Hypoglycemic Effect of *Kyllinga triceps* in STZ Induced Diabetic Rats

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**Abstract**

*Kyllinga triceps* is an Indian medicinal plant demonstrated to exert multiple pharmacological effects. In light of traditional claim of the plant in treatment of diabetes, the present study was carried out to evaluate the hypoglycemic activity of the plant in Streptozotocin (STZ) induced diabetic rats. The diabetic rats were given *Kyllinga triceps* extract (KTE, 100mg/kg & 200mg/kg) and Glibenclamide (0.5mg/kg) for 28 days. The effects of both treatments on body weight and blood glucose were assessed. Both doses of extract and Glibenclamide showed significant hypoglycemic activity in acute and sub acute study. Body weight of extract and glibenclamide treated rats were maintained during the study period. Where as body weight of untreated rats had gone down. The result obtained from the study scientifically proved the folkloric use of *Kyllinga triceps* as hypoglycemic agent. Thus the plant can be key contributor in treatment of Diabetes.

**Keywords:** Diabetes; Glibenclamide; Hypoglycemic; *Kyllinga triceps*

**Introduction**

Type 2 diabetes mellitus, also known as noninsulin dependent diabetes mellitus, develops in middle or later life and affects 2-6% of adults in most Western societies [1]. Diabetes mellitus has been recognized as a growing world-wide epidemic by many health's advocacy groups including the World Health Organization (WHO) [2]. The WHO has estimated that diabetes will be one of the worlds leading causes of death and disability within the next quarter century. The statistics are alarming; 30 million people were diagnosed with diabetes world-wide in 1985, by 1995 the number had risen to 135 million, and at the current rate there will be 300 million by the year 2025 as predicted by the WHO [3].

The limitations of currently-available oral antidiabetic agents either in terms of efficacy/safety coupled with the emergence of the disease into a global epidemic have encouraged a concerted effort to discover drugs that can manage type 2 diabetes more efficiently [4]. Also, with increasing incidence of diabetes mellitus in rural population throughout the world and due to adverse effects of synthetic medicine, there is a clear need for development of indigenous, inexpensive botanical sources for anti-diabetic crude or purified drugs [5].

Medicinal plants use has turned out to be an alternative method for treatment of disease such as diabetes mellitus. The recommended oral antidiabetic agents either in terms of efficacy/safety coupled with the emergence of the disease into a global epidemic have encouraged a concerted effort to discover drugs that can manage type 2 diabetes more efficiently [4]. Also, with increasing incidence of diabetes mellitus in rural population throughout the world and due to adverse effects of synthetic medicine, there is a clear need for development of indigenous, inexpensive botanical sources for anti-diabetic crude or purified drugs [5].

Medicinal plants use has turned out to be an alternative method for treatment of disease such as diabetes mellitus. Literature reports more than 800 plants have been utilized as empirical treatment for diabetes. One tenth of them have been characterized as hypoglycemic plants with active compounds such as mucilage gum, glycans, flavonoids, triterpenes and alkaloids [6].

Research conducted in last few decades on plants mentioned in ancient literature or used traditionally for diabetes has shown antidiabetic property.

The recommendations of the WHO committee on diabetes encouraging research on hypoglycemic agents of plant origin used in traditional medicine has greatly motivated research in this area [7]. Currently several hundred plants have been reported to have beneficial effect in treatment of diabetes [8-10].

*Kyllinga triceps* (Family: Cyperaceae) is common throughout India. It is also known as Nirvishaa, Nirbishi and Mustaa. Its root is used as diuretic (in polyuria), demulcent, antidermatosis and antidiabetic [11]. Decoction of the plant *Kyllinga triceps* showed antipyretic activity [12].

Different ethnic communities of Thoubal district of Manipur used to treat diabetes by 54 plant species. Boiled plant extract is used to treat diabetes by village priests cum doctors (Locally called Maida or Maidi) [13]. Jain et al reported that *Kyllinga triceps* is used to treat diabetes by local folklore and tribal of different districts of Madhya Pradesh and Chhattisgarh [14].

In the present study we have selected *Kyllinga triceps* to examine its hypoglycemic activity because traditional documents claimed that to be an antidiabetic and used as folkloric medicine in different region of India.

**Methodology**

**Collection of plants**

The plant was collected and authenticated by the comparison with the herbarium and voucher specimen (Voucher specimen No. 97767) was lodged in the department herbarium of National Botanical Research Institute, Lucknow.

**Preparation of plant extract**

Plants were shade dried, milled, and ground into coarse powder with the help of a mixer. The powdered material was subjected to cold maceration using sufficient quantity of ethanol and distilled water (1: 1) for 10 days with intermittent shaking in a round bottom flask. On 10th day, it was strained and marcs were pressed. The pressed liquid was

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added to the strained liquid and the combined liquid was clarified by filtration and the filtrate was subjected to distillation at temperature 60°C for removing ethanol and water. After distillation, the semi solid obtained was kept for experiment.

Experimental animals

Healthy wistar albino rats were used for the study. They were maintained at standard laboratory condition and fed with commercial pellet diet and water ad libitum. The animals were acclimatized to laboratory condition for one week before commencement of experiment.

Acute oral toxicity study

In present study observation of acute toxicity study carried according to OECD guideline 423, none of the rats showed observable signs of toxicity upon single administration of kyllinga extract (2 g/kg) and divided into following group consisting five animals each.

Group I – Non diabetic control
Group II – Diabetic Control
Group III –Diabetic rats administered with Glibenclamide at dose of (0.5 mg/kg)
Group IV –Diabetic rats administered with Kyllinga triceps extract (KTE) in dose of 100 mg/kg
Group V – Diabetic rats administered with Kyllinga triceps extract (KTE) in dose of 200 mg/kg

The acute study involved estimation of blood glucose levels at 0, 2, 4, 6 and 24 hrs after administration of Glibenclamide and KTE.

The sub acute study involved daily administration of Glibenclamide and KTE for 28 days and blood glucose levels were estimated on 0, 7,14,21 and 28 days.

Blood glucose estimation

Blood samples were obtained through puncture tail vein and collected blood samples were analyzed by Accu-Check Glucometer. Blood glucose levels were expressed in terms of mg/dl.

Body weight determination

Weight of rats was recorded during the study period of 28 days. Percentage (%) change in body weight was calculated and tabulated.

Results

In acute toxicity studies of Kyllinga extract given by oral route did not show any mortality and toxic effects up to the dose of 2000 mg/kg body weight. So 1/20th and 1/10th of safe dose is used for the experiments.

STZ- induced diabetic rats exhibiting persistent hyperglycemia (Blood Glucose>200 mg/dl) were selected for assessing the effect of KTE. Table 1 shows the blood glucose level at various time intervals to observe acute effect of two different doses of KTE and Glibenclamide. Data showed significant (p < 0.05) difference in blood glucose of all the treatments compared to diabetic control. Glibenclamide showed highest % decrease (45.7%) followed by KTE (200 mg/kg) (40%). KTE (100 mg/kg) showed lowest % decrease (36.4%) in blood glucose level. It is clear from the data that KTE (200 mg/kg) showed more hypoglycemic activity than KTE (100 mg/kg) in acute study.

To observe long term effect, sub acute study of 28 days was done. Table 2 shows the results of this study. Glibenclamide and KTE (200 mg/kg) showed highly significant (p < 0.01) hypoglycemic effect compared to diabetic control while KTE (100 mg/kg) also showed significant (p < 0.05) hypoglycemic activity. KTE (200 mg/kg) showed less % reduction (47.2%) than Glibenclamide (55.4%). In sub acute study also KTE (200mg/kg) is more effective than KTE (100 mg/kg).

Table 1: Acute effect of KTE on blood glucose level in STZ induced diabetic rats.

| GROUPS | BLOOD GLUCOSE (mg/dl) |
|--------|-----------------------|
|        | 0 hr | 2 hr | 4 hr | 6 hr | 24 hr |
| Normal Control | 93 ± 1.9 | 93 ± 2.7 | 95 ± 1.2 | 94 ± 3.1 | 91 ± 1.4 |
| Diabetic Control | 230 ± 2.4 | 237 ± 1.6 | 239 ± 3.2 | 244 ± 2.1 | 254 ± 4.1 |
| Diabetic+ Glibenclamide | 247 ± 2.3 | 194 ± 2.7* | 158 ± 1.8* | 134 ± 1.1* | 197 ± 3.6 |
| Diabetic+ KTE (100 mg/kg) | 222 ± 3.7 | 193 ± 2.4* | 164 ± 2.3* | 141 ± 1.9* | 202 ± 2.8 |
| Diabetic+ KTE (200 mg/kg) | 220 ± 2.4 | 185 ± 4.3* | 152 ± 3.1* | 131 ± 2.7* | 190 ± 3.8 |

All values are expressed as mean ± S.E.M (n=5). *P<0.05 as compared to diabetic control. One-way ANOVA followed by Dunnet multiple comparison test.

Table 2: Sub acute effect of KTE on blood glucose level in STZ induced diabetic rats.

| GROUPS | BLOOD GLUCOSE (mg/dl) |
|--------|-----------------------|
|        | 0 DAY | 7 DAY | 14 DAYS | 21 DAY | 28 DAY |
| Normal Control | 93 ± 2.7 | 101 ± 2.2 | 100 ± 2.6 | 98 ± 1.8 | 102 ± 3.1 |
| Diabetic Control | 230 ± 3.2 | 242 ± 2.8 | 258 ± 1.9 | 276 ± 3.8 | 298 ± 2.1 |
| Diabetic+ Glibenclamide | 247 ± 1.9 | 210 ± 2.9* | 180 ± 1.2** | 151 ± 1.3** | 110 ± 3.7** |
| Diabetic+ KTE (100 mg/kg) | 222 ± 2.2 | 206 ± 3.1* | 190 ± 3.6* | 174 ± 1.3* | 137 ± 3.9* |
| Diabetic+ KTE (200 mg/kg) | 220 ± 3.1 | 191 ± 2.9** | 164 ± 2.7** | 141 ± 4.2** | 116 ± 1.9** |

All values are expressed as mean ± S.E.M (n=5). *P<0.05 as compared to diabetic control. One-way ANOVA followed by Dunnet multiple comparison test.

Table 3: Effect of KTE on body weight in STZ induced diabetic rats.

| GROUPS | BODY WEIGHT (gm) |
|--------|------------------|
|        | 0 DAY | 7 DAY | 14 DAY | 21 DAY | 28 DAY |
| Normal Control | 140 ± 3.9 | 142 ± 4.2 | 144 ± 1.9 | 145 ± 3.9 | 145 ± 3.2 |
| Diabetic Control | 158 ± 4.7 | 154 ± 4 | 150 ± 3.7 | 148 ± 2.9 | 142 ± 1.7 |
| Diabetic+ Glibenclamide | 139 ± 1.8 | 139 ± 2.3* | 140 ± 3.5** | 144 ± 1.8*** | 148 ± 4.1*** |
| Diabetic+ KTE (100 mg/kg) | 139 ± 1.9 | 139 ± 2.8* | 143 ± 2.4* | 144 ± 3.4* | 146 ± 2.9* |
| Diabetic+ KTE (200 mg/kg) | 162 ± 2.3 | 164 ± 2.6** | 163 ± 3.6** | 167 ± 4.5** | 171 ± 4.1** |

All values are expressed as mean ± S.E.M (n=5). *P<0.05, **P<0.01 as compared to diabetic control. One-way ANOVA followed by Dunnet multiple comparison test.
Table 3 shows the body weight of rats at different time intervals during the study. The % change in body weight of all the treatments was calculated. Non diabetic control showed 3.4% increase in body weight while diabetic control showed 10.1% weight decrease. Glibenclamide, KTE (200 mg/kg) and KTE (100 mg/kg) showed 6.4, 5.0 and 5.5% increase in body weight respectively. All the treatments showed more % increase in body weight than non diabetic control. KTE (200 mg/kg) and KTE (100 mg/kg) showed nearly similar effect on body weight. So KTE not only showed significant hypoglycemic activity but also effective in preventing the weight loss due to diabetes.

Discussion

Streptozotocin is well known for its selective pancreatic islets β cell cytotoxicity and has extensively used to induce diabetes in experimental rat model. It interferes with cellular metabolic oxidative mechanism. Glibenclamide is often used as a standard antidiabetic drug in STZ induced moderate diabetes to compare the efficacy of variety of hypoglycemic compounds or plant extracts. The present study was conducted to assess the hypoglycemic activity of Kyllinga extract in STZ induced diabetic rats. Results shows both the doses of Kyllinga extract significantly improving the body weight and controlling the blood glucose level in diabetic rats. 200 mg/kg dose is more effective than 100 mg/kg and comparable to Glibenclamide. Result of the study scientifically validates the folkloric use of Kyllinga triceps in different ethnic communities of Manipur, Madhya Pradesh and Chhattisgarh.

It may be concluded that Kyllinga extract possess hypoglycemic activities and the plant extract may be used as hypoglycemic agent. The plants should be considered as an excellent candidature for further studies on diabetes mellitus.

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