

## Journal of Global Pharma Technology

Available Online at: www.jgpt.co.in

**RESEARCH ARTICLE** 

# The Effect of Type II Diabetes Mellitus on Some Prostate Markers in Iraqi Men

## Khalid Maseer Rmaidh\*, Omar S. I. Al-janabi\*\*, Mustafa Salim Ibrahim\*\*\*

<sup>1.</sup> Medical Department, Lecturer of Medicine. College of Medicine / University Of Anbar/Iraq.

<sup>2.</sup> Medical Laboratory Techniques Department, Lecturer of biochemistry. AL-Maaref University College/Iraq.

#### \*Corresponding Author: Khalid Maseer Rmaidh

#### Abstract

Aim: Diabetes mellitus is a spread disease in the whole world and this study was conducted on Iraqi diabetic men to know the relation between type II diabetic and some prostate related markers including (prostate size, total prostate antigen, random blood sugar and zinc). Material and methods: This study included one hundred men their age ranges between (40-60 year), fifty males were suffering from type II diabetes mellitus, and other fifty men were non-diabetics. Some parameters such as (Prostate size, total prostate specific antigen (tPSA), random blood sugar (RBS) and Zinc (Zn<sup>+2</sup>) were studied. Results: The results have showed significant increase in prostate size 50.5000 ± 3.748 and random blood sugar 203.55 ± 13.629 mg/dl compared with non-diabetic group  $39.650 \pm 2.484$  and  $125.750 \pm 5.396$  mg/dl, respectively but there is non-significant difference in total prostate specific antigen 1.543 ± 0.087 ng/ml and zinc  $9.725 \pm 0.344 \mu$ mol/l in diabetic group compared with non-diabetic group  $1.612 \pm 0.152$  ng/ml and  $11.025 \pm 0.582 \mu$ mol/l respectively at P<0.05.Conclusions: Patients with type II diabetes mellitus are more liable to suffer from increase in prostate size and increase in RBS levels in Iraqi men under study.

Keywords: Diabetic patient, Prostate size, RBS, Zinc and tPSA.

#### Introduction

Generally, mellitus diabetes (DM)considered of one the main incommunicable disorders and the prevalence of this disease in developing countries was highly over the past decades [1]. The incidence of this disease in 2015 was more than 415 million people worldwide. while in Chinese was one fourth in the population are affected with diabetes [2].

Occasionally, people with inactive lifestyles and insanity dietary behavior showed highly documented to have risk factors of diabetes. In addition, several studies indicated that some environmental chemicals like (heavy metals) play a crucial role in the enhancement of T2DM [3].

Therefore, the general people during the daily life are in direct contact with these kinds of metals such as dietary intake, inhalation of ambient air, water drinking and dermal contact of consumable goods [4].

The correlation between diabetes and prostate illnesses (for instance prostate cancer and benign prostatic hyperplasia (BPH) is complex.

Diabetes has the possibility to decrease the danger of prostate cancer. Men who diagnosed with type II DM have lower androgen levels; therefore, they have been immediately connected with prostate cancer hazards. Accordingly, diabetes relates to changes in insulin levels and demand, which seems to be higher in pre-and early diabetic cases. These procure to reduced insulin-like growth factor-I (IGF-I), binding protein levels and eventually, to higher IGF-I, therefore, all factors are probably relating to BPH development and the risk of prostate cancer. Thus, in this presented study, we tested the association between type II diabetes mellitus and some prostate markers (P. Size and tPSA) and Zinc (Zn) on some of the Iraqi men who affected by diabetic and who has not.

## Material and Method

The study took place in Anbar province in Iraq at the department of medicine of Ramadi Teaching Hospital. It started from Nov2017 to Apr2018. In the beginning, we got the approval from all men who would be used as samples in this study. The ages of those men range between (40-60 years old). Our study performed on fifty men who diagnosed by diabetes type II, and another fifty men who doesn't have diabetics. All the data results have been recorded according to administering a questionnaire, and all men's size have been checked and prostates examined.

Also, we drawn five milliliters of venous blood and the serum separated and send to our laboratories at Ramadi Teaching Hospital. Serum samples have been divided into Eppendorf and stored at -20 C until used.

In this study, the following aspects have been considered for all the patients:

• Prostate size was calculated by using abdominal ultrasonography

- Total prostate specific antigen (tPSA) was determined by (ELISA) method followed using kit purchased from bioactiva diagnostica company (Germany) [5, 9].
- Random blood sugar (RBS) was measured by enzymatic colorimetric method [10].
- Zinc was determined by using atomic absorption spectrophotometric (AAS) [11].

## **Statistical Analysis**

Data were analyzed by using version 18.0 of SPSS program (SPSS, IBM Corporation, Chicago, IL, USA). The P value was supposed to be significant at  $\leq 0.05$ .

#### Results

The results of this study (Table 1 and Fig. 1) found out that there is a significant increase in prostate size and random blood sugar but, they showed no significant difference in total prostate specific antigen and zinc in diabetic group compared with non-diabetic group at  $P \leq 0.05$ .

Table.1: Shows the relationship between P. Size, TPSA, RBS and Zinc in type II diabetic and non-diabetic men at P ≤0.05

Groups	<b>Diabetic</b> (Mean± SE) <b>no= 50</b>	Non-diabetic (Mean± SE) no=50	Р
P. Size	$50.5000 \pm 3.748$	$39.650 \pm 2.484$	0.220
tPSA (ng/ml	$1.543 \pm 0.087$	$1.612 \pm 0.152$	0.696
RBS (ng/dl)	$203.550 \pm 13.629$	$125.750 \pm 5.396$	0.000
Zinc (µmol/l)	$9.725 \pm 0.344$	$11.025 \pm 0.582$	0.640



Figure 1: shows the relationship between P. Size, TPSA, RBS and Zinc in type II diabetic and non-diabetic men

## Discussion

The results of this study in (Table 1 and Fig. 1) showed significant increase in prostate size and random blood sugar in diabetic group compared with non-diabetic group. These outcomes were in agreement with other global study among aged men are above 50% related to the occurrence of benign prostatic hyperplasia (BPH) [12], BPH became as important medical public health problem. Nevertheless, there are some etiologic agents need to understand for developing the mechanism of BPH. Previous studies suggested that some factors play a main role in this matter such as diabetes, obesity, alcohol consumption, physical activity, dyslipidemia, hypertension, diet, smoking, and the environment [13, 14]. Since the most common benign tumor in men is BPH and these events are age development related.

Pathologically the BPH exhibit a type of nonmalignant cells growth with unregulated shape leading to an increase in stromal cells size and prostate epithelial especially in the end stage. Therefore, there are some corresponding systems are likely to be involved in the etiology of BPH to be solved. These systems in addition to the resident aspects are nerve, endocrine, immune, and vascular system [15]. A study by [15] indicated that the pathogenic of BPH is controlled by endocrine mechanism; this will keep the loss of specific pathway which remains to be more investigated. Regarding medical opinions about some the dissimilarity of the pathogenesis between T2DM and BPH, these still misgivings the relation between these public diseases.

On the contrary, however, T2DM and BPH are different in some clinical features; both diseases have the same epidemiological characters related with age and diet which probably associated in the pathogenic pathways [16, 17]. The results of our study are consistent with many studies that found the large prostate volume is associated with the components of metabolic syndrome [18, 19]. In fact, both type II DM and BPH seem to share similar epidemiological features, possibly related to aging and diet [20] Barnard et al [21] found that the reduction of insulin level leads to decrease of stem epithelial prostate cells growth.

Other possible mechanisms have been suggested to correlate the improvement of type II diabetes mellitus with BPH, [22] such as the enhancement activity in the autonomic nervous system caused by hyper-insulinemia, and peripheral sympathetic nerve tone and hypoxia caused by DM-induced vascular damage [23].

Also, many previous studies found a positive correlation between the levels of fasting blood glucose and prostate size [24]. Serum PSA concentration is age- dependent, i.e. it tends to increase with age because the prostate enlarges with years and contains more PSAproducing tissue [25]. Results from epidemiologic findings on the relationship between prostate cancer and diabetes risk are often confusing. Some studies suggested that diabetic's patient has a low risk of prostate cancer. This relationship investigated in the follow-up study conducted from 1986 till 1994 in the United States.

The basis of this relationship is unclear, however, it may reflect hormonal changes associated with diabetes or it may due to being low testosterone level [26]. The theory related to the increasing total or bioavailable testosterone levels and prostate cancer riskfactor have been supported by some studies [27]. Results of the Cancer Prevention Study (1959-1972) have indicated that men without diabetes had a low risk of prostate cancer than men who had DM for five years or more [28]. Another study also found a relationship among diabetes and prostate cancer [29]. Results of a population-based study of Chinese men documented that in Chinese men with high levels of serum insulin may encourage the risk of prostate cancer [30].

In addition, another study in north Italy found no association between the risk of cancer and diabetic prostate patient (increased risk of liver cancer, pancreas and endometrium was observed) [31].Our study has shown no significant high defect in Zinc element between the diabetic and nondiabetic group. Since, the metabolic characteristics and biological functions are affected by some elements such as selenium, zinc [32]. Numerous manganese, and studies indicated that glucose homeostasis process regulated by zinc elements.

The association of type II diabetes and plasma zinc concentrations has been illustrated in a study among Chinese population in 1796 [33, 34]. However, our study has found no significant results in the reduction of zinc levels in serum among men with diabetes. In contrast, different findings observed by a study among postmenopausal women indicated that a woman with diabetic has significant increase level in serum zinc than in the healthy group [35].

Recent evidence indicated that the role of zinc transporter is diverse between ISG zinc ZnT8, this different pathway perhaps considered with protection or risk of diabetes [33]. A study by [36] confirmed that the sensitivity of peripheral insulin was improved by zinc through the role of activated insulin stimulated glucose transports [36]. In addition, zinc plays an important action in the physiology of  $\beta$  cells [37]. The recent genetic study demonstrated that the specific  $\beta$  cells islet of zinc ZnT8 (SLC30A8) transporter expression which is

### References

- 1. Guariguata L, Whiting DR, Hambleton I, Beagley J, Linnenkamp U, Shaw JE (2014) Global estimates of diabetes prevalence for 2013 and projections for 2035. Diabetes research and clinical practice, 103(2): 137-149.
- 2. Atlas ID (2017) In., 6th edn. Brussels, Belgium: International Diabetes Federation; 2013. URL: <u>https://www.idf.org/component/attachments/att</u> <u>achments.html</u>.
- 3. Thayer KA, Heindel JJ, Bucher JR, Gallo MA (2012) Role of environmental chemicals in diabetes and obesity: a National Toxicology Program workshop review. Environmental health perspectives, 120 (6): 779.
- 4. Rehman K, Fatima F, Waheed I, Akash MSH (2018) Prevalence of exposure of heavy metals and their impact on health consequences. Journal of cellular biochemistry, 119(1): 157-184.
- 5. Giovannucci E (2001) Medical history and etiology of prostate cancer. Epidemiologic reviews, 23(1): 159-62.
- 6. Tavani A, Gallus S, Bertuzzi M, Dal Maso L, Zucchetto A, Negri E, La Vecchia C (2005) Diabetes mellitus and the risk of prostate cancer in Italy. European urology, 47(3): 313-317.
- 7. Donnell RF (2011) Benign prostate hyperplasia: a review of the year's progress from bench to clinic. Current opinion in urology, 21(1): 22-26.
- 8. Vikram A, Jena G, Ramarao P (2010) Insulinresistance and benign prostatic hyperplasia: the connection. European journal of pharmacology, 641(2-3): 75-81.
- Espana F, Sanchez-Cuenca J, Estelles A, Gilabert J, Griffin JH, Heeb MJ (1996) Quantitative immunoassay for complexes of prostate-specific antigen with alpha2macroglobulin. Clinical chemistry, 42(4): 545-550.
- 10. Barham D, Trinder P (1972) An improved colour reagent for the determination of blood glucose by the oxidase

the best regulator of the insulin secretion might change the risk of T2DM development [38].

## Conclusions

Patients with type II diabetes mellitus are more liable to suffer from increase in prostate size and increase in RBS levels in Iraqi men under study.

system. Analyst, 97(1151): 142-145.

- 11. Taylor A, Bryant TN (1981) Comparison of procedures for determination of copper and zinc in serum by atomic absorption spectroscopy. Clinica chimica acta, 110(1): 83-90.
- 12. Marberger M, Harkaway R, De la Rosette J (2004) Optimizing the medical management of benign prostatic hyperplasia. European urology, 45(4): 411-419.
- Parsons JK (2007) Modifiable risk factors for benign prostatic hyperplasia and lower urinary tract symptoms: new approaches to old problems. The Journal of urology, 178(2): 395-401.
- 14. Ibrahim MS, Abd ZH, AL-Rawi KF (2018) Effect of obesity on renal function test, uric acid, and gamma glutamyl transferase (GGT) in sample of Iraqi men. Kufa Journal for Chemistry, 2: 3.
- Lee C, Kozlowski JM, Grayhack JT (1995) Etiology of benign prostatic hyperplasia. The Urologic clinics of North America, 22(2): 237-246.
- Gades NM, Jacobson DJ, Girman CJ, Roberts RO, Lieber MM, Jacobsen SJ (2005) Prevalence of conditions potentially associated with lower urinary tract symptoms in men. BJU international, 95(4): 549-553.
- 17. Resistance I (1991) A Multifaceted Syndrome Responsible for NIDDM, Obesity, Hypertension, Dyslipidemia, and Atherosclerotic Cardiovascular Disease. Diabetes Care, 14(3): 173-194.
- 18. Hammarsten J, Högstedt B, Holthuis N, Mellström D (1998) Components of the metabolic syndrome-risk factors for the development of benign prostatic hyperplasia. Prostate cancer and prostatic diseases, 1(3): 157.
- 19. Ozden C, Ozdal OL, Urgancioglu G, Koyuncu H, Gokkaya S, Memis A (2007) The correlation between metabolic syndrome and prostatic growth in patients with benign prostatic hyperplasia. European urology, 51(1): 199-206.

- 20. Gades N M, Jacobson DJ, Girman CJ, Roberts RO, Lieber MM, Jacobsen SJ (2005) Prevalence of conditions potentially associated with lower urinary tract symptoms in men. BJU international, 95(4): 549-553.
- 21. Barnard RJ, Kobayashi N, Aronson WJ (2008) Effect of diet and exercise intervention on the growth of prostate epithelial cells. Prostate Cancer and Prostatic Diseases, 11(4): 362.
- 22. Berger AP, Deibl M, Halpern EJ, Lechleitner M, Bektic J, Horninger W, Frauscher F (2005) Vascular damage induced by type 2 diabetes mellitus as a risk factor for benign prostatic hyperplasia. Diabetologia, 48(4): 784-789.
- 23. Kasturi S, Russell S, McVary KT (2006) Metabolic syndrome and lower urinary tract symptoms secondary to benign prostatic hyperplasia. Current Prostate Reports, 4(3), 127-131.
- 24. Kim WT, Yun SJ, Choi YD, Kim GY, Moon SK, Choi YH, Kim WJ (2011) Prostate size correlates with fasting blood glucose in nondiabetic benign prostatic hyperplasia patients with normal testosterone levels. Journal of Korean medical science, 26(9): 1214-1218.
- 25. Oesterling JE, Jacobsen SJ, Chute CG, Guess HA, Girman CJ, Panser LA, Lieber MM (1993) Serum prostate-specific antigen in a community-based population of healthy men: establishment of age-specific reference ranges. Jama, 270(7): 860-864.
- Giovannucci E, Rimm EB, Stampfer MJ, Colditz GA, Willett WC (1998) Diabetes mellitus and risk of prostate cancer (United States). Cancer Causes & Control, 9(1): 3-9.
- 27. Kaaks R, Lukanova A, Sommersberg B (2000) Plasma androgens, IGF-1, body size, and prostate cancer risk: a synthetic review. Prostate cancer and prostatic diseases, 3(3): 157.
- 28. Will JC, Vinicor F, Calle EE (1999) Is diabetes mellitus associated with prostate cancer incidence and survival? Epidemiology, 313-318.
- 29. Ilić M, Vlajinac H, Marinković J (1996) Casecontrol study of risk factors for prostate cancer. British journal of cancer, 74(10): 1682.

- 30. Hsing AW, Chua Jr S, Gao YT, Gentzschein E, Chang L, Deng J, Stanczyk F Z (2001) Prostate cancer risk and serum levels of insulin and leptin: a population-based study. Journal of the National Cancer Institute, 93(10): 783-789.
- 31. La Vecchia C, Negri E, Franceschi S, D'avanzo B, Boyle P (1994) A case-control study of diabetes mellitus and cancer risk. British Journal of Cancer, 70(5): 950.
- Nordberg GF, Fowler BA, Nordberg M (Eds.) (2014) Handbook on the Toxicology of Metals. Academic press.
- 33. Rutter GA, Chabosseau P, Bellomo EA, Maret W, Mitchell RK, Hodson DJ, Hu M (2016) Intracellular zinc in insulin secretion and action: a determinant of diabetes risk? Proceedings of the Nutrition Society, 75(1): 61-72.
- 34. Shan Z, Chen S, Sun T, Luo C, Guo Y, Yu X, Liu L (2016) U-shaped association between plasma manganese levels and type 2 diabetes. Environmental health perspectives, 124(12): 18-76.
- 35. Skalnaya MG, Skalny AV, Tinkov AA (2017) Serum copper, zinc, and iron levels, and markers of carbohydrate metabolism in postmenopausal women with pre diabetes and type 2 diabetes mellitus. Journal of Trace Elements in Medicine and Biology, 43: 46-51.
- 36. Tang XH, Shay NF (2001) Zinc has an insulinlike effect on glucose transport mediated by phosphoinositol-3-kinase and Akt in 3T3-L1 fibroblasts and adipocytes. The Journal of nutrition, 131(5): 1414-1420.
- 37. Wijesekara N, Chimienti F, Wheeler MB (2009) Zinc, a regulator of islet function and glucose homeostasis. Diabetes, Obesity and Metabolism, 11: 202-214.
- Rutter GA (2010) Think zinc: new roles for zinc in the control of insulin secretion. Islets, 2(1): 49-50.