Volume 1 Issue 2 2012

ISSN: 2320-480X

The Journal of Phytopharmacology

(Pharmacognosy and Phytomedicine Research)

Indian medicinal plants with hypoglycemic potentials: Review Part I

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Abstract: Blood glucose lowering activity of aqueous extract in normal and glucose loaded rats. Antihyperglycemic activity of the extract in streptozotocin-diabetic rats along with reduction in the increased plasma thiobarbituric acid reactive substance and blood urea. Exhibits significant reduction in blood glucose level, a decrease in the activities of glucose-6-phosphatase and fructose-1,6-bishosphatase and an increase in the activity of liver hexokinase, resulting in potent hypoglycemic activity.

Keywords: Plant, Hypogleemia, Blood level, Glucose, Family

Introduction: Indian medicinal plants with hypoglycemic activity:

1. Acacia Arabica

Common name: Indian Gum Arabic tree

Family: Leguminosae

Activity with route of

administration/dosage:

- i. Hypoglycaemic activity of 94% seed diet in normal rats orally with no blood sugar lowering activity in alloxanized rats at the same dose level.¹
- ii. Hypoglycemic effect of powdered seeds in normal rabbits (2, 3 and 4 mg/kg) administered orally.²

Reported mechanism of action:

Acts through release of insulin from pancreatic beta cells, which accounts for the hypoglycemic activity.^{1,2}

2. Aegle marmelos (L.)

Common name: Holy Fruit Tree

Family: Rutaceae

Activity with route of administration/dosage:

- i. Hypoglycemic effect of aqueous decoction of the plant root bark (1 ml/100 mg) in normal fasted rats.³
- ii. Antihyperglycemic activity of aqueous leaf extract in alloxanized rats.⁴
- iii. Antihyperglycemic activity of aqueous leaf extract in streptozotocin induced diabetic rats. 5,6
- iv. Hypoglycemic and antioxidant activity of leaves in diabetic male albino rats.⁷
- v. Antihyperglycemic and antioxidant activity of the plant in alloxanized rats.⁸

- vi. Antihyperglycemic activity of the leaves in glucose induced hyperglycemic rat at an oral dose equivalent to 250 mg/kg.⁹
- vii. Antihyperglycemic activity of aqueous fruit extract (250 mg/kg, twice daily for 1 month) in streptozotocin induced female albino Wistar diabetic rats. 10
- viii. Hypoglycaemic activity of water extract of fruits in streptozotocin-induced diabetic Wistar rats (125 and 250 mg/kg) twice a day for 4 weeks, orally.¹¹
- ix. Antioxidant activity of the aqueous extract of fruits in streptozotocin diabetic rats (125 and 250 mg/kg), orally for 30 days. 12

Reported mechanism of action:

Increases utilization of glucose; either by direct stimulation of glucose uptake or via the mediation of enhanced insulin secretion⁹ and also decreases the elevated glucose and glycosylated hemoglobin levels.¹⁰

3. Allium cepa L.

Common name: Onion

Family: Liliaceae

Activity with route of administration/dosage:

- i. Hypoglycemic activity of ether soluble fraction of onion (0.25 mg/kg p.o.) in normal rabbits.¹³
- ii. Hypoglycemic activity of the bulb in rabbits in an oral glucose tolerance test at 2 g/kg.¹⁴
- iii. Antihyperglycemic, antioxidant and hypolipidemic activity of a diet containing 3% freeze dried onion powder upon prolonged administration in STZ diabetic rats. 15

Reported mechanism of action:

Lowers blood glucose level and has potent antioxidant activity, which may account for the hypoglycemic potential.¹³

4. Allium sativum L.

Common name: Garlic

Family: Alliaceae

Activity with route of administration/dosage:

- i. Antihyperglycemic activity of ethanol, petroleum ether and ethyl acetate extract in alloxanized rabbits at a dose of 0.25 mg/kg, orally. 16
- ii. Antioxidant activity of allicin, isolated from garlic. 17

Reported mechanism of action:

Has strong antioxidant activity and rapid reactivity with thiol containing proteins responsible for the hypoglycemic property.¹⁷

5. Aloe vera (L.)

Common name: Aloe

Family: Aloaceae

Activity with route of administration/dosage:

- i. Hypoglycemic activity of the plant (200 and 300 mg/kg p.o.) on normal fasted rats, oral glucose-loaded rats and streptozotocin-induced diabetic rats. ¹⁸
- ii. Hypoglycaemic activity of leaf pulp extracts in type I and type II diabetic rats. 19

- iii. Hypoglycemic effect of aloe and its bitter principle in alloxanized mice. ²⁰
- iv. Antihyperglycemic activity of dried sap in five non-insulin-dependent diabetic patients and in alloxanized Swiss albino mice.²¹

Reported mechanism of action:

Maintains glucose homeostasis by controlling the carbohydrate metabolizing enzymes 18 and stimulates insulin release from pancreatic beta cells.20

6. Artemisia pallens

Common name: Davana

Family: Compositae

Activity with route of administration/dosage:

i. Antihyperglycemic activity of aerial parts (100 mg/kg, orally) in glucose-fed hyperglycaemic and alloxan-induced diabetic rats.

Moderate hypoglycaemic effect (1000 mg/kg) in fasted normal rats.²²

Reported mechanism of action:

Inhibits glucose re-absorption or increase in peripheral glucose utilization.²²

7. Annona squamosa L.

Common name: Sugar apple

Family: Annonaceae

Activity with route of administration/dosage:

- i. Hypoglycemic activity of aqueous leaf extracts in streptozotocin-nicotinamide induced diabetic rats.²³
- ii. Hypoglycemic and antihyperglycemic activities of ethanolic leaf-extract (350 mg/kg, orally) in normal, streptozotocin (STZ)-diabetic rats and alloxanized rabbit.²⁴

Reported mechanism of action:

Lowers blood glucose level.²³

8. Andrographis paniculata

Common name: King of Bitter

Family: Acanthaceae

Activity with route of administration/dosage:

 Hypoglycemic and antihyperglycemic activity of Andrographis paniculata and andrographolide in normal and streptozotocin induced diabetic rats, orally.²⁵

Reported mechanism of action:

Prevents glucose absorption from gut.²⁵

9. Azadirachta indica

Common name: Neem

Family: Meliaceae

Activity with route of administration/dosage:

- i. Hypoglycemic activity of hydro alcoholic plant extract in normal rats and hypoglycemic activity in glucose fed and streptozotocin induced diabetic rats.²⁶
- ii. Hypoglycemic and antihyperglycemic activities of leaf extract in normal and streptozotocin-induced diabetic rat.²⁷
- iii. Hypoglycemic activity of crude ethanolic extract of the plant in alloxan diabetic albino rats.²⁸

Reported mechanism of action:

Inhibits action of epinephrine on glucose metabolism, resulting in increased utilization of peripheral glucose²⁶ and exhibits hypoglycaemic activity without altering the serum cortisol concentration.²⁷,

10. Biophytum sensitivum (L.)

Common name: Life Plant

Family: Oxalidaceae

Activity with route of administration/dosage:

- i. Hypoglycemic activity of the plant leaf extract in alloxan diabetic male rabbits.²⁹
- ii. Hypoglycemic activity of the plant on glucose homeostasis in rabbits.³⁰

Reported mechanism of action:

Stimulates pancreatic beta cells to release insulin.²⁹

11. Beta vulgaris L.

Common name: Garden beet

Family: Chenopodiaceae

Activity with route of administration/dosage:

i. Hypoglycemic activity of Betavulgarosides II–IV, isolated from the root of Beta vulgaris L. in an oral glucose tolerance test in rats.³¹

Reported mechanism of action:

Lowers blood glucose level.³¹

12. Brassica juncea (L.)

Common name: Brown Mustard

Family: Brassicaceae

Activity with route of administration/dosage:

i. HHypoglycemic activity of Brassica juncea diet (10%, w/w) in normal rats upon oral administration for 60 days.³²

Reported mechanism of action:

Increases the concentration of hepatic glycogen and glycogenesis and suppressed the activity of glycogen phosphorylase and gluconeogenic enzymes, lead to reduction in glycogenolysis and gluconeogenesis.³²

13. Boerhavia diffusa L.

Common name: Tar vine

Family: Nyctaginaceae

Activity with route of administration/dosage:

- i. Hypoglycemic activity of aqueous leaf extract at a dose of 100, 200 and 400 mg/kg in alloxan induced diabetic rats.³³
- ii. Hypoglycemic and antihyperglycemic activity of aqueous leaf extract (200 mg/kg p.o., daily for 4 weeks) in normal and alloxan induced diabetic rats. 34, 35

Reported mechanism of action:

Increases plasma insulin levels and improves glucosentolerance, produced significant antioxidant activity. 34, 35

14. Cassia auriculata L.

Common name: Tanner's

Family: Leguminosae

Activity with route of administration/dosage:

- i. Antihyperglycemic and antihyperlipidemic activity of aqueous flower extract in streptozotocin-induced diabetic rats upon oral administration at different doses for 30 days. 36, 37
- ii. Antioxidant activity of aqueous flower extract in the brain of streptozotocin diabetic rats. 38,39

Reported mechanism of action:

Suppresses enhanced gluconeogenesis during diabetes and enhance utilization of glucose through increased glycolysis^{36, 37} in addition to pronounced alpha-glucosidase inhibitory actions resulting in a significant and potent lowering of blood glycemic response.^{38, 39}

15. Caesalpinia bonducella (L.)

Common name: Chinese Cinnamon

Family: Caesalpiniaceae

Activity with route of administration/dosage:

- i. Hypoglycemic and antihyperglycemic activities of the aqueous and 50% ethanolic seed extracts in normal and streptozotocin-diabetic rats. 40
- ii. Antihyperglycemic activity of the seed extracts in type II diabetic Long Evans rat. 41
- iii. Hypoglycemic activity of aqueous and ethanolic extracts in chronic type II diabetic model with an increase in secretion of insulin from isolated islets.⁴²

Reported mechanism of action:

Increases the release of insulin from pancreatic cells. 40

16. Cajanus cajan (L.).

Common name: Pigeon pea

Family: Fabaceae

Activity with route of

administration/dosage:

- Glucose tolerance enhancing activity of aqueous leaf and stem extract in oral glucose tolerance test.⁴³
- ii. Hypoglycemic activity of cooked diet in healthy human volunteers. 44

Reported mechanism of action:

Lower plasma glucose level. 45

17. Citrullus colocynthis (L.)

Common name: Bitter apple

Family: Cucurbitaceae

Activity with route of administration/dosage:

- i. Hypoglycemic activity of aqueous extract (300 mg/kg), glycosidic and saponin extract (50 mg/kg), orally in normalnrabbits. 46,47
- ii. Blood glucose lowering activity of aqueous seed extract in normal and streptozotocin (STZ)-induced diabetic rats upon daily oral administration for 2 weeks. 48

Reported mechanism of action:

Exerts an insulinotropic effect. 46, 47

18. Coccinia indica

Common name: Ivy gourd

Family: Cucurbitaceae

Activity with route of administration/dosage:

- i. Hypoglycemic activity of alcoholic leaf extract in normoglycemic guinea pig. 49
- ii. Hypoglycemic activity of the leaves in alloxanized dogs upon oral administration. 50
- iii. Hypoglycemic and antihyperglycemic activity of the ethanolic root extract in fasted and glucose-loaded animal models.⁵¹
- iv. Hypoglycemic effect of 95% ethanol extract of the leaves in normal fed and 48 h starved rats. 52
- v. Blood glucose lowering activity of 60% ethanol leaf extract (200 mg/kg, orally). ⁵³

vi. Hypoglycemic activity of the leaf extract in a double blind control trial in human subjects. 54,55

vii. Antihyperglycemic activity of dried extract (500 mg/kg p.o., for 6 weeks) in 30 diabetic patients.⁵⁶

Reported mechanism of action:

Suppresses glucose synthesis, through depression of the key gluconeogenic glucose-6-phosphatase enzymes fructose-1,6-bisphosphatase and enhances glucose oxidation by shunt pathway through activation of its principal enzyme glucose-6phosphate dehydrogenase.52 Also has an insulin secretagogue effect^{54, 55} and acts like insulin by correcting elevated enzymes in glycolytic pathway and restoring LPL activity in lypolytic pathway with control of hyperglycemia in diabetes.⁵⁶

19. Casearia esculenta Roxb.

Common name: Carilla

Family: Flacourtiaceae

Activity with route of

administration/dosage:

i. Antihyperglycaemic activity of root extract (300 mg/kg p.o. for 45 days) in normal and streptozotocin-induced diabetic rats.⁵⁷

Blood glucose lowering activity ii. of aqueous extract in normal glucose loaded and rats. Antihyperglycemic activity of the extract in streptozotocindiabetic rats along with reduction in the increased plasma thiobarbituric acid reactive substance and blood urea.⁵⁸

Reported mechanism of action:

Exhibits significant reduction in blood glucose level, a decrease in the activities of glucose-6-phosphatase and fructose-1,6-bishosphatase and an increase in the activity of liver hexokinase, resulting in potent hypoglycemic activity.⁵⁷

20. Catharanthus roseus (L.)

Common name: Madagascar periwinkle

Family: Apocynaceae

Activity with route of administration/dosage:

- i. Hypoglycemic activity of ethanolic leaf extract in normal rats upon oral administration at graded dose. Hypoglycemic activity of the extract (500 mg/kg) in streptozotocin rats and in oral glucose tolerance test.⁵⁹
- ii. The hypoglycemic activity of dichloromethane: methanol nextract of leaves and twigs in streptozotocin (STZ) induced diabetic rat (500 mg/kg p.o., for 7 and 15 days). 60
- iii. Hypoglycemic and antihyperglycemic activity of leaf juice or water decoction of the plant in normal and alloxaninduced diabetic rabbits. 61

Reported mechanism of action:

Increases metabolization of glucose 60 and enhances secretion of insulin either from the beta cells of Langerhans or through extrapancreatic mechanism.⁶¹

21. Camellia sinensis

Common name: Green tea

Family: Theaceae

Activity with route of administration/dosage:

i. Antihyperglycemic activity of hot water extract of green tea in streptozotocin (STZ)-diabetic rats. 62, 63

Reported mechanism of action:

Epigallocatechin gallate, present in tea increases insulin activity and prevent oxidative damages, responsible for the hypoglycemic activity. 62, 63

Conclusion:

From the above work it is clear that number of medicinal plants used to treat hyperglycemia.

Acknowledgements:

The authors are very thankful to the director of the college.

Reference:

 Singh, K.N., Chandra, V., Barthwal, K.C., 1975. Letter to the editor: hypoglycemic activity of Acacia arabica, Acacia benthami and Acacia modesta leguminous seed diets in

- normal young albino rats. Indian Journal of Physiology and Pharmacology 19, 167–168.
- 2. Wadood, A., Wadood, N., Shah, S.A., 1989. Effects of Acacia arabica and Caralluma edulis on blood glucose levels of normal and alloxan diabetic rabbits. Journal of Pakistan Medical Association 39, 208–212
- 3. Karunanayake, E.H., Welihinda, J., Sirimanne, S.R., Sinnadorai, G., 1984. Oral hypoglycemic activity of some medicinal plants of Sri Lanka. Journal of Ethnopharmacology 11, 223–231.
- Ponnachan, P.T., Paulose, C.S., Panikkar, K.R., 1993. Effect of leaf extract of Aegle marmelose in diabetic rats. Indian Journal of Experimental Biology 31, 345–347.
- Das, A.V., Padayatti, P.S., Paulose,
 C.S., 1996. Effect of leaf extract of
 Aegle marmelose (L.) Correa ex
 Roxb. on histological and
 ultrastructural changes in tissues of
 streptozotocin induced diabetic rats.
 Indian Journal of Experimental
 Biology 34, 341–345.
- 6. Seema, P.V., Sudha, B., Padayatti, P.S., Abraham, A., Raghu, K.G.,

- Paulose, C.S., 1996. Kinetic studies of purified malate dehydrogenase in liver of streptozotocin-diabetic rats and the effect of leaf extract of Aegle marmelose (L.) Correa ex Roxb. Indian Journal of Experimental Biology 34, 600–602.
- 7. Upadhya, S., Shanbhag, K.K., Suneetha, G., Balachandra Naidu, M., Upadhya, S., 2004. A study of hypoglycemic and antioxidant activity of Aegle marmelos in alloxan induced diabetic rats. Indian Journal of Physiology and Pharmacology 48, 476–480.
- 8. Sabu, M.C., Kuttan, R., 2004. Hypoglycemic activity of Aegle marmelos and its relationship with its antioxidant properties. Indian Journal of Physiology and Pharmacology 48, 81–88.
- 9. Sachdewa, A., Raina, D., Srivastava, A.K., Khemani, L.D., 2001a. Effect of Aegle marmelos and Hibiscus rosa sinensis leaf extract on glucose tolerance in glucose induced hyperglycemic rats (Charles foster). Journal of Environmental Biology 22, 53–57.

- Kamalakkanan, N., Rajadurai, M., Prince, P.S., 2003. Effect of Aegle marmelos fruits on normal and streptozotocin-diabetic Wistar rats. Journal of Medicinal Food 6, 93–98.
- 11. Kamalakkannan, N., Prince, P.S., 2003. Hypoglycaemic effect of water extracts of Aegle marmelos fruits in streptozotocin diabetic rats. Journal of Ethnopharmacology 87, 207–210.
- 12. Kamalakkannan, N., Stanely, P., 2003. Effect of Aegle marmelos Correa. (Bael) fruit extract on tissue antioxidants in streptozotocin diabetic rats. Indian Journal of Experimental Biology 41, 1285–1288.
- 13. Augusti, K.T., 1973. Studies on the effects of a hypoglycemic principle from Allium Cepa Linn. Indian Journal of Medical Research 61, 1066–1071.
- 14. Gupta, R.K., Gupta, S., Samuel, K.C., 1977. Blood sugar lowering effect of various fractions of onion. Indian Journal of Experimental Biology 15, 313–314.
- 15. Babu, P.S., Srinivasan, K., 1997.
 Influence of dietary capsaicin and onion on the metabolic abnormalities

- associated with streptozotocin induced diabetes mellitus. Molecular and Cellular Biochemistry 175, 49–57
- 16. Jain, R.C., Vyas, C.R., 1975. Garlic in alloxan-induced diabetic rabbits. American Journal of Clinical Nutrition 28, 684–685.
- 17. Rabinkov, A., Miron, T., Konstantinovski, L., Wilchek, M., Mirelman, D., Weiner, L., 1998. The mode of action of allicin: trapping of radicals and interaction with thiol containing proteins. Biochimica et Biophysica Acta 1379, 233–244.
- 18. Rajasekaran, S., Sivagnanam, K., Ravi, K., Subramanian, S., 2004. Hypoglycemic effect of Aloe vera gel on streptozotocin-induced diabetes in experimental rats. Journal of Medicinal Food 7, 61–66.
- 19. Okyar, A., Can, A., Akev, N., Baktir, G., Sutlupinar, N., 2001. Effect of Aloe vera leaves on blood glucose level in type I and type II diabetic rat models. Phytotherapy Research 15, 157–161.
- 20. Ajabnoor, M.A., 1990. Effect of aloes on blood glucose levels in normal and alloxan diabetic mice.

- Journal of Ethnopharmacology 28, 215–220.
- 21. Ghannam, N., Kingston, M., Al-Meshaal, I.A., Tariq, M., Parman, N.S., Woodhouse, N., 1986. The hypoglycemic activity of aloes: preliminary clinical and experimental observations. Hormone Research 24, 288–294.
- 22. Subramoniam, A., Pushpangadan, P., Rajasekharan, S., Evans, D.A., Latha, P.G., Valsaraj, R., 1996. Effects of Artemisia pallens Wall. on blood glucose levels in normal and alloxan-induced diabetic rats. Journal of Ethnopharmacology 50, 13–17.
- 23. Shirwaikar, A., Rajendran, K., Dinesh Kumar, C., Bodla, R., 2004. Hypoglycemic activity of aqueous leaf extract of Annona squamosa in streptozotocin-nicotinamide type 2 diabetic rats. Journal of Ethnopharmacology 91, 171–175.
- 24. Gupta, R.K., Kesari, A.N., Murthy, P.S., Chandra, R., Tandon, V., Watal, G., 2005. Hypoglycemic and hypoglycemic effect of ethanolic extract of leaves of Annona squamosa L. in experimental

- animals. Journal of Ethnopharmacology 99, 75–81.
- 25. Borhanuddin, M., Shamsuzzoha, M., Hussain, A.H., 1994. Hypoglycaemic effects of Andrographis paniculata Nees on non-diabetic rabbits. Bangladesh Medical Research Council Bulletin 20, 24–26.
- 26. Chattopadhyay, R.R., 1996. Possible mechanism of antihyperglycemic effect of Azadirachta indica leaf extract, part IV. General Pharmacology 27, 431–434.
- 27. Gholap, S., Kar, A., 2004. Hypoglycaemic effects of some plant extracts are possibly mediated through inhibition in corticosteroid concentration. Pharmazie 59, 876–878.
- 28. Kar, A., Choudhary, B.K., Bandyopadhyay, N.G., 2003. Comparative evaluation of hypoglycaemic activity of some Indian medicinal plants in alloxan diabetic rats. Journal of Ethnopharmacology 84, 105–108.
- 29. Puri, D., Baral, N., 1998.Hypoglycemic effect of Biophytum sensitivum in the alloxan diabetic

- rabbits. Indian Journal of Physiology and Pharmacology 42, 401–406.
- 30. Puri, D., 2001. The insulinotropic activity of a Nepalese medicinal plant Biophytum sensitivum: preliminary experimental study. Journal of Ethnopharmacology 78, 89–93.
- 31. Yoshikawa, M., Murakami, T., Kadoya, M., Matsuda, H., Muraoka, O., Yamahara, J., Murakami, N., 1996. Medicinal foodstuff. III. Sugar beet. Hypoglycemic oleanolic acid oligoglycosides, betavulgarosides I, II, III, and IV, from the root of Beta vulgaris L. (Chenopodiaceae). Chemical and Pharmaceutical Bulletin (Tokyo) 44, 1212–1217.
- 32. Khan, B.A., Abraham, A., Leelamma, S., 1995. Hypoglycemic action of Murraya koeingii (curry leaf) and Brassica juncea (mustard): mechanism of action. Indian Journal of Biochemistry and Biophysics 32, 106–108.
- 33. Chude, M.A., Orisakwe, O.E.,Afonne, O.J., Gamaniel, K.S.,Vongtau, O.H., Obi, E., 2001.Hypoglycaemic effect of the aqueous extract of Boerhavia diffusa leaves.

- Indian Journal of Pharmacology 33, 215–216.
- 34. Pari, L., Amarnath Satheesh, M., 2004. Hypoglycemic activity of Boerhaavia diffusa L.: effect on hepatic key enzymes in experimental diabetes. Journal of Ethnopharmacology 91, 109–113.
- 35. Satheesh, M.A., Pari, L., 2004.
 Antioxidant effect of Boerhavia diffusa L. in tissues of alloxan induced diabetic rats. Indian Journal of Experimental Biology 42, 989–992.
- 36. Pari, L., Latha, M., 2002. Effect of Cassia auriculata flowers on blood sugar levels, serum and tissue lipids in streptozotocin diabetic rats. Singapore Medical Journal 43, 617–621.
- 37. Latha, M., Pari, L., 2003b.

 Preventive effects of Cassia auriculata L. flowers on brain lipid peroxidation in rats treated with streptozotocin. Molecular and Cellular Biochemistry 243, 23–28.
- 38. Latha, M., Pari, L., 2003a. Modulatory effect of Scoparia dulcis in oxidative stress-induced lipid peroxidation in streptozotocin

- diabetic rats. Journal of Medicinal Food 6, 379–386.
- 39. Abesundara, K.J., Matsui, T., K., 2004. Matsumoto, Alphaglucosidase inhibitory activity of some Sri Lanka plant extracts, one of which, Cassia auriculata, exerts a strong antihyperglycemic effect in rats comparable to the therapeutic drug acarbose. Journal of Agricultural and Food Chemistry 52, 2541-2545.
- 40. Sharma, S.R., Dwivedi, S.K., Swarup, D., 1997. Hypoglycemic, antihyperglycemic and hypolipidemic activities of Caesalpinia bonducella seeds in rats. Journal of Ethnopharmacology 58, 39–44.
- 41. Chakrabarti, S., Biswas, T.K., Rokeya, B., Ali, L., Mosihuzzaman, M., Nahar, N., Khan, A.K., Mukherjee, B., 2003. Advanced studies on the hypoglycemic effect of Caesalpinia bonducella F. in type 1 and 2 diabetes in Long Evans rats. Journal of Ethnopharmacology 84, 41–46.
- 42. Chakrabarti, S., Biswas, T.K., Seal, T., Rokeya, B., Ali, L., Azad Khan,

- A.K., Nahar, N., Mosihuzzaman, M., Mukherjee, B., 2005. Hypoglycemic activity of Caesalpinia bonducella F. in chronic type 2 diabetic model in Long-Evans rats and evaluation of insulin secretagogue property of its fractions on isolated islets. Journal of Ethnopharmacology 97, 117–122.
- 43. Esposito Avella, M., Diaz, A., de Gracia, I., de Tello, R., Gupta, M.P., 1991. Evaluation of traditional medicine: effects of Cajanus cajan L. and of Cassia fistula L. on carbohydrate metabolism in mice. Revista Medica de Panama 16, 39–45.
- 44. Panlasigui, L.N., Panlilio, L.M., Madrid, J.C., 1995. Glycemic response in normal subjects to five different legumes commonly used in the Philippines. International Journal of Food Science and Nutrition 46, 155–160.
- 45. Amalraj, T., Ignacimuthu, S., 1998a. Hypoglycemic activity of Cajanus cajan (seeds) in mice. Indian Journal of Experimental Biology 36, 1032–1033.
- 46. Abdel-Hassan, I.A., Abdel-Barry, J.A., Tariq Mohammeda, S., 2000.

- The hypoglycemic and antihyperglycemic effect of Citrullus colocynthis fruit aqueous extract in normal and alloxan diabetic rabbits. Journal of Ethnopharmacology 71, 325–330.
- 47. Nmila, R., Gross, R., Rchid, H., Roye, M., Manteghetti, M., Petit, P., Tijane, M., Ribes, G., Sauvaire, Y., 2000. Insulinotropic effect of Citrullus colocynthis fruit extracts. Planta Medica 66, 418–423.
- 48. Al-Ghaithi, F., El-Ridi, M.R., Adeghate, E., Amiri, M.H., 2004. Biochemical effects of Citrullus colocynthis in normal and diabetic rats. Molecular and Cellular Biochemistry 261, 143–149.
- 49. Mukherjee, K., Ghosh, N.C., Datta, T., 1972. Coccinia indica Linn. as potential hypoglycemic agent. Indian Journal of Experimental Biology 10, 347–349.
- 50. Singh, N., Singh, S.P., Vrat, S., Misra, N., Dixit, K.S., Kohli, R.P., 1985. A study on the anti-diabetic activity of Coccinia indica in dogs. Indian Journal of Medical Science 39, 27–29.

- 51. Chandrasekar, B., Mukherjee, B., Mukherjee, S.K., 1989. Blood sugar lowering potentiality of selected Cucurbitaceae plants of Indian origin. Indian Journal of Medical Research 90, 300–305.
- 52. Hossain, M.Z., Shibib, B.A., Rahman, R., 1992. Hypoglycemic effects of Coccinia indica: inhibition of key gluconeogenic enzyme, glucose-6- phosphatase. Indian Journal of Experimental Biology 30, 418–420.
- 53. Shibib, B.A., Khan, L.A., Rahman, R., 1993. Hypoglycemic activity of Coccinia indica and Momordica charantia in diabetic rats: depression of the hepatic gluconeogenic enzymes glucose-6-phosphatase and fructose-1,6- bisphosphatase and elevation of both liver and red-cell shunt enzyme glucose-6-phosphate dehydrogenase. Biochemistry Journal 292, 267–270.
- 54. Azad Khan, A.K., Akhtar, S., Mahtab, H., 1979. Coccinia indica in the treatment of patients with diabetes mellitus. Bangladesh Medical Research Council Bulletin 5, 60–66.

- 55. Platel, K., Srinivasan, K., 1997. Plant foods in the management of diabetes mellitus: vegetables as potential hypoglycemic agents. Die Nahrung 41, 68–74.
- 56. Kamble, S.M., Kamlakar, P.L., Vaidya, S., Bambole, V.D., 1998. Influence of Coccinia indica on certain enzymes in glycolytic and lipolytic pathway in human diabetes. Indian Journal of Medical Science 52, 143–146.
- 57. Prakasam, A., Sethupathy, S., Pugalendi, K.V., 2002.

 Antihyperglycaemic effect of Casearia esculenta root extracts in streptozotocin-induced diabetic rats.

 Pharmazie 57, 758–760.
- 58. Prakasam, A., Sethupathy, S., Pugalendi, K.V., 2003a. Effect of Casearia esculenta root extract on blood glucose and plasma antioxidant status in streptozotocin diabetic rats. Polish Journal of Pharmacology 55, 43–49.
- 59. Chattopadhyay, R.R., Sarkar, S.K., Ganguly, S., Banerjee, R.N., Basu, T.K., 1991. Hypoglycemic and antihyperglycemic effect of leaves of Vinca rosea Linn. Indian Journal of

- Physiology and Pharmacology 35, 145–151.
- 60. Singh, S.N., Vats, P., Suri, S., Shyam, R., Kumria, M.M., Ranganathan, S., Sridharan, K., 2001. Effect of an hypoglycemic extract of Catharanthus roseus on enzymic activities in streptozotocin induced diabetic rats. Journal of Ethnopharmacology 76, 269–277.
- 61. Nammi, S., Boini, M.K., Lodagala, S.D., Behara, R.B., 2003. The juice of fresh leaves of Catharanthus roseus Linn. reduces blood glucose in normal and alloxan diabetic rabbits. BMC Complementary and Alternative Medicine 3, 4.
- 62. Gomes, A., Vedasiromoni, J.R., Das, M., Sharma, R.M., Ganguly, D.K., 1995. Anti-hyperglycemic effect of black tea (Camellia sinensis) in rat. Journal of Ethnopharmacology 45, 223–226.
- 63. Anderson, R.A., Polansky, M.M., 2002. Tea enhances insulin activity. Journal of Agricultural and Food Chemistry 50, 7182–7186.