2004;91:109-13.

25. Jeychandran R, Mahesh A. Enumeration of antidiabetic herbal flora of Tamil Nadu. Res J Medicinal Plant 2007;1:144-8.

26. Prabhuj, Lobo R, Shirwaikar A. Antidiabetic properties of the alcoholic extract of Sphaeranthus indicus in streptozotocin-nicotinamide diabetic rats. J Pharm Pharmacol 2008;60:909-16.

27. Stanely Mainzen Prince P, Menon VP. Hypoglycaemic and hypolipidaemic action of alcohol extract of Tinospora cordifolia roots in chemical induced diabetes in rats. Phytother Res 2003;17:410-3.

28. Prince PS, Menon VP. Antioxidant activity of Tinospora cordifolia roots in experimental diabetes. J Ethnopharmacol 1989;56:277-81.

29. Rathi SS, Grover JK, Vikrant V, Biswas NR. Prevention of experimental diabetic cataract by Indian Ayurvedic plant extracts. Phytother Res 2002;16:774-7.

30. El-Tantawy WH, Hassanin LA. Hypoglycemic and hypolipidemic effects of alcoholic extract of Tribulus alatus in streptozotocin-induced diabetic rats: a comparative study with T. terrestris (Caltrop). Indian J Exp Biol 2007;45:785-90.

31. Islam A, Mazumder UK, Gupta M, Ghosal S. Phytopharmacology of Phyllanthus amarus: an overview. Pharmacognonline 2008;3:202-9.

32. Raphael KR, Sabu MC, Kuttan R. Hypoglycemic effect of methanol extract of Phyllanthus amarus Schum and Thonn on alloxan induced diabetes mellitus in rats and its relation with antioxidant potential. Indian J Exp Biol 2002;40:905-9.

33. Srividya N, Periwal S. Diuretic, hypotensive and hypoglycaemic effect of Phyllanthus amarus. Indian J Exp Biol 1995;33:861-4.

34. Asolkar LV, Kakkar KK, Chakre OJ. Gmelina arborea: glossary of Indian Medicinal plants with Active principles. CSIR Publication, New Delhi, Second supplement, Part-I (A-K), 1992:335.

35. Mitra SK, Gopumadhavan S, Muralidhar TS, Anturlikar SD, Sujatha MB. Effect of a herbomineral preparation D-400 in streptozotocin-induced diabetic rats. J Ethnopharmacol 1996;54:41-6.

36. Tanaka M, Misawa E, Ito Y, Habara N, Nomaguchi K, Yamada M, et al. Identification of five phytosterols from Aloe vera gel as anti-diabetic compounds. Bioll Pharm Bull 2008;29:1418-22.

37. Kim K, Kim H, Kwon J, Lee S, Kong H, Im SA, et al. Hypoglycemic and hypolipidemic effects of processed Aloe vera gel in a mouse model of non-insulin-dependent diabetes mellitus. Phytomedicine 2009;16:1-8 (In Press)

38. Lanks CA. Ethnomedicines used in Trinidad and Tobago for urinary problems and diabetes mellitus. J Ethnobiol Ethnomed 2006;2:45.

39. Sabu MC, Kuttan R. Anti-diabetic activity of medicinal plants and its relationship with their antioxidant property. J Ethnopharmacol 2002;81:155-60.

40. Ojewole JA, Adewole SO, Oluyiwola G. Hypoglycaemic and hypotensive effects of Momordica charantia Linn (Cucurbitaceae) whole-plant aqueous extract in rats. Cardiovasc J S Afr 2006;17:227-32.

41. Kaleem M, Sheema, Sarmad H, Bano B. Protective effects of Piper nigrum and Vinca Rosea in alloxan induced diabetic rats. Indian J Physiol Pharmacol 2005;49:65-71.

42. Hannan JM, Marenah L, Ali L, Rokeya B, Flatt PR, Abdel-Wahab YH. Ocimum sanctum leaf extracts stimulate insulin secretion from perfused pancreas, isolated islets and clonal pancreatic-cells. J Endocrinol 2006;189:127-36.

43. Eshrat EM, Mukhopadhyay AK. Effect of Ocimum sanctum (Tulsi) and vitamin-E on biochemical parameters and retinopathy in streptozotocin induced diabetic rats. Indian J Clin Biochem 2006;21:181-8.

44. Seetharam YN, Chalageri G, Setty SR, Bheemachar. Hypoglycemic activity of Abelitum indicum leaf extracts in rats. Fitoterapia 2002;73:156-9.

45. Roshan A, Ali S, Khan A, Tazneem B, Purohit MG. Wound healing activity of Abelitum indicum. Phcog Mag 2008;4:585-8.

46. Suryanarayana P, Saraswat M, Mridula T, Krishna TP, Krishnaswamy K, Reddy GB. Curcumin and Turmeric Delay Streptozotocin-Induced Diabetic Cataract in Rats. Invest Ophthalmo Vis Sci 2005;46:2092-9.

47. Itokawa H, Shi Q, Akiyama T, Morris-Natschke SL, Lee KH. Recent advances in the investigation of curcuminoids. Chin Med 2008;3:1-13.

48. Yoganarasimhan SN. Rumex maritimus: Medicinal Plants- Tamil Nadu. Bangalore. 1st edition, Vol. 2. 2000:471.

49. Modak M, Dixit P, Londhe J, Ghaskadbi S, Paul A, Devasagayam T. Indian herbs and herbal drugs used for the treatment of diabetes. J Clin Biochem Nutr 2007;40:163-73.

50. Atal CK, Zutshi U, Rao PG. Scientific evidence on the role of Ayurvedic herbas on bioavailability of drugs. J Ethnopharmacol 1981;4:229-32.