

# Hypoglycaemic Activity of Bitter Gourd Fruits Extracts in Normal and Alloxan-Induced Diabetic Rats

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## Abstract

The purpose of the study was to investigate the effect of bitter gourd fruits (*Momordica charantia*) extracts on blood glucose concentration and liver enzymes in normal and alloxan-induced diabetic rats.

Oral administration of an aqueous extract of bitter gourd fruits (BGF) to alloxan diabetic rats caused a significant reduction of blood glucose and serum transaminases (aminotransferases). However, bitter gourd fruits extract didn't affect the blood glucose in normal rats.

Maximum reduction in serum glucose was observed after 4 h at a dose level of 50 mg extract/kg of body weight and considers the optimum dose. Chronic administration of the extract significantly reduced the blood glucose in alloxan-induced diabetic rats for fifteen days. The bitter gourd fruits extract also lowers the aspartate transaminase (sAST) and alanine transaminase (sALT) in diabetic rats.

Thus results suggest that the bitter gourd fruits not only reduce blood glucose level, but may be able to ameliorate biochemical damages reduced by alloxan in diabetic rats.

**Keywords:** Bitter gourd; Hypoglycaemic activity; Transaminases

## Introduction

Diabetes mellitus is a major disease that affects more than one hundred million people and may attain about five times more subjects in the next ten years [1]. Its control involves exercise, diet and chemotherapy. However, the pharmaceutical drugs at the authors disposal are either too expensive or have undesirable side effects and contraindications [2-4]. Historically, the study of the pathogenesis of diabetes mellitus was in the traditional pattern of endocrinology, the major dysfunction of  $\beta$ -cells associated with the pre-overt phase of type I diabetes or with type II diabetes is a decreased secretory response to glucose [5-7].

After insulin became available, evidence emerged suggesting that human diabetes mellitus has a multifactorial etiology. Insulin was and still the major hypoglycaemic medication used in diabetes mellitus treatment. In order to discover other hypoglycaemic agents, many investigations have been performed in traditional medicine testing eventual hypoglycaemic plants [8-12]. In this view, many studies have demonstrated hypoglycaemic activity of some plants in different animal models [13-16].

The bitter gourd was widely distributed in New Valley Governorate (Egypt). Its water extract was largely used in Egypt on traditional medicine as anti-diabetic without scientific background.

In this study it has been investigated the effect of extract of bitter gourd on blood glucose and serum transaminases in normal and alloxan induced diabetic rats.

## Materials and Methods

### Plant materials

Bitter gourd fruits were collected from New Valley Governorate in July-September 2002, and dried. Plant material was prepared according to the traditional method 10 g of dried sample was boiled at 100°C in 100 ml of distilled water for 10 min. Then cooled at room temperature for 20 min and filtered. The water extract was prepared daily just before administration.

### Animals

Male albino rats (*Rattus norvegicus*) were examined well to ensure

that they are free from any signs of microbial or parasitic diseases. Their body weights were about (120 ± 10 g). Rats were adapted in large cages with sawdust bottom coating to the controlled laboratory conditions for three weeks before carrying out the inoculation procedures and they were provided with commercial rodent food pellets and water.

### Effect of bitter gourd fruits extract on blood glucose and liver enzymes levels of normal rats

Normal rats were divided into four groups of eight rats per groups.

Group I: (Untreated controls) Group rats receive distilled water and served as control group.

Group II: (Treated controls) Normal rats receiving (25 mg/kg body weight) orally the BGF extract.

Group III: (Treated controls) Normal rats receiving (50 mg/kg body weight) orally the BGF extract.

Group IV: Normal rats receiving (100 mg/kg body weight) orally the BGF extract.

Blood samples for plasma glucose and liver enzymes determination was obtained by a tail vein at 2, 4 and 6 hours after giving drug extract [17-19].

### Alloxan-induced hyperglycemia in rats

Hyperglycemia was induced by the intraperitoneally injection of

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alloxan (120 mg/kg body weight) dissolved in distilled water. Five days after injection of alloxan blood glucose levels of all surviving rats were determined only animals with blood glucose levels over than 350 mg/dl were used.

### Effect of the bitter gourd fruits extraction on alloxan induced diabetic rats

The diabetic rats were divided into four groups each group contain eight rats group I (control group) were given distilled water orally, while group II, group III and group IV were given water extract 25, 50 and 100 mg/kg body weight. Blood sample were collected from the tail vein 2, 4 and 6 h after the water extract administration.

### Hypoglycaemic effect of water extract on alloxan-induced diabetic rats through longer duration

Male albino rats were divided five groups of eight rats in each group. Group I (normal rats) were given distilled water and served as normal. Group II (control diabetic) diabetic rats were given distilled water instead of extract. Groups III-V was given BGF extract orally at dose 25, 50 and 100 body weights. The oral administration of BGF extract was continued for fifteen days. Blood samples were collected from tail vein after 5, 10 and 15 days.

### Biochemical analysis

Blood sugar was determined according to the method described by Trinder, ALT and AST by Reitman and Frankel [20-23].

### Statistical analysis

Results are represented as mean  $\pm$  standard deviation (S.D). A two way analysis variance (ANOVA) according to Mason [19] for testing the significance between alloxan controls, treated and control group was performed.

### Results

Change in blood glucose level in normal and treated rats with different dose of BGF were presented in Table 1. Administration of BGF extract was found didn't reduce blood glucose level in normal rats.

Table 2 show the effect of various doses of the extract of BGF on blood glucose level in diabetic rats. The blood glucose was increased

Groups	Treatment	Mean Blood Glucose (mg/dl)			
		Initial	2 h	4 h	6 h
I	Control	74 $\pm$ 3	77 $\pm$ 2	75 $\pm$ 2	76 $\pm$ 3
II	BGF 25 X	71 $\pm$ 2	69 $\pm$ 2	72 $\pm$ 2	71 $\pm$ 2
III	BGF 50 X	73 $\pm$ 3	71 $\pm$ 2	67 $\pm$ 2	69 $\pm$ 3
IV	BGF 100 X	72 $\pm$ 2	70 $\pm$ 1	69 $\pm$ 3	70 $\pm$ 2

Table 1: Effect of bitter gourd fruits extract on blood glucose levels in normal rats.

Groups	Treatment	Mean Blood Glucose (mg/dl)			
		Initial	2 h	4 h	6 h
I	Control (distilled water)	350 $\pm$ 5	352 $\pm$ 6	348 $\pm$ 8	346 $\pm$ 7
II	BGF 25 X	355 $\pm$ 9	352 $\pm$ 7	349 $\pm$ 8	342 $\pm$ 8
III	BGF 50 X	352 $\pm$ 6	315 $\pm$ 7	245 $\pm$ 9	260 $\pm$ 8
IV	BGF 100 X	360 $\pm$ 9	352 $\pm$ 8	301 $\pm$ 7	307 $\pm$ 7

Result Mean  $\pm$  S.D  
Significant p<0.05  
X mg/kg body weight

Table 2: Effect of bitter gourd fruits extract on blood glucose levels in diabetic rats.

Groups	Treatment	Mean Blood Glucose (mg/dl)			
		Initial	5 days	10 days	15 days
I	Normal	78 $\pm$ 2	76 $\pm$ 2	80 $\pm$ 2	79 $\pm$ 2
II	Diabetic (Distilled water)	360 $\pm$ 8	352 $\pm$ 8	311 $\pm$ 9	315 $\pm$ 7
III	BGF 25 X	358 $\pm$ 9	249 $\pm$ 6	188 $\pm$ 8	178 $\pm$ 9
IV	BGF 50 X	355 $\pm$ 10	225 $\pm$ 9	165 $\pm$ 7	155 $\pm$ 6
V	BGF 100 X	370 $\pm$ 7	235 $\pm$ 8	175 $\pm$ 7	168 $\pm$ 8

Table 3: Hypoglycaemic effect of BGF extract on blood glucose levels in diabetic rats through long duration.

Groups	Treatment	Body weight (g)			
		Initial	5 days	10 days	15 days
I	Normal	118 $\pm$ 2	122 $\pm$ 2	132 $\pm$ 1	141 $\pm$ 2
II	Diabetic	120 $\pm$ 3	102 $\pm$ 2	104 $\pm$ 2	106 $\pm$ 3
III	BGF 25 X	116 $\pm$ 3	112 $\pm$ 2	114 $\pm$ 3	118 $\pm$ 3
IV	BGF 50 X	117 $\pm$ 4	122 $\pm$ 3	126 $\pm$ 3	129 $\pm$ 2
V	BGF 100 X	118 $\pm$ 3	120 $\pm$ 2	124 $\pm$ 2	128 $\pm$ 2

Result Mean  $\pm$  S.D  
Significant p<0.05  
X mg/kg body weight

Table 4: Effect of BGF on body weight of normal and diabetic rats.

in alloxan diabetic rats as compared to normal rats. Administration of BGF extract decreased blood glucose as compared to diabetic rats. The maximum reduction was rated after 4 h after administration of the BGF extract. There was reduction of blood glucose in these rats four hours after the administration of BGF; 25 mg/kg body weight (349  $\pm$  8 mg/dl), 50 mg/kg body weight (245  $\pm$  9 mg/dl) and 100 mg/kg body weight (301  $\pm$  7 mg/dl) compared to diabetic rats (355  $\pm$  9 mg/dl, 352  $\pm$  6 mg/dl and 360  $\pm$  9 mg/dl respectively).

Oral administration of all the doses of the test drug extract through 5; 10 and 15 days on blood glucose and body weight were given in Tables 3 and 4. The blood glucose level of diabetic rats group significantly decreased from 355  $\pm$  10 mg/dl to 225  $\pm$  9 mg/dl on days 5 after treated with BGF extract. Oral administration of BGF (25, 50 and 100 mg/kg body weight) through long duration decreased blood glucose as compared to diabetic rats. Continuous administration of BGF extract was found to significantly (p<0.05) decrease the blood glucose level to 140 mg/dl. The body weight was lowered in diabetic rats as compared to normal rats. But when the animals were treated with BGF extract increased body weight as compared to diabetic rats. BGF extract (50 mg/kg body weight) increased body weight nearly as the normal compared to diabetic rat.

The data of the present work clearly demonstrated that, there was a significant increase in the mean value of sAST (71  $\pm$  3 U/ml) and sALT (76  $\pm$  4 U/ml) of the diabetic rats compared to that of the normal group (Tables 5 and 6). On the other hand, administration of BGF extract changes the sAST and sALT values to approximately control value.

For all parameters studies BGF at doses 25, 50 and 100 mg/kg body weight showed significant effect. But BGF at 50 mg/kg body weight showed highest effect. The optimum dose of bitter gourd extract given to diabetic rat 50 mg/kg body weight.

### Discussion

Alloxan has been observed to cause reduction of the beta cell of islets of Langerhans and induced hyperglycemia. The normal beta cells count was markedly reduce after alloxan treatment [24-29]. The data revealed that the subcutaneous injection of alloxan in a dose of 120 mg/

Groups	Treatment	Mean AST U/ml			
		Initial	5 days	10 days	15 days
I	Normal	32 ± 2	33 ± 3	33 ± 2	33 ± 2
II	Diabetic	71 ± 3	70 ± 3	70 ± 3	72 ± 2
III	BGF 25 X	66 ± 3	62 ± 2	58 ± 3	55 ± 3
IV	BGF 50 X	65 ± 4	52 ± 3	45 ± 3	41 ± 2
V	BGF 100 X	67 ± 3	58 ± 4	52 ± 3	47 ± 2

Table 5: Effect of BGF extract in AST in diabetic rats.

Groups	Treatment	Mean ALT U/ml			
		Initial	5 days	10 days	15 days
I	Normal	38 ± 3	39 ± 3	38 ± 3	39 ± 3
II	Diabetic	76 ± 4	79 ± 4	82 ± 4	83 ± 4
III	BGF 25 X	76 ± 2	71 ± 3	70 ± 4	62 ± 3
IV	BGF 50 X	77 ± 4	62 ± 4	58 ± 3	50 ± 2
V	BGF 100 X	77 ± 3	68 ± 3	60 ± 3	54 ± 2

Result Mean ± S.D  
Significant p<0.05  
X mg/kg body weight

Table 6: Effect of BGF extract in ALT in diabetic rats.

kg body weight was a convenient dose for the induction of diabetes in rats. These results are in agreement with those reported data by Eskander and Won Jun [9] Sheweta et al. [25] who used 120 mg/kg body weight to produce diabetic rats.

The result of the present study indicates that BGF decrease blood glucose in alloxan diabetic rats. The possible mechanism include the stimulation of β-cells and release of insulin and activation of the insulin receptors. Alloxan has been shown to induce free radical production and cause tissue injury [28]. The pancreas is especially susceptible to the action alloxan induced free radical damage. Other possible mechanism BGF extract can act as a free radical scavenger. Administration of BGF extracts reduction the activity of the liver enzyme. The decrease in the concentration of enzymes in alloxan treated rats given BGF may be as hepatoprotective agent and improvement of liver function. In the present study it was observed the body weight was decrease in alloxan diabetic rats [26] and administration of BGF extract increases body weight on alloxan diabetic. The possible mechanism the ability of BGF to protect body weight loss seems to be as a result of its ability to reduce hyperglycemia [30-32].

The isolation and structure determination of these compounds are currently in progress. The present study indicates that BGF possessed a significant hypoglycaemic effect and may be as hepatoprotective agent

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