Comparison the effect of Various Cinnamon plant Extracts with Metformin in Blood Glucose level of alloxan-induced diabetic laboratory rats

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Abstract:
The present study aims to compare the hypoglycemic activity of blood sugar of three types of cinnamon plant extract. Which is methanol, hexane and chloroform extract with metformin drug which is used for type 2 diabetic mellitus in laboratory rats of . The study showed presence of significant $p \geq 0.05$ hypoglycemic activity in all cinnamon plant extracts compared with control group. The result also showed highly significant $p \geq 0.05$ hypoglycemic activity of hexane extract compared with metformin drug than other extracts after (4, 6, 9)hr of the treatment.

Introduction:
Diabetes mellitus (DM) is a serious health problem with high rates of incidence and mortality. It is a serious endocrine syndrome. DM is characterized by elevated plasma glucose concentrations and discharge of large amount of sugar in urine by the patient resulting from relative or absolute insulin deficiency, insulin resistance, or both, leading to metabolic abnormalities in carbohydrates, lipids and proteins. (1,2,3). DM is a disorder that cannot be cured, but can only be managed. In spite of tremendous
progress in the management of diabetes using synthetic drugs, potential new in expensive treatments should be used to reduce global morbidity and mortality, as most of the people with diabetes lives in areas of the world, where existing treatments are unavailable or are too expensive. It is well documented that insulin sensitivity can be modulated by various dietary compounds and exercise regimes (4,5,6). Despite important progress in the management of diabetes using synthetic drugs, many traditional plant treatments are still used throughout the world. (7). However, few traditional antidiabetic plants have received proper scientific validation. derivatives have hypoglycemic properties are used in folk medicine and traditional healing systems around the world (8). Many pharmaceuticals used in modern medicine are also of natural, plant origin (9). Cinnamon is amongst the world’s oldest and most frequently consumed spices, and is used as an herbal remedy (10). The genus Cinnamomum consists of 250 species of aromatic evergreen trees and shrubs, primarily located in Asia and Australia. The term Cinnamomum is derived from Greek kinnamomon, meaning “sweet wood”. Cinnamon is classified in the botanical division: Magnoliophyta, class: Magnoliopsida, order: Magnoliales and family: Lauraceae. The cinnamon of commerce is the dried inner stem-bark of a small evergreen tree 10-15 meters tall. It is native to tropical southern India and Srilanka. There are two types of cinnamon, common cinnamon (vernacular name: dalchini) or true cinnamon (Cinnamomum zeylanicum, C. verum) and cassia (Cinnamomum aromaticum). Cinnamon has been used for centuries, as flavor modifiers to make food more palatable. Its ingredients impart characteristic flavor and spicy aroma to food(11). The most constituent of commercial importance is the volatile oil. Volatile components are present in all parts of cinnamon and can be classified broadly into monoterpenes, sesquiterpenes and phenylpropenes. Cinnamaldehyde (more precisely trans-cinnamaldehyde or 3-phenyl-2-propenal) is the main constituent in cinnamon bark oil, whereas, that of leaf oil is eugenol (12). Extensive investigation in recent years suggests that cinnamon possess numerous pharmacological activities including reported to possess potent antioxidant (13;14), antimicrobial (15), and antipyretic (16) properties. Much attention has also been paid to the influence of cinnamon on insulin action, which may provide benefits for diabetic patients. Interest in cinnamon as a potentially useful treatment for type-2 diabetes began almost 20 years ago (17). Since that time, numerous in vitro and in vivo studies have elucidated cinnamon’s effect on insulin signal transduction (18, 19,20,21). Most experiments claimed that cinnamon is a natural...
insulin sensitizer (22) and an inhibitor of advanced glycation endproducts(23). The aim of the present work is to determine the hypoglycemic effect cinnamon extract and compared with metformin which is hypoglycemic drug in hyperglycemic induced laboratory rats.

Materials and Methods:
1. Plant extraction:
The plant material was of cinnamon brought from the local market. The bark of the plant was powdered and extracted in Soxhlet apparatus with arrange of solvents, starting with hexane and chloroform (to separate lipids and terpenoids if present that otherwise may precipitate on the wall of the flask) then we used methanol to extract the other constituents of cinnamon then each extract was dried and collected.(24).

2. Animals:
75 Adult Male albino rats weighing 150-200 g were used in the present study. All rats were kept at room temperature. They were fed with standard rat pellet diet and provided water ad libitum. The animal were treated with 300 mg /kg and this dose was selected after a series of primary experiments.

3-Alloxan-induced diabetes:
The rats weighing 150-200 g were allowed to fast for 24 hours prior to experimentation and rendered diabetic by a single dose of intraperitoneal injection of alloxan 150 mg/kg body weight dissolved in normal saline (25) After 18 hours of injection of alloxan, diabetes was confirmed by testing blood sugar. The level more than 200 mg/dl were selected for the further study. then the animal were divided into the following groups each with 15 rats and treated the plant extract orally using stomach tube.(26).

Group 1: rats treated with 300 mg /kg of methanolic extract mg/kg.
Group 2: rats treated with 300 mg /kg of chloroform extract.
Group 3: rats treated with 300 mg /kg of hexane extract.
Group 4: rats treated with normal saline as control group
Group 5: rats treated with 650mg/kg of metformin drug(27).

4-blood sampling:
Blood samples from rats were collected by direct heart punctu Serum glucose level was measured by using (glucose enzymatic colorimetric test kit) from Biocon Diagnostik (Germany) Blood glucose was measured at , 3, 6 and 9hr (28).

Results:
The present result showed significant $p\geq0.05$ decreasing in blood glucose level in the group treated with methanol and metformin drug compared with control group (normal saline ) fig(1).With in the period of time the result showed significant $p\geq0.05$ decreasing in blood glucose level in the group treated with methanol and metformin drug compared with control group. While there are no significant
decreasing in the group treated with metformin and methanol extract in the 3 and 6hr period also there is significant p≥0.05 decreasing in 9hr period between metformin and methanol group. Fig(2) the result showed significant p≥0.05 decreasing in blood glucose level in the group treated with hexane and metformin drug compared with control group. With in the period of time the result showed significant p≥0.05 decreasing in blood glucose level in the group treated with hexane and metformin drug compared with control group. Also there are highly significant p≥0.05 decreasing in the group treated with hexane compared with metformin drug in the 6 and 9hr period. the result in fig.(3) showed significant p≥0.05 decreasing in blood glucose level in the group treated with chloroform extract and metformin drug compared with control group. With in the period of time the result showed significant p≥0.05 decreasing in blood glucose level in the group treated with chloroform and metformin drug compared with control group. While there are significant p≥0.05 decreasing in the group treated with metformin compared with chloroform extract in the 3,6,9hr period. Fig (4) showed the comparison between the hypoglycemic activity of metformin drug and methanol,hexane,and chloroform of cinnamon extract it is appeared from the fig(4) significant p≥0.05 decreasing in blood glucose level in the group treated with hexane extract compared with metformin drug and methanol or chloroform extract within the period there is significant decreasing in the group treated with hexane extract compared metformin drug and methanol or chloroform extract specially in 9hr period.

Discussion:

The present study has detected the antidiabetic effect of different extract of Cinnamomum bark in alloxan induced diabetic rats. And compared with the hypoglycemic drug metformin. Intraperitonial injection of alloxan monohydrate caused diabetes mellitus in adult male rats. Results of the present study showed that diabetic rats exhibited a significant increase in blood glucose level. This result is agree with other studies in rats (29, 30, 31, 32,33,34).the result showed that cinnamon has antidiabetic activity in all extract and drug compared with control group. The result in fig.(4) showed that hexane extract highly significant hypoglycemic activity compared with metformin and control group and this may be due to the oil substances in these extract . and this result agree with (35) who reported that the oil in cinnamon significantly reduces blood glucose levels in STZ-induced diabetic rats after 3 weeks of treatment . Thus this study showed that administration of different extract of cinnamon bark are reduces blood glucose levels .And this reduction may be due to
the active antihyperglycemic agents present in the extract helps in overcoming the diabetic complications by increasing the insulin secretion (36). However the exact mechanism is not clear and further biochemical and pharmacological investigations are needed to isolate and identify the active ingredient(s) in these extract. These findings are partially similar to those reported by (37) who concluded that intake of 3 gram or 6 gram of cinnamon reduces the fasting serum glucose in people with type 2 diabetes. The hypoglycemic effect cinnamon extract which reported her it may be due to its hyperinsulinemia that evident in this study (38). Other study suggest that Antidiabetic properties have also been reported for Vaccinium angustifolium, the Canadian lowbush blueberry, which also contains oligomeric procyanidins as possible antidiabetic agents (39). The experiments also suggested that the possible mechanism of its hypoglycemic action is may from potentiating the effect of insulin in serum or increasing either the pancreatic secretion of insulin from the existing beta cells or its release from the bound form. (40). Several animal studies have also reported insulin potentiating effects after cinnamon administration. In vivo, administration of aqueous extracts of cinnamon improves glucose metabolism and potentiates the action of insulin. These results suggest that increased glucose uptake in vivo is a result of enhancing the insulin signaling pathway. Other showed that Cinnamon extract fed to high fructose-induced insulin resistant male Wistar rats indicated that insulin stimulated glucose uptake was significantly greater in cinnamon fed rats and that the rate of insulin resistance was reversed by cinnamon feeding.. The mode of action for this hypoglycemic effect of cinnamonn may be attributed to an increase in serum insulin levels, hepatic glycogen storage, or insulin-receptor signaling, an insulinomimetic effect, or a reduction in intestinal α-glycosidase activity (41).
Figure (1). Hypoglycemic activity of methanolic extract of cinnamon in laboratory rats. n = 5

Figure (2). Hypoglycemic activity of hexane extract of cinnamon in laboratory rats. n = 5
Figure (1). Hypoglycemic activity of chloroform extract of cinnamon in laboratory rats. n = 5

Figure (4). Comparison the hypoglycemic activity of methanol, hexane and chloroform of cinnamon extract with metformin drug n = 5.
References:


prevent the formation of advanced glycation endproducts. J. Agric. Food Chem. 56: 1907–1911.


