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Evaluation of antidiabetic and related actions of some Indian medicinal plants in diabetic rats

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ABSTRACT

Objective: To evaluate antidiabetic activity of chloroform extracts of *Acacia arabica* bark, *Benincasa hispida* fruit, *Tinospora cordifolia* stem, *Ocimum sanctum* areal parts and *Jatropha curcas* leaves. **Methods:** The chloroform extracts of *Acacia arabica* bark, *Benincasa hispida* fruit, *Tinospora cordifolia* stem, aerial part of *Ocimum sanctum* and *Jatropha curcas* leaves were evaluated at different doses (250 and 500 mg/kg body weight.) for antidiabetic potentials in alloxan induced diabetic albino rats. The extracts were administered for two weeks in different groups whereas tolbutamide (80 mg/kg body weight) was used as reference standard throughout study. **Results:** The result of present study showed test compounds significantly decreases elevated level of serum glucose and also caused to reverse the cholesterol, triglyceride, HDL and LDL values when compared to untreated diabetic rats. **Conclusions:** Our finding indicates that different test extracts were able to ameliorate the derangements in lipid metabolism caused by diabetes mellitus in alloxan induced diabetic rats towards normal level.

1. Introduction

Diabetes mellitus is a chronic metabolic disorder caused by an absolute or relative lack of insulin and/or reduced insulin activity which results in hyperglycemia and abnormalities in carbohydrate, fat and protein metabolism^[1,2]. Management of diabetes without any side effects is still a challenge for medical system. This leads to an increasing search for improved antidiabetic drugs. Few of plant treatments used in traditional medicine for diabetes received scientific scrutiny and WHO has recommended that this area warrants attention^[3]. *Acacia arabica* (Mimosaceae); *Benincasa hispida* (Cucurbitaceae); *Tinospora cordifolia* (Menispermaceae); *Jatropha curcas* (Euphorbiaceae) and *Ocimum sanctum* Linn are traditionally used for treatment of diabetes^[4]. *Acacia arabica* possesses antibacterial^[5], antiviral^[6] and antifertility activities^[7]. *Tinospora cordifolia* was reported to have adaptogenic^[8], antiviral^[9], antihyperglycemic^[10], immunomodulatory^[11], antineoplastic^[12], cardioprotective^[13] and antioxidants^[14,15],

activities. *Ocimum sanctum* showed antioxidant, antifungal^[16], antimicrobial^[17] and wound healing^[18] activities. *Jatropha curcas* possesses anti-diabetic^[19], wound-healing^[20], antitumor^[21], anti-inflammatory^[22] and antifungal^[23] activities. *Benincasa hispida* was reported to have antinociceptive, anti-pyretic^[24] and anti-stress^[25] activities. The objective of present study was to evaluate antidiabetic activity of chloroform extracts of *Acacia arabica* bark, *Benincasa hispida* fruit, *Tinospora cordifolia* stem, *Ocimum sanctum* areal parts and *Jatropha curcas* leaves.

2. Material and methods

2.1. Plant material

The bark of *Acacia arabica*, fruit of *Benincasa hispida*, stem of *Tinospora cordifolia*, aerial part of *Ocimum sanctum* and leaves of *Jatropha curcas* were collected from Pune District, Maharashtra, India in August 2008. The plants were identified and authenticated by Dr. R. B. Deshmukh, Department of Botany, Sharadabai Pawar College, Malegaon(bk) Tal- Baramati Dist- Pune.(India). The voucher specimens (AA-08, BH-08, TC-08, OS-08 and

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JC-08 respectively) are deposited in herbarium. The samples were cleaned, dried in shade and powdered. The powdered samples were kept in airtight container for further studies.

2.2. Chemicals and drugs

Alloxan monohydrate, tolbutamide and chloroform were purchased from Sigma Chemicals, St. Louis. Chemical kits for estimation of blood glucose, cholesterol, triglyceride, LDL and HDL were purchased from Erba Diagnostics, Mannheim.

2.3. Animals

The male albino rats (Wistar strain weighing 150–180 g) and albino mice (weighing 20–30 g) were procured from SVPM's College of Pharmacy, Malegaon (bk), Tal– Baramati Dist– Pune, India and housed in animal house. They were kept at 27 ± 3 °C (Relative humidity: $65\% \pm 10\%$ and light / dark cycle for 12 h. All the animals were fed with rodent pellet diet (Gold Mohr, Lipton India Ltd.) and water was allowed ad–libitum under strict hygienic conditions. Institutional Animal Ethics Committee (IAEC) approved all the protocol of study (Reg. No. 1214/ac/08/CPCSEA).

2.4. Preparation of extract

Dried powders each of 100 g of *Acacia arabica* bark (CEAA), *Benincasa hispida* fruit (CEBH), *Tinospora cordifolia* stem (CEOS), *Ocimum sanctum* aerial part and leaves of *Jatropha curcus* (CETC) were extracted with chloroform by Soxhlet extraction and solvents were evaporated by rotary evaporator. Concentrated extracts were dried on water bath at controlled temperature to yield CEAA 12.30%w/w, CEBH 14.08%w/w, CETC 9.69%w/w, CEOS 15.07%w/w and CEJC 11.31%w/w respectively.

2.5. Toxicity studies

An acute toxicity of CEAA, CEBH, CETC, CEOS and CEJC were carried out in female albino mice (25–30 g). The mice were divided in to six groups ($n=6$). The animals were fasted for overnight prior to the experimentation. The fixed dose method of CPCSEA (Mrs. Prema Veeraraghavan. Expert consultant, CPCSEA, OECD guide line No. 420; Oct 2000) was adopted for toxicity studies.

2.6. Preparation of diabetic rats

Alloxan monohydrate dissolved in saline was injected in rats intraperitoneally at dose of 150 mg/ kg body weight. After a fortnight, rats with marked hyperglycemia were selected and used for the study.

2.7. Experimental design

The rats were divided into thirteen groups ($n=5$). Group I (controlled group) administered 0.5 mL saline, Group II (untreated diabetic rats), group III (diabetic rats receiving

tolbutamide orally at 80 mg/kg body weight in saline), Groups IV to XIII (diabetic rat treated with CEAA, CEBH, CETC, CEOS and CEJC at dose of 250 and 500/mg kg body weight). Tolbutamide was used as the standard antidiabetic throughout the experimentation. The animals were carefully monitored at every day. Animals described as fasted were deprived of food for at least 12 h but allowed to free access for drinking water. Fasting blood glucose measurement was done on day 0, 7th and 14th of the study. Blood samples were collected by retro orbital method at weekly intervals till the end of study and processed for estimation of serum glucose and serum lipid[26, 27, 28].

2.8. Biochemical estimation

Serum samples from all the experimental rats were collected for estimation of biochemical parameters, serum glucose (GOD–POD method), cholesterol (CHOD–PAP method), triglycerides (GPO–Triender method), HDL and LDL[29].

2.9. Statistics

All values are expressed as Mean \pm SEM. The differences were compared using one way analysis of variance (ANOVA) followed by Tukey's multiple comparison tests. P values < 0.05 were considered as significant.

3. Result

In untreated diabetic rats, glucose, cholesterol, triglycerides, LDL values were elevated to high level during the study where as HDL value was decreased. Chronic treatment with CEAA, CEBH, CETC, CEOS and CEJC at 250 and 500 mg/kg significantly ($P < 0.0001$) causes decrease in serum glucose on 7th and 14th day as shown in Figure 1, 2 and 3. Serum cholesterol triglycerides HDL and LDL levels also reversed with treatment of test extracts and standard as shown in table 1.

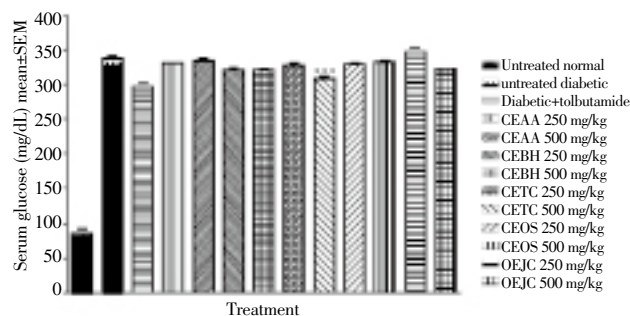


Figure 1. Effect of CEAA, CEBH, CETC, CEOS and CEJC on serum glucose level in diabetic rats on 0 day.

All values are expressed as Mean \pm SEM ($n=5$). Statistical comparisons between each treatment and untreated diabetic were carried out by one way ANOVA followed by Tukey's multiple comparison tests. *** $P < 0.0001$ when compare to untreated diabetic group.

Table 1

Comparative effect of CEEA, CEBH, CETC, CEOS and CEJC on serum cholesterol, triglycerides, HDL and LDL (mg/dl) in diabetic rats

Groups	Treatment	Average serum lipid profile (mg/dL)			
		Total cholesterol	Triglycerides	HDL cholesterol	LDL cholesterol
I	Normal	64.28±3.51	79.64±3.12	48.60 ±3.72	48.53± 4.36
II	Diabetic	95.28±4.01	143.88±3.52	26.14 ±5.61	108.23±7.59
III	Diabetic + Tolbutamide 80 mg/ kg	58.78±3.61 ^{***}	81.10±4.00 ^{***}	41.86 ± 5.94 ^{**}	55.34±5.46 ^{**}
IV	Diabetic+CEEA 250 mg/kg	72.46±2.69 ^{***}	109.60±3.06 ^{***}	30.14±4.18	81.34±2.30 ^{***}
V	Diabetic+CEEA 500 mg/kg	61.39±2.06 ^{***}	97.59±2.33 ^{***}	35.24±4.84	65.71±0.60 ^{***}
VI	Diabetic+CEBH 250 mg/kg	72.01±2.21 ^{***}	109.56±1.23 ^{***}	31.16±5.14	86.73±1.58 ^{**}
VII	Diabetic+CEBH 500 mg/kg	64.69±1.08 ^{***}	87.41±2.95 ^{***}	39.30±4.19	63.24±1.60 ^{**}
VIII	Diabetic+CETC 250mg/kg	79.43±3.07 [*]	100.79±2.82 ^{***}	30.18±3.51	89.24±2.60 ^{**}
IX	Diabetic+CETC 500mg/kg	66.82±1.62 ^{***}	82.91±2.70 ^{***}	36.13±3.24	71.10±0.60 ^{***}
X	Diabetic+CEOS 250mg/kg	80.23±2.65 [*]	92.80±1.18 ^{***}	31.24±4.14	91.46±1.69 [*]
XI	Diabetic+CEOS 500mg/kg	65.26±2.71 ^{***}	83.47±2.91 ^{***}	37.62±3.31	64.18±0.50 ^{***}
XII	Diabetic+CEJC 250mg/kg	81.52±2.69 [*]	87.06±2.02 ^{***}	29.92±4.12	93.24±1.65
XIII	Diabetic+CEJC 500mg/kg	72.18±3.01 ^{***}	95.48±3.74 ^{***}	38.81±5.21	72.09±1.90 ^{***}

Values are expressed as Mean ±SEM; (n = 5), a P < 0.05, b p < 0.01, c P<0.001 compared with untreated diabetic rats.

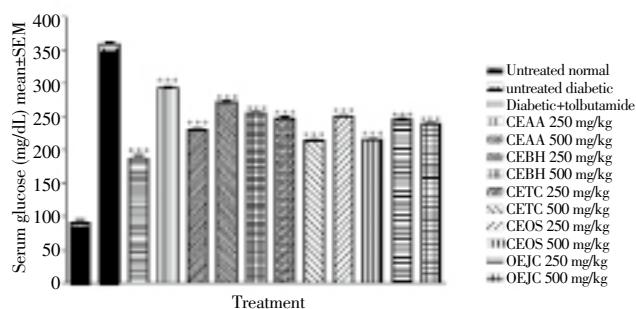


Figure 2. Effect of CEEA, CEBH, CETC, CEOS and CEJC on serum glucose level in diabetic rats on 7th day.

All values are expressed as Mean ± SEM (n=5). Statistical comparisons between each treatment and untreated diabetic were carried out by one way ANOVA followed by Tukey's multiple comparison tests. *** P < 0.0001 when compare to untreated diabetic group.

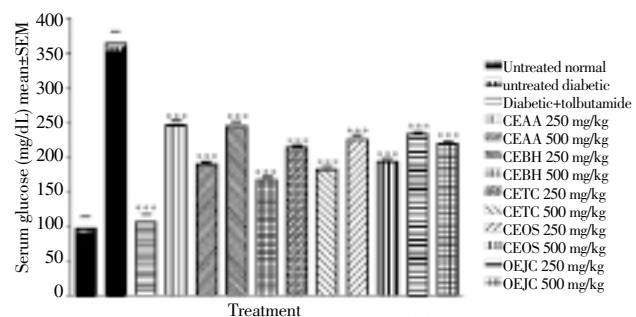


Figure 3. Effect of CEEA, CEBH, CETC, CEOS and CEJC on serum glucose level in diabetic rats on 14th day.

All values are mean ± SEM (n=5). Statistical comparisons between each treatment and untreated diabetic were carried out by one way ANOVA followed by Tukey's multiple comparison tests. *** P < 0.0001 when compare to untreated diabetic group.

4. Discussion

Alloxan induces diabetes by damaging the insulin secreting cells of pancreas leading to hyperglycemia^[30].

Diabetes mellitus is a group of syndromes characterized by hypoglycemia, altered metabolism of lipids, carbohydrates and proteins. It is an increased risk of complications from vascular diseases^[31].

An observation in this study correlates with the previous research finding, in that the blood glucose levels significantly increased in alloxan induced untreated diabetic rats. In the present study, the continuous treatment of different test compounds for a period of two weeks caused a significant decrease in blood glucose of treated diabetic rats. These results have confirmed the earlier results of our preliminary studies^[32]. We have also observed an increase in concentration of cholesterol, triglyceride, LDL and decrease in HDL in alloxan untreated diabetic rats. Hyperlipidaemia is recognized consequence of diabetes mellitus^[33–35]. Chronic administration of different tests extracts normalized serum lipid profile i.e. secondary to diabetic state. Diabetes induced hyper lipidaemia is attributable to excess mobilization of fat from adipose due to the under utilization of glucose^[36]. The regression of the diabetic state due to chronic administration of different test extract at different doses increased the utilization of glucose, thereby depressing the mobilization of fat.

In conclusion, the present investigation supports the hypoglycemic effects of extracts of ethnic folk medicine which mitigates pathological states like diabetes mellitus.

Conflict of interest statement

We declare that we have no conflict of interest.

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