

Beneficial Role of Taurine on Biochemical Parameters of Diabetic Female Rats

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ABSTRACT

For studying the positive effects of taurine (TAU) on lipid and glucose metabolism. Moreover, the present paper examines the positive roles of glucose and lipid on the correction of oxidative stress diabetes-related complications in alloxan diabetic rats. To achieve the objective of study, 24 of female rats (*Rattus norvegicus*) have been used. The division of animals was done in 4 groups (6 each). Diabetes was enhanced by injected intraperitoneally with alloxan at a single dose in body weight; 125 mg/kg. Diabetic rats go through a specific rise ($p \leq 0.05$) in the glucose levels, triglyceride, total cholesterol, very-low-density lipoprotein, low-density lipoprotein, and malondialdehyde and an important noticeable decrease in high-density lipoprotein, glutathione, and albumin. In addition, taurine supplementation caused a significant reduction in the glucose, total cholesterol, triglycerides, low-density lipoprotein, and very low-density lipoprotein levels. The obtained results revealed that taurine exhibited an inhibitory effect on oxidative stress indices (MDA) and improved antioxidant levels. Taurine could have potential as a pharmaceutical drug for diabetes mellitus (DM), and this invites further studies in this field.

Keywords: Antioxidants, Diabetes, Lipid profile, Malondialdehyde, Taurine.

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INTRODUCTION

Diabetes can be defined as a complex, chronic illness that needs consistent medical care and treatment to control blood sugar levels and prevent hard and unexpected or long-term complications of the disease.¹ There is a rapid growth of this disease and can harm 340 million population; it is expected that the number of diabetes patients could reach 552 million in 2030.² Therefore, diabetes indicates a real challenge to healthcare systems globally. An absolute reduction of the secretion of insulin is seen in type-1 diabetes, associated to the destruction of pancreatic β -cells. It is majorly inherited from the previous generations.³ Furthermore, a combination of resistance of insulin action and impaired secretion of insulin causes type 2 diabetes, and more than 90% of cases fall under this category.

As there is a rapid increase in the global diabetic population, new therapies are required for impactful yet less adverse effects. There are various oral antihyperglycemic agents that have beneficial side effects and some do not possess an effect on chronic diabetes.⁵ Despite introducing hypoglycemic drugs, diabetes and associated complications continue to be a complicated medical problem.⁶ So, there is an increasing need for new natural antihyperglycemic products,

especially nutraceuticals with less side effects, safe, and high antihyperglycemic potential.

Taurine or 2-aminoethylsulfonic acid is a non-protein and non-essential amino acid. Its presence is important in the cases of diet, disease, and aging.⁷⁻⁸ Taurine is a vital acid in insulin immunity and sensitivity.⁹⁻¹⁰ Because of its benefits, it is considered as wonder molecule,¹¹ the control on diabetes and its consequences. The results studies in animals concluded that having suitable taurine levels helps the control of diabetes. This can be by the reduction of blood glucose and restoring insulin sensitivity.¹²

In this paper, the anti-diabetic activity of taurine was evaluated. Moreover, a role in the correction of oxidative stress diabetes-related complications in alloxan diabetic rats was also assessed.

Materials and Methods

Animals

Female rats (190-200 g), were collected from the department of biology (animal house unit), college of Science, University of Thi-Qar, Iraq. An air-conditioned room was used by the animals ($22 \pm 3^\circ\text{C}$) with 12 hours light/dark cycle and $55 \pm 5\%$ humidity. The diet had free access to water and was standard.

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The local committee gave permission for the design of the experiments, and the protocols were conducted based on the guidelines of the National Institutes of Health (NIH).

Chemicals

Alloxan, taurine, trichloroacetic acid (TCA), thiobarbituric acid (TBA), paraphenylenediamine (PPD), azide sodium, and sodium carbonate anhydrate was purchased from BDH, England, Sodium phosphate acidic NaH_2PO_4 , Sodium phosphate basic Na_2HPO_4 were purchased from FlukaGarnatie) Switzerland).

Diabetic Increase

A single intraperitoneal injection of 125 mg/kg body weight alloxan-induced diabetes, which went through advanced preparation, seven days later, determination of serum glucose levels was done. The experiments included rats with fasting blood glucose over 250 mg/dL, and they were deemed.

Experimental Design

Four groups (6 rats/group) were used to divide twenty-four female rats as follows:

Group 1: Treatment was given to animals with distilled water for 15 days.

Group 2: Animals were injected intraperitoneally with alloxan at single dose 125 mg/kg body weight.

Group 3: Intraperitoneal injections of taurine were given to animals at a dose of 100 mg/kg bodyweight for 15 days.

Group 4: Intraperitoneal injections were given to animals with alloxan at a single dose (125 mg/kg body weight) and taurine was injected at a dose (100 mg/kg body weight) after 7 days for 15 days.

Biochemical Parameters

The measurement of glucose was done by using serum. The method of Barham *et al.* (1972)¹³ was used to measure it (Randox, UK) supplied the required reagents. Total cholesterol (TC) was measured according to the method of Allan and Dawson (1979),¹⁴ the utilized reagents were supplied by (Biolabo, France). Triglycerides levels were observed by applying the method of Tietz *et al.* (1999),¹⁵ the used reagents

were supplied by (Biolabs, France), high-density lipoprotein (HDL) was determined according to the method of Lopes-Virella (1977),¹⁶ the utilized reagents were supplied by (Biolabo, France), very low-density lipoprotein (VLDL), and low-density lipoprotein (LDL) were measured by the use of method of Friedwald *et al.* (1972),¹⁷ malondialdehyde (MDA) was measured based on the method of Fong *et al.* (1973).¹⁸ Albumin was also measured, and it is performed by the method of (Doumas *et al.* (1971),¹⁹ the used reagents was supplied by (Biolabo, France), decreased glutathione was measured based on the method of Ellman's (1959).²⁰

Statistical Analysis

The software SPSS version 15.0 was used to obtain the statistical analysis; mean \pm standard deviations (mean \pm SD) and LSD were used to express the results. One-way ANOVA-test was examined for the comparison parameters of the various groups that have been studied. p-values ($p \leq 0.05$) were statistically important for the study.

RESULTS

Effect of Taurine on Serum Glucose and Lipid Profile in Alloxan- Diabetic Rats

The results showed a specific increase ($p \leq 0.05$) in the concentration of serum glucose, triglycerides (TG), TC, very low-density lipoprotein (VLDL) and low-density lipoprotein (LDL) in group 2 compared with group 1. A nonspecific difference in the serum glucose concentration, TG, TC, VLDL and LDL in group 3 in comparing with group 1. Also, there was a specific reduction of ($p \leq 0.05$) in the concentration of serum glucose, TC, TG, LDL and VLDL in group 4 compared with group 2. While there was a specific reduction of ($p \leq 0.05$) in the concentration of serum high-density lipoprotein (HDL) in group 2 in comparison with group 1. Furthermore, there was no specific variation in the serum HDL concentration in group 3 in comparing with group 1. Furthermore, there was a specific enhancement of ($p \leq 0.05$) in the concentration of serum HDL in group 4 in comparison with group 2. (Table 1).

Table 1: Effect of taurine on serum glucose and lipid profile in diabetic rats.

Parameters/groups	Group 1	Group 2	Group 3	Group 4	LSD
Glucose(mg/dL)	96.20 \pm 10.77 ^c	270.70 \pm 31.24 ^a	95.96 \pm 5.70 ^c	116.49 \pm 19.78 ^b	19.44
TC (mg/dL)	131.34 \pm 16.90 ^c	279.25 \pm 16.55 ^a	120.92 \pm 14.47 ^c	174.75 \pm 11.70 ^b	21.70
TG (mg/dL)	87.52 \pm 3.94 ^c	248.30 \pm 16.56 ^a	82.46 \pm 5.50 ^c	131.28 \pm 7.98 ^b	13.26
HDL (mg/DL)	46.79 \pm 3.11 ^b	31.61 \pm 3.68 ^c	48.91 \pm 2.03 ^b	55.12 \pm 4.25 ^a	4.23
LDL (mg/dL)	67.04 \pm 14.49 ^c	197.48 \pm 19.84 ^a	55.53 \pm 14.34 ^c	93.38 \pm 10.71 ^b	21.83
VLDL (mg/dL)	17.51 \pm 1.79 ^c	49.55 \pm 3.31 ^a	16.49 \pm 1.09 ^c	26.26 \pm 1.60 ^b	2.65

- The mean \pm standard division value of 6 rats was presented.

- Different letters refer to a significant difference at ($p \leq 0.05$).

Table 2- Effect of taurine on oxidant-antioxidant markers in diabetic rats.

Parameters/Groups	Group	1			LSD
		Group 2	Group 3	Group 4	
MDA ($\mu\text{mol/L}$)	1.67 \pm 0.34 ^c	3.05 \pm 0.54 ^a	1.52 \pm 0.15 ^c	1.87 \pm 0.16 ^b	0.48
Albumin (g/dL)	4.41 \pm 0.42 ^a	3.36 \pm 0.41 ^b	4.04 \pm 0.38 ^a	4.31 \pm 0.39 ^a	0.51
Glutathione ($\mu\text{mol/L}$)	327.60 \pm 28.90 ^a	200.32 \pm 12.36 ^d	292.85 \pm 10.29 ^b	275.31 \pm 17.87 ^c	24.89

- The mean \pm standard deviation value of 6 rats was represented.

- Different letter refers to a significant difference at ($p \leq 0.05$).

Effect of Taurine on Serum Malondialdehyde (MDA) and Antioxidants in Alloxan- Diabetic Rats.

The results showed a noticeable increase ($p \leq 0.05$) in the serum MDA concentration in group 2 by comparing with group 1). No specific difference in the serum MDA concentration in group 3 was noticed by comparing with group 1. Additionally, there was a specific reduction of ($p \leq 0.05$) in the serum malondialdehyde (MDA) concentration in group 4 by comparing it with group 2. While there was a specific reduction of ($p \leq 0.05$) in the serum albumin and glutathione (GSH) concentration in group 2 by comparing with group 1. Furthermore, no specific difference in the serum albumin concentration in group 3 by comparing with group 1. Also, there was a specific reduction of ($p \leq 0.05$) in the serum albumin and glutathione (GSH) concentration in group 4 by comparing with group 2 (Table 2).

DISCUSSION

The mechanism through which alloxan brings about its diabetic state included selective destruction of pancreatic β -cells that secrete insulin, which reduces the activity of cells, leading a weakness in the utilization of glucose by tissues.²¹ Taurine exerts hypoglycemic effects by regulation of gene expression needed for the insulin secretion of stimulated glucose and development of insulin action,²² as well as by facilitating the insulin interaction with its receptor.⁹ On the other hand, taurine increases glycolysis, glucose oxidation, and glycogenesis.²³ It has hypoglycemic effects, and this forms a reduction in the fructose amine and glucose levels along with an enhancement in the glycogen, C-protein and insulin level, and glycogen in the liver.²⁴

Hyperlipidemia is a relatively common challenge in patients with poorly controlled diabetes mellitus²⁵ and coexists with hyperglycemia, which is attributed by enhanced triglycerides, cholesterol, and phospholipids levels, and also lipoprotein modification. The diabetic hyperlipidemia was characterized to the imbalance of regulation of hormone of metabolism of glucose.²⁶ The hypolipidemic impact of TAU could be partial because of the inhibition of the absorption of cholesterol in intestine or the enhancement of cholesterol conversion to bile acid. Moreover, it might be because that TAU is responsible for the HDL increase or ending the serum lipoprotein balance that contains cholesterol.²⁷ Moreover, it is likely possible that drinking TAU can enhance clearance of serum cholesterol

and reduce TC level in liver in high-fat/cholesterol dietary hamsters, which may be because of LDL receptor regulations, thus increases fecal TC and bile acids output.²⁸ Yang *et al.* (2010)²⁹ stated that TAU could alleviate blood lipids and hepatic damage induced by a high-fat/cholesterol-dietary diet. The data of Saleh (2012)³⁰ indicated that the treatment of diabetic rats with TAU induced a decrease in lipid profile except for HDL-cholesterol.

The present study revealed that the elevation in MDA level might be a reflection of a decrease in the enzymatic and non-enzymatic antioxidants of defense systems. Previous studies have reported that there was an increase in lipid peroxidation in liver, kidney, brain, heart,³¹ pancreas,²⁷ and erythrocytes³² of diabetic rats. Patel *et al.* (2009)³³ suggested this elevation in hepatic MDA level might be due to high concentration of lipid, which was found in the liver of diabetic rats, and resulted in the activation of nicotinamide adenine dinucleotide phosphate (NADPH) dependent microsomal lipid peroxidation in the liver. The decreasing in albumin level is due to the increase of the synthesis of lipid peroxide and increase the formation of free radicals, which result in increasing of membranes permeability and leaking the proteins outside the vascular system.³⁴ Hyperglycemia causes an increase in reactive oxygen metabolites and their derivatives.³⁵ Since GSH is an important antioxidant, GSH deficiency causes increase in oxidative stress.³⁶ Hyperglycemia induced oxidative stress and reduction in the levels of GSH in the vascular straight muscles.³⁷ The prevention of oxidative stress by taurine was also reported in alloxan-induced type 1 diabetic.³⁸ Interestingly, taurine supplementation for 2 days later of STZ injection, prolonged survival in diabetic rats.³⁹ This observation indicates that taurine may confer resistance against some stresses induced by hyperglycemia, which may associate with the beneficial role against the complications. Administration of taurine protected the tissue damage produced by the acute sublethal dose of γ - irradiation in rats by decreasing oxidative stress.^{8,40-41}

CONCLUSION

The benefits of the taurine amino acid appear to be due to its various actions on cellular functions, while toxicity seems relatively low. The possible taurine impact in improving diabetic complications is showed generally post-treatment supplementation for diabetes induced by alloxan.

REFERENCES

- Atlanta GA, Centers for Disease Control and Prevention. National Diabetes Fact Sheet: Centers for Disease Control and prevention. *US Department of Health and Human Services* (2012). <http://www.cdc.gov/diabetes/pubs/pdf/ndfs>.
- Bergman M, Buysschaert M, Schwarz PEH, Albright A&Yach D, Diabetes prevention: global health policy and perspectives from the ground. *Diab. Manage*, 2 (2012), 309.
- Bottini N, Vang T, Cucca F&Mustelin T, Role of PTPN22 in type 1 diabetes and other autoimmune diseases. *Seminars in Immunol*, 18(2006) 207.
- Warren RE, The stepwise approach to the management of type 2 diabetes, *Diabetes Research and Clinical Practice*, 65(2004)53.
- Dubey GP, Dixit SP&Alok S, Alloxan-induced diabetes in rabbits and effect of herbal formulation D-400. *Indian J. Pharmacol*, 26(1994)225.
- Prince PS, Menon VP&Pari L, Hypoglycemic activity of *Syzgiumcumini* seeds: effect on lipid peroxidation in alloxan diabetes rats. *J. Ethnopharmacol*, 61(1998) 1.
- Pierno S, De Luca A, Camerino C, Huxtable RJ&Camerino DC. Chronic administration of taurine to aged rats improves the electrical and contractile properties of skeletal muscle fibers. *J. Pharm. Exp. Therap*, 286(1998)1183.
- Auda MA, Majid A& Al-Fartosi KG, Protective role of polyphenolic compounds extracted from *Cyperusrotundus* rhizomes and taurine on troponin-I and some oxidant/antioxidants parameters of female rats treated with an isoproterenol-induced myocardial infarction. *Journal of Thi-Qar University* 2018; 13(2): 61-76.
- Yamori Y, Food factors for atherosclerosis prevention: Asian perspective derived from analyses of worldwide dietary biomarkers. *Exp. Clin. Cardiol*, 11(2006)11. 2006.
- Yamori Y, Liu L& Mori M, Taurine as the nutritional factor for the longevity of the Japanese revealed by a world-wide epidemiological survey. *Adv. Exp. Med. Biol*, 643(2009)13.
- Yamori Y, Taguchi T, Hamada A, Kunimasa K, Mori H, *et al*, Taurine in health and diseases: consistent evidence from experimental and epidemiological studies. *J. Biomed. Sci*, 17(2010).
- Franconi F, Di Leo MAS, Bennardini F & Ghirlanda G, Is taurine beneficial in reducing risk factors for diabetes mellitus? *Neurochem. Res*. 29(2004)143.
- Barham D&Trinder P. *Analyst*, 97(1972)142.
- Allan C& Dawson JG, Enzymatic Assay of Total Cholesterol Involving Chemical or Enzymatic Hydrolysis-a Comparison of Methods. *Clin. Cem*, 25(1979) 976.
- Tietz NW, Burtis CA, Ashwood ER& Saunder WB, *Text Book of Clinical Chemistry*. 3rd Ed.(1999)809.
- Lopes-Virella MF, Cholesterol Determination in High-density Lipoproteins Separated by Three Different Methods. *Clin. Chem*, 23(1977)882.
- Friedwald WT, Levy RI& Fredrickson DS, Estimation of Concentration of LDL-C in Plasma without the Use of Preparative Ultracentrifuge. *Clinical Chemistry*, 18(1972)499.
- Fong KL, McCay PB& Poyer JL, Oxidative Stress. *Free Rad J Biol Chem*, 248(1973)7792.
- Doumas BT, Watson W A& BIGGS HG, Albumin Standards and the Measurement of Serum Albumin with Bromocresol Green. *Clin Chem. Acta*, 31(1971)87.
- Ellman GL, Tissue Sulfhydryl Groups. *Arch Biochem Biophys*, 82(1959)70.
- Anusuya N, Rajarathinam NI&Nungampakkam SM, Antidiabetic, antihyperlipidemic and antioxidant effect of ethanolic extract of *Curcuma raktakantha* on streptozotocin induced diabetic rats. *International Journal of Pharmacy and Pharmaceutical Sciences*, 5(2013) 201.
- Carneiro EM, Latorraca MQ, Araujo E, Beltra M, Oliveras MJ, *et al.*, Taurine supplementation modulates glucose homeostasis and islet function. *Journal of Nutritional Biochemistry*, 20(2009) 503.
- Higo S, Miyata S, Jiang QY, Kitazawa R, Kitazawa S, *et al.*, Taurine administration after appearance of proteinuria retards progression of diabetic nephropathy in rats. *Kobe J Med Sci*, 54(2008)35.
- Gavrovskaya LK, Ryzhova OV, Safonova AF, Matveev AK&Sapronov NS, Protective effect of taurine on rats with experimental insulin-dependent diabetes mellitus. *Bull Exp Biol Med*, 146(2008)226.
- Shivanand KG, Manjunath ML&Jeganathan PS, Lipid profile and its complications in diabetes mellitus. *International Journal of Biomedical and Advance Research*, 03(2012)775.
- Bagri P, Ali M, Aeri V, Bhowmik M& Sultana S, Antidiabetic effect of *Punicagranatum* flowers: Effect on hyperlipidemia, pancreatic cells lipid peroxidation and antioxidant enzymes in experimental diabetes. *Food Chem. Toxicol*, 47(2009) 50.
- Heibashy MIA, Hypolipidemia effect of taurine and L-carnitine on rats fed a high cholesterol diet. *J. Union Arab Biol. Cairo*, 14(2000)11.
- Chang YY, Chou CH, Chiu CH, Yang KT, Lin YL, *et al*, Preventive Effects of Taurine on Development of Hepatic Steatosis Induced by a High-Fat/Cholesterol Dietary Habit. *J. Agric. Food Chem*, 59(2011)450.
- Yang SF, Bor-Show T, Kuo-Tai Y, Yuan-Chao H, Yuan-Yen C, *et al.*, Taurine alleviates dyslipidemia and liver damage induced by a high-fat/cholesterol-dietary habit. *Food Chemistry*, 120(2010) 156.
- Saleh AA, Effect of taurine and/or ginseng and their mixture on lipid profile and some parameters indicative of myocardial status in streptozotocin-diabetic rats. *Journal of basic and applied zoology*, 65(2012)267.
- Kumar G, Banu S&Murugesan AG, Influence of *Helicteresisora* administration for diabetes mellitus: Its effect on erythrocyte membrane and antioxidant status. *Food Chem. Toxicol*. 47(2009)1803.
- Chandramohan G, Al-Numair KS&Pugalendi KV, Restoration of altered plasma, erythrocyte and liver antioxidant levels by 3-hydroxymethyl xylitol in streptozotocin-diabetic rats. *Int. J. Integr. Biol*, 5 (2009) 176.
- Patel SS, Shah RS&Goyal RK, Antihyperglycemic, antihyperlipidemic and antioxidant effects of Dihar, a polyherbal ayurvedic formulation in streptozotocin-induced diabetic rats. *Ind. J. Exp. Biol*, 47(2009)564.
- Vlassara H, Brownle M&Cerami A, Nonenzymatic glycosylation of peripheral nerve protein in diabetes mellitus. *Proceedings of the National Academy of Sciences of the United States of America*, 78(1981)5190.
- Ceriello A&Giugliano D, Oxidative stress and Diabetic complications. In: Alberti KGMM, Zimmet P, Fronzo RA, editors. *International Textbook of Diabetes Mellitus*. Chichester, UK: John Wiley, (2001) 1453.
- Seghrouchni I, Draï J, Bannier E, Riviere J, Calmard P, *et al*, Oxidative stress parameters in type 1, type 2 and insulin-treated

- type 2 diabetes mellitus; insulin treatment efficiency. *Clin ChimActa*, 321(2002)89.
37. Powell LA, Nally SM, McMaster D, Catherwood MA& Trimble ER, Restoration of glutathione levels in vascular smooth muscle cells exposed to high glucose conditions. *Free RadicBiolMed*, 31(2001)1149.
38. Di Leo MA, Santini SA, GentiloniSilveri N, Giardina B, Franconi F&Ghirlanda G, Long-term taurine supplementation reduces mortality rate in streptozotocin-induced diabetic rats. *Amino Acids*, 27(2004)187.
39. Monira AA, Manal HE, Amina MAT, Mamdouh MA&Anisa SM, Evaluation of Taurine Role on Some Biochemical and Histological Alterations in γ - Irradiated Rats. *Int. J. Pharm. Sci. Rev. Res*, 30(2015)263.
40. Selvan VT, Manikandan L, Senthil KGP, Suresh R, Kakoti BB&Gomathi P, Antidiabetic and antioxidant effect of methanol extract of *Artanemesamoides* in streptozotocin-induced diabetic rats. *International Journal Applied Research in Natural Products*, 1(2008)25.
41. Majid A, Auda MA& Al-Fartosi KG, Cardioprotective Activity of Polyphenolic Extract of Tubers of *CyperusRotundus* and Taurine against Isoproterenol – induced Myocardial Infraction in Rats: Troponin-I and Histological Findings. *Journal of Global Pharma technology*, 10(2018)479.