

# Effects of *Cinnamomum zeylanicum* (Ceylon cinnamon) on blood glucose and lipids in a diabetic and healthy rat model

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## ABSTRACT

**Objectives:** To evaluate short- and long-term effects of *Cinnamomum zeylanicum* on food consumption, body weight, glycemic control, and lipids in healthy and diabetes-induced rats.

**Materials and Methods:** The study was conducted in two phases (Phase I and Phase II), using Sprague-Dawley rats in four groups. Phase I evaluated acute effects on fasting blood glucose (FBG) (Groups 1 and 2) and on post-oral glucose (Groups 3 and 4) blood glucose. Groups 1 and 3 received distilled-water and Groups 2 and 4 received cinnamon-extracts. Phase II evaluated effects on food consumption, body weight, blood glucose, and lipids over 1 month. Group A ( $n = 8$ , distilled-water) and Group B ( $n = 8$ , cinnamon-extracts) were healthy rats, while Group C ( $n = 5$ , distilled-water) and Group D ( $n = 5$ , cinnamon-extracts) were diabetes-induced rats. Serum lipid profile and HbA1c were measured on D-0 and D-30. FBG, 2-h post-prandial blood glucose, body weight, and food consumption were measured on every fifth day. **Results:** Phase I: There was no significant difference in serial blood glucose values in cinnamon-treated group from time 0 ( $P > 0.05$ ). Following oral glucose, the cinnamon group demonstrated a faster decline in blood glucose compared to controls ( $P < 0.05$ ). Phase II: Between D0 and D30, the difference in food consumption was shown only in diabetes-induced rats ( $P < 0.001$ ). Similarly, the significant difference following cinnamon-extracts in FBG and 2-h post-prandial blood glucose from D0 to D30 was shown only in diabetes-induced rats. In cinnamon-extracts administered groups, total and LDL cholesterol levels were lower on D30 in both healthy and diabetes-induced animals ( $P < 0.001$ ). **Conclusions:** *C. zeylanicum* lowered blood glucose, reduced food intake, and improved lipid parameters in diabetes-induced rats.

**Key words:** Blood glucose, Ceylon cinnamon, *Cinnamomum zeylanicum*, diabetes mellitus, lipids, Sprague-Dawley rats

## INTRODUCTION

Diabetes mellitus is a leading cause of morbidity and mortality worldwide, with an estimated 80% of the world population with diabetes living in developing countries.<sup>[1]</sup> Most patients with the disease have type 2 diabetes to which the South Asians are known to have an increased predisposition.<sup>[2]</sup> The causes of type 2 diabetes are multi-factorial, and the diet plays an important role on its' incidence, severity, and management.<sup>[3]</sup> Hence, studies have

frequently focused on dietary components beneficial in the prevention and treatment of diabetes. Recent studies have demonstrated that herbal products have beneficial effects in patients with diabetes by improving glucose and lipid metabolism, antioxidant status, and capillary function.<sup>[4]</sup> Cinnamon is one such a dietary component that has shown to have biologically active substances with insulin-mimetic properties. *In vitro*<sup>[5,6]</sup> and *in vivo*<sup>[7,8]</sup> studies have shown that cinnamon enhances glucose uptake by activating the insulin receptor kinase activity, auto-phosphorylation of the insulin receptor, and glycogen synthase activity.

Cinnamon, the inner bark of a tropical evergreen tree has two main types, Ceylon cinnamon (*Cinnamomum zeylanicum*

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Blume) and Chinese Cassia (*Cinnamomum aromaticum* Ness) and which when dried, rolls into a tubular form known as a quill. Cinnamon is available in either its whole quill form (cinnamon sticks) or as ground powder. In native Ayurvedic medicine, cinnamon is considered a remedy for respiratory, digestive, and gynecological ailments. Recent studies emerging from western countries have shown many potentially beneficial health effects of cinnamon such as anti-inflammatory properties, anti-microbial activity, blood glucose control, reducing cardiovascular disease, boosting cognitive function, and reducing risk of colonic cancer.<sup>[9-11]</sup> Sri Lanka is a middle-income country in the South Asian region produces the largest quantity of *C. zeylanicum* and the best quality cinnamon. *C. zeylanicum*, also known as Ceylon cinnamon (the source of its Latin name, zeylanicum) or 'true cinnamon', is indigenous to Sri Lanka. One important difference between 'true' cinnamon and the 'cassia' cinnamon is their coumarin content.<sup>[12]</sup> Coumarins are naturally occurring plant compounds with strong anticoagulant properties. The coumarin content in Ceylon cinnamon appears to be very small to cause health risks, whereas the coumarin level in *C. aromaticum* appears to be much higher and may pose health risks if consumed in higher quantity on a regular basis.<sup>[12]</sup> In addition, coumarins also have potentially toxic effects on the liver.<sup>[13]</sup>

Previous studies have explored the anti-diabetic effects of Cinnamon cassia (*C. aromaticum*) extract *in vivo* and *in vitro*.<sup>[5-8]</sup> However, there is a relative lack of knowledge regarding the effects of the indigenous species of Sri Lankan Cinnamon (*C. zeylanicum*) on blood glucose and lipids. The aim of this study was to evaluate the short- and medium-term effects of *C. zeylanicum* extracts on food consumption, body weight, blood glucose and glycemic control, and lipids in both healthy and diabetes-induced rats as an animal model.

## MATERIALS AND METHODS

The study was conducted in two phases from June to August 2010. Ethical approval was obtained from the Ethics Review Committee of the Faculty of Medicine, University of Colombo, Sri Lanka.

### Preparation of cinnamon extract

The cinnamon extracts were prepared at the Industrial Technology Institute (ITI), Colombo, Sri Lanka. A cinnamon sample was obtained from the collection center of S.D.S. Spices (Pvt) Ltd, Kosgodda, Sri Lanka. The voucher specimen is deposited at the Herbal Technology Section of Industrial Technology Institute under voucher specimen number HTS-CIN-01 (Ham), and was identified by Dr Sirimal Premakumara. The cinnamon was extracted to distilled water from the stem barks of *C. zeylanicum* using

a Soxhlet apparatus and the resulting hot water extract was freeze-dried to obtain crude water extract (yield 8.3% w/w dry weight basis). This solid freeze-dried extract in a dosage form in distilled water was administered to experimental rats. The dosage for rats was calculated according to the body surface area as shown below, using the human dosage (6 g) obtained from previous studies.<sup>[9]</sup> The daily cinnamon dose given for a 200 g rat was 120 mg (600 mg/kg) in 1 ml of distilled water administered via a metal oro-gastric tube.  $\text{Dose (mg/kg)} = \text{Human dose (mg/kg)} \times [\text{Human } Km \text{ factor/Rat } Km \text{ factor}]^{[14]}$

### Phase I

The aim of Phase I was to determine the short-term effects of *C. zeylanicum* on blood glucose in the unfed and post-prandial state. Healthy Sprague-Dawley rats having a mean body weight of  $190 \pm 25$  g and aged 3–4 months ( $n = 32$ , male:female = 1:1) were used as the animal model. They were divided into four equal groups of eight animals each. In Groups 1 and 2 blood glucose levels in the unfed state were measured, while in Groups 3 and 4 blood glucose levels following a standard oral glucose load (1.25 g/kg) was measured.

In all four groups, the animals were kept fasting for 12 h overnight. In Groups 1 and 2 the fasting blood glucose (FBG) value was determined at '0' h and the cinnamon extract and distilled water, respectively, were administered immediately. Subsequently the blood glucose levels in the unfed state were measured at 0.5, 1, 2, 4, 8, 12, and 24 h. Similarly in Groups 3 and 4 the FBG values were determined at '0' h and an oral glucose load was administered. The cinnamon and distilled water were administered after 0.5 h following the glucose load with measuring of blood glucose values at that time. Subsequently blood glucose levels were measured at 1, 2, 4, 8, 12 and 24 h. In all four groups, the animals were fed at 24 h and kept fasting for 12 h for repetition of this cycle the next day. This protocol was carried out for three consecutive cycles and mean blood glucose values were analysed. The percentage reductions of the blood glucose values of each hour from the respective FBG of Groups 1 and 2 were compared. The percentage reductions were calculated as follows:

$$\text{Percentage reduction} = \frac{(\text{FBG} - \text{Blood glucose value})}{\text{FBG}} \times 100$$

### Phase II

The aim of Phase II was to determine the medium-term effects of *C. zeylanicum* on food consumption, body weight, blood glucose and glycemic control, and lipids. Group A ( $n = 8$ ) and Group B ( $n = 8$ ) were healthy Sprague-Dawley rats, while Group C ( $n = 5$ ) and Group D ( $n = 5$ ) were

diabetes-induced Sprague-Dawley rats. In groups A and B distilled water and cinnamon were administered, respectively, on a daily basis for a period of 1 month. Similarly, in Groups C and D distilled water and cinnamon were administered, respectively, for 1 month. In all four groups, the animals were kept fasting overnight for a period of 12 h, and cinnamon or distilled water were administered, followed by a 30 min interval prior to commencing of the standard daily pellet feed. In all four groups' blood for estimation of serum total cholesterol, HDL cholesterol, LDL cholesterol, triglycerides, and HbA1c were drawn on day 0 and day 30 from the tail vein of the rats. FBG and 2 h post-prandial blood glucose were measured from day 0 every fifth day till day 30.

### Induction of diabetes

In Groups C and D ( $n = 16$ ), diabetes was induced by a single intravenous injection of streptozotocin (Sigma Chemicals Co, St. Louis, MO, USA) to the tail vein at a dose of 75 mg/kg body weight. Confirmatory FBG tests were performed after 14 days of injection and animals with a FBG of  $>200$  mg/dl (six females and four males) after induction were randomly allocated to Groups C ( $n = 5$ ) and D ( $n = 5$ ) with an equal gender distribution.

### Measurements: Biochemical and physiological

The blood glucose values were determined using an Optium Xceed glucometer and strips (Abbott Diabetes Care Ltd, Oxon, UK). A cross-validation to establish the accuracy of the glucometer was done by comparing with 10 laboratory plasma glucose tests. Blood for estimation of lipid parameters were collected to plain tubes while for estimation of HbA1c blood was collected to K3 EDTA tubes. HbA1c was determined by the HPLC Bio-Rad variant method (Bio-Rad, USA), cholesterol by the CHOD-PAP method (Bio System S.A., Barcelona, Spain), HDL cholesterol by an enzymatic colorimetric direct HDL assay (Analyticon Biotechnologies AG, Lichtenfels, Germany), LDL cholesterol by a homogeneous enzymatic direct LDL assay (Analyticon Biotechnologies AG, Lichtenfels, Germany) and triglycerides by the GPO-PAP method (Bio System S.A., Barcelona, Spain).

During the study period, the animals were fed with a standard pellet diet and water daily with added vitamins as required. The animals were housed in separate cages in a barrier procedure room at ambient temperature with a relative humidity of 40–60% and a 12 h light/dark cycle. The daily food and water consumption was measured, while body weight was measured on every fifth day.

### Determination of the effects on organs

On completion of Phase II, a histo-pathological study was conducted on 3 animals from each group. The selected

animals were dissected under ether anesthesia by a trained researcher and the liver (10% formalin preservative), both kidneys (modified Bouin's solution) and pancreas (10% formalin preservative) were harvested to determine the effects of cinnamon on these organs. The liver was examined for spotty/zonal liver cell necrosis, hepatocyte swelling, fatty changes, and inflammation. In the kidneys, tubular epithelial cell damage, presence of casts/crystals, glomerular changes, and inflammation were examined. The pancreas was examined for cell necrosis, inflammation, islet cell hyperplasia, and/or atrophy.

### Statistical analysis

Data from the experiments are presented as mean  $\pm$  standard deviation. Statistical analysis was performed using the Statistical Package for Social Science (SPSS) software for windows version 14 (SPSS Inc., Chicago, Illinois, USA). In all statistical analysis a  $P$  values of  $<0.05$  were considered significant.

## RESULTS

### Phase I

The mean body weight of animals in Groups 1–4 were  $185 \pm 12$  g,  $192 \pm 8$  g,  $180 \pm 14$  g and  $188 \pm 10$  g, respectively. There was no statistically significant difference in the mean body weights of the respective groups. There were no statistically significant differences observed between the mean blood glucose values at each hour of Groups 1 and 2 [Table 1]. When comparing the percentage reductions of the blood glucose values of each hour from the respective FBG of Groups 1 and 2 only the cinnamon administered group demonstrated a greater mean percentage reduction at 0.5, 1 and 2 h. However, this difference was not statistically significant [Figure 1].

When a standard loading dose of glucose was administered to Groups 3 and 4, in both groups the peak plasma glucose

**Table 1: Effect of *Cinnamomum zeylanicum* on fasting blood glucose and post-oral glucose load**

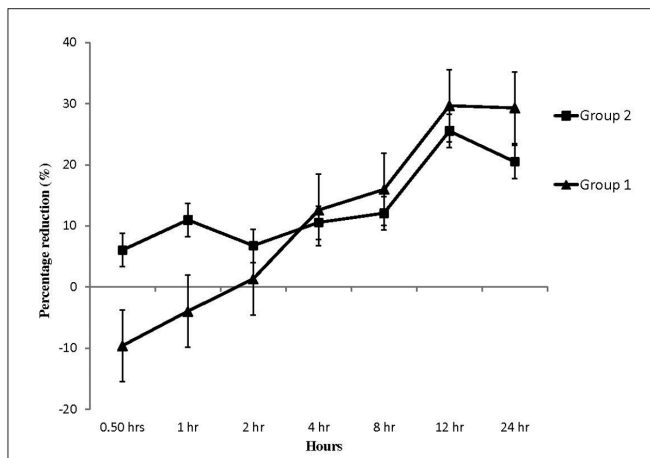
Time (h)	Mean blood glucose ( $\pm$ SD) mg/dl			
	Fasting		Post-oral glucose load	
	Group 1 (Distilled water)	Group 2 (Cinnamon)	Group 3 (Distilled water)	Group 4 (Cinnamon)
0	92 ( $\pm$ 9)	96 ( $\pm$ 11)	96 ( $\pm$ 11)	93 ( $\pm$ 10)
0.5	101 ( $\pm$ 18)	90 ( $\pm$ 19)	<sup>a</sup> 149 ( $\pm$ 27)*	<sup>a</sup> 190 ( $\pm$ 39)*
1	96 ( $\pm$ 24)	86 ( $\pm$ 14)	137 ( $\pm$ 14)	137 ( $\pm$ 48)
2	91 ( $\pm$ 11)	90 ( $\pm$ 14)	99 ( $\pm$ 21)	109 ( $\pm$ 37)
4	81 ( $\pm$ 12)	86 ( $\pm$ 10)	88 ( $\pm$ 8)	80 ( $\pm$ 11)
8	85 ( $\pm$ 11)	78 ( $\pm$ 11)	<sup>a</sup> 79 ( $\pm$ 5)*	<sup>a</sup> 69 ( $\pm$ 7)*
12	65 ( $\pm$ 8)	72 ( $\pm$ 4)	<sup>a</sup> 74 ( $\pm$ 4)*	<sup>a</sup> 65 ( $\pm$ 8)*
24	65 ( $\pm$ 11)	76 ( $\pm$ 10)	71 ( $\pm$ 8)	67 ( $\pm$ 10)

\* $P < 0.05$ , <sup>a</sup>Comparison between Groups 3 and 4

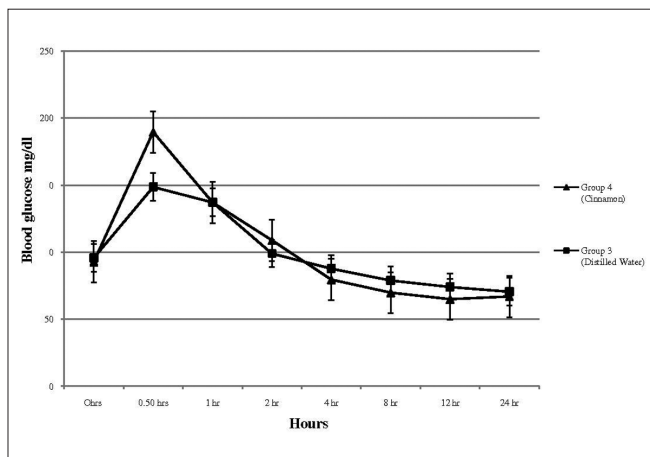
following the standard oral glucose load was recorded at 0.5 h. When the test substance (cinnamon or distilled water) was administered to the respective groups at 0.5 h, Group 4 (cinnamon) demonstrated a faster decline in blood glucose in comparison to Group 3 (distilled water) [Figure 2]. The mean percentage reduction in blood glucose from 0.5 to 1 h in Groups 3 and 4 were 7.7% and 27.7%, respectively, with the cinnamon administered group demonstrating a significantly higher percentage reduction ( $P < 0.001$ ). In addition, the cinnamon group also demonstrated a lower blood glucose value at 8 and 12 h post-oral glucose load ( $P < 0.05$ ) [Table 1].

### Phase II

In healthy animals the mean body weight ( $\pm$ SD) of the cinnamon group and control group on day 0 was  $193.9 \pm 9.2$  g and  $191.6 \pm 8.6$  g, respectively. At the end of the study period (D-30), the mean weight of the control group



**Figure 1:** Percentage reduction of hourly blood glucose values from fasting blood glucose in Groups 1 (distilled water) and 2 (cinnamon extract)

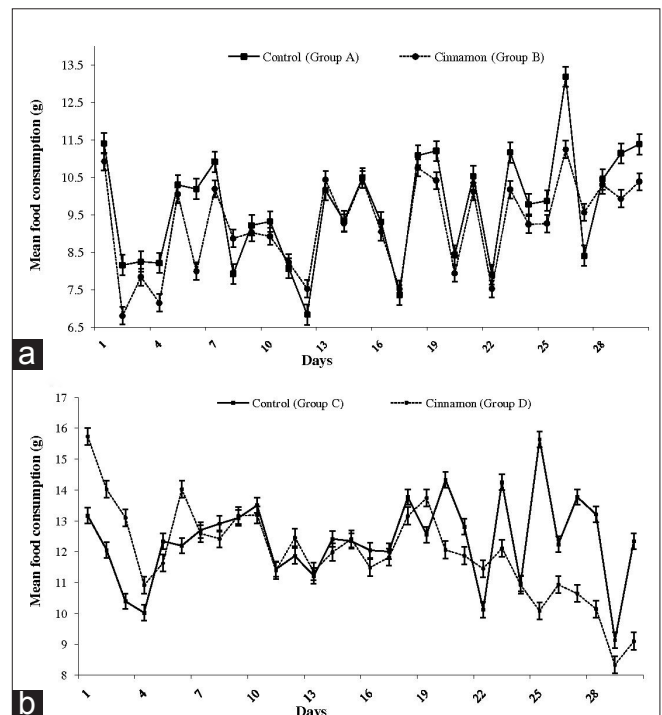


**Figure 2:** Blood glucose values following post-oral glucose load in Groups 3 (distilled water) and 4 (cinnamon extract)

increased significantly ( $P < 0.001$ ) to  $210.0 \pm 16.8$  g, while in the cinnamon group the weight gain was not significant ( $202.5 \pm 15.7$  g,  $P > 0.05$ ). The mean food consumption of healthy rats on day 0 in the cinnamon and control groups was  $10.9 \pm 1.2$  g and  $11.4 \pm 1.9$  g, respectively, whereas on day 30 it was  $10.3 \pm 1.5$  g and  $11.4 \pm 1.3$  g, respectively, indicating no significant difference in the mean food consumption between day 0 and day 30 [Figure 3a].

The diabetes-induced animals had mean body weights of  $176.5 \pm 13.9$  g in the cinnamon group and  $169.9 \pm 6.5$  g in the control group at the beginning of the study (day 0). On day 30, the mean body weight of the cinnamon group was  $168.8 \pm 33.4$  g and in the control group it was  $163.1 \pm 17.6$  g. The weight reduction observed in both groups during the study period was not statistically significant. The mean food consumption per animal had significantly reduced from day 0 to day 30 in the cinnamon group, but not in the control group of diabetes-induced animals ( $P < 0.001$ ) (cinnamon group D0:  $15.7 \pm 1.2$  g, D30:  $9.1 \pm 0.5$  g; control group D0:  $13.2 \pm 1.6$  g, D30:  $12.3 \pm 1.3$  g) [Figure 3b].

In healthy animals, there was no significant difference in the FBG or 2-h post-prandial blood glucose between the cinnamon and control groups from day 0 to day 30. However, in the diabetes-induced animals the FBG was significantly lower in the cinnamon group than the control



**Figure 3:** Mean food consumption of animals during the study period, (a) in healthy animals and (b) diabetes-induced animals



group on day 20 and day 30, while the 2-h post-prandial glucose was significantly lower on day 30 in the cinnamon group [Table 2]. In both healthy and diabetes-induced animals, the cinnamon group showed a significantly lower total and LDL cholesterol level on day 30 compared to day 0. However, this difference was not observed in the control groups [Table 2]. The HDL cholesterol and triglycerides remained unchanged in all four groups of animals in Phase II. The mean HbA1C remained unchanged in healthy animals in day 0 and day 30, while in diabetes-induced animals the mean HbA1C in the control group increased significantly from 2.2% ( $\pm 0.1$ ) in day 0 to 4.5% ( $\pm 0.9$ ) on day 30 ( $P < 0.001$ ). In the cinnamon group of diabetes-induced animals, the mean HbA1C on day 0 and day 30 was not significantly different [Table 2]. The histological analysis of the liver, kidney, and pancreas were within normal histological limits with no evidence indicative of toxicity. The liver histology demonstrated no evidence of spotty/zonal liver cell necrosis, hepatocyte swelling, fatty changes, and inflammation. In the kidneys, tubular epithelial cell damage, presence of casts/crystals, glomerular changes, and inflammation were not seen. The pancreas demonstrated no evidence of cell necrosis,

inflammation, islet cell hyperplasia, and/or atrophy.

## DISCUSSION

To our knowledge, this is the first report on the effects of authentic Ceylon cinnamon collected from native cultivators on blood glucose, although there are studies on *C. zeylanicum*.<sup>[10]</sup> Our results demonstrate that *C. zeylanicum* improves the body's ability to handle a glucose load as demonstrated by the rapid metabolism shown in the cinnamon group of healthy animals following a standard oral glucose load. This probably occurs as a result of the ability of cinnamon to potentiate insulin-regulated glucose utilization *via* enhancing insulin signaling.<sup>[5,8,15]</sup> In the study by Qin *et al.*,<sup>[8]</sup> enhanced cellular glucose uptake due to the stimulation of the insulin receptor activity by the increasing concentrations of the phosphorylated intracellular protein IRS-1 and the increasing of PI 3-kinase has been shown with cinnamon. Studies have shown that this potentiating effect of cinnamon on insulin causes a dose-dependent reduction in the serum insulin concentration.<sup>[16]</sup> This rapid reduction in blood glucose concentration and the reduced insulin requirement could help prevent  $\beta$ -cell destruction

**Table 2: Effect of *Cinnamomum zeylanicum* on blood glucose, lipids, and HbA1C in healthy and diabetes-induced rats**

Biochemical test	Mean ( $\pm$ SD) mg/dl			
	Healthy rats		Diabetes-induced rats	
	Group 1 (Distilled water)	Group 2 (Cinnamon)	Group 3 (Distilled water)	Group 4 (Cinnamon)
<i>Fasting blood glucose</i>				
Day 0	72 ( $\pm 6$ )	70 ( $\pm 8$ )	296 ( $\pm 20$ )	271 ( $\pm 42$ )
Day 10	81 ( $\pm 8$ )	83 ( $\pm 9$ )	322 ( $\pm 36$ )	290 ( $\pm 66$ )
Day 20	75 ( $\pm 7$ )	76 ( $\pm 13$ )	<sup>b</sup> 372 ( $\pm 74$ )*	<sup>b</sup> 254 ( $\pm 28$ )*
Day 30	77 ( $\pm 8$ )	77 ( $\pm 10$ )	<sup>b</sup> 320 ( $\pm 53$ )*	<sup>b</sup> 247 ( $\pm 63$ )*
<i>Post-prandial blood glucose</i>				
Day 0	106 ( $\pm 15$ )	102 ( $\pm 19$ )	395 ( $\pm 63$ )	378 ( $\pm 72$ )
Day 10	101 ( $\pm 13$ )	103 ( $\pm 8$ )	403 ( $\pm 108$ )	346 ( $\pm 110$ )
Day 20	99 ( $\pm 11$ )	96 ( $\pm 9$ )	419 ( $\pm 52$ )	384 ( $\pm 115$ )
Day 30	94 ( $\pm 14$ )	90 ( $\pm 9$ )	<sup>b</sup> 493 ( $\pm 21$ )*	<sup>b</sup> 369 ( $\pm 111$ )*
<i>Total cholesterol</i>				
Day 0	68.4 ( $\pm 12.3$ )	<sup>a</sup> 74.5 ( $\pm 9.5$ )*	69.7 ( $\pm 11.8$ )	<sup>a</sup> 64.3 ( $\pm 9.8$ )*
Day 30	67.3 ( $\pm 10.3$ )	<sup>a</sup> 59.2 ( $\pm 13.2$ )*	72.0 ( $\pm 15.2$ )	<sup>a</sup> 50.8 ( $\pm 14.7$ )*
<i>LDL cholesterol</i>				
Day 0	12.3 ( $\pm 8.6$ )	<sup>a</sup> 13.6 ( $\pm 6.2$ )*	( $\pm 4.6$ )	<sup>a</sup> 15.9 ( $\pm 3.0$ )*
Day 30	8.8 ( $\pm 3.8$ )	<sup>a</sup> 3.0 ( $\pm 1.3$ )*	12.9 ( $\pm 18.8$ )	<sup>a</sup> 3.1 ( $\pm 0.9$ )*
<i>HDL cholesterol</i>				
Day 0	40.2 ( $\pm 8.0$ )	45.4 ( $\pm 10.7$ )	25.4 ( $\pm 11.8$ )	32.0 ( $\pm 9.5$ )
Day 30	51.6 ( $\pm 8.1$ )	46.5 ( $\pm 11.5$ )	19.5 ( $\pm 5.3$ )	26.9 ( $\pm 4.6$ )
<i>Triglycerides</i>				
Day 0	78.9 ( $\pm 21.3$ )	77.6 ( $\pm 16.2$ )	197.0 ( $\pm 87.5$ )	121.4 ( $\pm 54.9$ )
Day 30	72.8 ( $\pm 33.1$ )	59.7 ( $\pm 20.5$ )	198.2 ( $\pm 80.1$ )	131.0 ( $\pm 76.9$ )
<i>HbA1C</i>				
Day 0	1.9 ( $\pm 0.1$ )	2.3 ( $\pm 0.5$ )	<sup>a</sup> 2.2 ( $\pm 0.1$ )*	4.2 ( $\pm 2.2$ )
Day 30	3.0 ( $\pm 0.3$ )	3.2 ( $\pm 0.3$ )	<sup>a</sup> 4.5 ( $\pm 0.9$ )*	4.9 ( $\pm 1.4$ )

\*  $P < 0.05$ , <sup>a</sup>Comparison between days 0 and 30, <sup>b</sup>Comparison between groups 3 and 4

occurring as a result of  $\beta$ -cell exhaustion from chronic hyperglycemia in diabetes.<sup>[17]</sup>

The cinnamon group of diabetes-induced animals demonstrated a significant reduction in mean food consumption implying that cinnamon also tends to promote satiety. In addition with long-term usage in diabetes-induced animals the cinnamon group demonstrated a lower FBG (on day 20 and 30) and 2-h post-prandial blood glucose (on day 30). These results are compatible with results from previous studies.<sup>[9,18]</sup> However, the same result was not observed in healthy animals where normal glucose homeostatic mechanisms are intact indicating that the effects of cinnamon are under the regulation of normal physiological feedback mechanisms. The fact that cinnamon lowers blood glucose usually within physiological levels without hypoglycemia and increases satiety leads to the hypothesis that it may act by potentiating the effects of incretin hormones.<sup>[19]</sup>

The cinnamon extracts also promoted a better glycemic control in diabetes-induced animals as demonstrated by the stable HbA1c in the cinnamon group as opposed to the diabetic control group. According to the United Kingdom Prospective Diabetes Study (UKPDS), a drop of HbA1C from 7.9% to 7% lowers the risk of macro- and micro-vascular disease significantly;<sup>[20]</sup> thus, a better control of HbA1c levels in patients with cinnamon supplementation might be expected to yield similar beneficial effects on the long-term complications of diabetes. In addition, both in healthy and diabetes-induced animals, cinnamon caused a very significant reduction in the total and LDL cholesterol levels. This cholesterol lowering effect of cinnamon together with its antioxidant properties<sup>[21]</sup> has the potential to reduce the long-term cardiovascular complications associated with diabetes,<sup>[22]</sup> while also being effective as a supplement in patients with hyperlipidemia and cardiovascular diseases without diabetes.

The usage of cinnamon as a regular supplement with meals was not advocated or the daily dosage was restricted in many countries due to the toxic effects of *C. aromaticum* on the liver and coagulation.<sup>[23]</sup> In contrast, *C. zeylanicum* has shown to contain a lesser content of coumarin<sup>[11,24]</sup> and thus it may be possible that Ceylon cinnamon could be used in higher doses without toxic effects for longer durations. In addition our pathological analysis also revealed no evidence indicative of toxicity of *C. zeylanicum*. Our study has several limitations, the study was conducted with a single dose of cinnamon the dose-dependent effect of cinnamon extracts on the serum insulin level was not evaluated. The serum insulin levels that were measured in Phase II were <2 mIU/l in all animals (the sensitivity limit of the assay).

The study involved both healthy and diabetes-induced male and female rats, the difference in the body composition and lipid profile between genders were not considered during analysis, however since all comparative groups had equivalent numbers of males and females this effect is likely to be minimal.

## CONCLUSION

In conclusion, *C. zeylanicum* lowered blood glucose, reduced food intake, and reduced atherogenic LDL cholesterol. Its effect on humans and the possibility of incretin mimetic effects need further study.

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