

19(10): 1-7, 2021; Article no.AJMAH.74081 ISSN: 2456-8414

The Frequency of Thyroid Dysfunctions among Patients of Diabetes Mellitus Presenting in Tertiary Care Hospital of a Developing Country

Rabia Arshad¹, Shahzaib Maqbool^{1*}, Sara Arshad², Fatima Rehman³, Muhammad Nadeem¹, Ruqyia Shabir¹, Abdul Mateen⁴, Rabia Rehman⁵, Arham Ihtesham¹ and Waleed Inayat Mohamed¹

¹Internal Medicine Holy Family Hospital, Rawalpindi, Pakistan.
²Forensic Medicine Sheikh Zaid Hospital, Rahim Yar Khan, Pakistan.
³Biochemistry Department, Shahida Islam Medical College, Bahawalpur, Pakistan.
⁴Internal Medicine Shifa International Hospital, Islamabad, Pakistan.
⁵Internal Medicine Sheikh Zaid Hospital, Rahim Yar Khan, Pakistan.

Authors' contributions

This work was carried out in collaboration among all authors. All authors read and approved the final manuscript.

Article Information

DOI: 10.9734/AJMAH/2021/v19i1030376 <u>Editor(s)</u>: (1) Dr. Ashish Anand, GV Montgomery Veteran Affairs Medical Center, USA. <u>Reviewers:</u> (1) Riju Ramachandran, Amrita Viswavidyapeetham, India. (2) Zainab Sajid Mohammed, University of Ahl al-Bayt, Iraq. (3) Syeda Nayab Batool Rizvi, University of the Punjab, Pakistan. Complete Peer review History: <u>https://www.sdiarticle4.com/review-history/74081</u>

Original Research Article

Received 10 July 2021 Accepted 20 September 2021 Published 25 September 2021

ABSTRACT

Aims: Thyroid disease is a pathological state associated significantly with diabetes mellitus (DM) Type 1 and Type 2. As the prevalence of diabetes mellitus is on the rise in our population, so the purposed significance of our study was to evaluate the frequency of thyroid dysfunction in diabetic patients of our local population.

Study Design: Cross-sectional study.

Place and Duration of Study: This study conducted among patients of diabetes (Type 1 and Type 2) from September 2020 to March 2021 while their visit in Holy family hospital, Rawalpindi, for a routine clinical check-up on an OPD basis.

Methodology: A total of 96 patients with diabetes mellitus on regular medication, 20 to 60 years of age, were included. Patients with diabetic ketoacidosis, hyperlipidemia, pregnant females, and those taking medication that can affect thyroid functions (dopamine antagonists, antiepileptics, oral



contraceptives, lithium, glucocorticoids) were excluded. A venous blood sample was drawn and sent to the laboratory to analyze thyroid function tests for the presence or absence of any thyroid dysfunction. Data analysis was done through SPSS.v.23. Descriptive statistics and Chi-square analysis was used, and a p-value of ≤ 0.05 was considered significant.

Results: The mean age of the patients was 42.97 ± 10.29 years. The majority of the patients, 37.50%, were between 41 to 50 years of age. Out of 96 patients, 61 (63.54%) were male, and 35 (36.46%) were females with a male to female ratio of 1.7:1. The majority of patients, 69 (71.88%), had type II diabetes mellitus. The frequency of thyroid dysfunction found in diabetic patients was 30 (31.25%), with hypothyroidism in 19 (19.79%) and hyperthyroidism in 11 (11.46%) patients.

Conclusion: This study deduced that there is a high frequency of thyroid dysfunction in diabetic patients. In the same vein, hypothyroidism was the common thyroid dysfunction associated with diabetes mellitus.

Keywords: Diabetes mellitus; hypothyroidism; hyperthyroidism; thyroid dysfunction; tertiary care hospital.

1. INTRODUCTION

Diabetes mellitus (DM) is regarded as a single disease entity. It is a heterogeneous group of diseases characterized by chronic hyperglycemia multifactorial associations such with as environmental, genetic and geographical factors acting simultaneously or jointly [1]. There are three main types of diabetes mellitus (DM): (i) Type I or insulin-dependent DM or juvenile diabetes, (ii) Type II or non-insulin-dependent DM or adult-onset diabetes, (iii) gestational diabetes occurs when pregnant women without a previous diagnosis of diabetes develop a high blood glucose level. Other forms of diabetes mellitus include congenital diabetes due to genetic defects of insulin secretion, cystic fibrosis-related diabetes, steroid diabetes induced by high doses of glucocorticoids, and several forms of monogenic diabetes [2]. There is currently no cure for diabetes. The condition, however, can be managed so that patients can live relatively normal life. Treatment of diabetes focuses on two goals: keeping blood glucose within normal range and preventing long-term complications. The maior lona-term complications relate to damage to blood vessels. Diabetes doubles the risk of cardiovascular The primary "macrovascular" disease [3]. diseases (related to atherosclerosis of larger arteries) are ischemic heart disease (angina and myocardial infarction), stroke, and peripheral vascular disease. Diabetes also damages the capillaries (causes microangiopathy) [4].

Occasionally other endocrine disorders like abnormal thyroid hormone levels are found in diabetes. Various studies have reported a significant association between Diabetes and thyroid disorders [5]. On the one hand, studies have also reported the significance of thyroid hormones' contribution to regulating

carbohvdrate metabolism and pancreatic function; in the same vein, studies are also showing the paramount significance of DM in regulating the thyroid function tests [6,7]. The Thyroid hormones and pancreatic hormones like insulin antagonize each other, and excess and deficit of any one of them can lead to functional derangement of the other [8]. DM appears to influence thyroid function in two sites; firstly, at the level of the hypothalamus where TSH release is regulated, and secondly, at the level of the peripheral tissues where T4 is converted into T3 [9]. A study by M V Jali et al. demonstrated an overall prevalence of 16.2% of thyroid diseases in diabetics [10]. In another study, Udiong CEJ et al. [11] showed hypothyroidism in 26.6% and hyperthyroidism in 19.8% of patients with diabetes mellitus.

As the prevalence of diabetes mellitus is on the rise in our population, so the purposed significance of our study is to evaluate the frequency of thyroid dysfunction in diabetic patients of our local population. As routinely thyroid functions in diabetic patients are not assessed in our general practice, and failure to recognize the presence of abnormal thyroid hormone levels in diabetics might be a primary cause of poor management often encountered in some treated diabetics, so the results of our study would not only provide the magnitude of the local population but also help the clinicians to design a proper management protocol for these particular patients in order to reduce the morbidity of our population.

2. MATERIALS AND METHODS

2.1 Study Design and Setting

It is a cross-sectional study conducted among patients of diabetes (Type 1 and Type 2) from

19th September 2020 to 26th March 2021 while their visit in Holy family hospital, Rawalpindi, for a routine clinical check-up on an OPD basis.

2.2 Sample Size and Study Sampling

A total of 96 patients were included in our study, and consecutive sampling technique was followed during data collection. Sample size calculation was done by utilizing these values (Z=1.96, p=19.8%, q=100-p, d=8%) [11].

2.3 Inclusion and Exclusion Criteria

All patients, either males or females with age group between 20-60 years, diagnosed with diabetes mellitus and on regular medication as per operational definition of >2 years duration were included in our study. Similarly, the patients with diabetic ketoacidosis, chronic renal failure, taking medication that can affect thyroid functions (dopamine antagonists, antiepileptics, oral contraceptives, lithium, glucocorticoids), cases of thyroid disorders. known and hyperlipidemic patients were excluded from our sample population.

2.4 Data Collection Technique

After the approval of the study by the hospital committee of faculty members, 96 patients with diabetes mellitus (as per operational definition) presented to the outpatient department of medicine, Holy Family Hospital, Rawalpindi, fulfilling the inclusion/exclusion criteria were selected. Informed written consent was taken from each patient. A venous blood sample was taken and sent to the laboratory for analysis of thyroid function tests presence or absence of any thyroid dysfunction such as hypothyroidism defined as TSH more than 4.0mIU/L and

hyperthyroidism defined as TSH less than 0.4mIU/L. All the data obtained from patients were recorded on the predesigned Performa.

2.5 Statistical Analysis

Data was entered and analyzed using the program SPSS computer version 23.0. Descriptive statistics were applied to calculate the mean and standard deviation for the age of the patients and the duration of diabetes mellitus. Frequencies and percentages were calculated for categorical variables such as gender, type of diabetes (Type I or II), hypothyroidism (yes/no), and hyperthyroidism (yes/no). Effect modifiers like age, gender, duration of diabetes mellitus. type of diabetes, and BMI (<27kg/m2 ≥27kg/m2) were controlled by stratification. chisquare test was applied post-stratification, and pvalue ≤ 0.05 was considered as significant.

3. RESULTS AND DISCUSSION

3.1 Results

According to demographic characterizations, the age range in this study was from 20 to 60 years, with a mean age of 42.97 ± 10.29 years. The majority of the patients, 37.50%, were between 41 to 50 years of age, as shown in Table 1. Out of 96 patients, 61 (63.54%) were male, and 35 (36.46%) were females with a male to female ratio of 1.7:1, as shown in Fig. 1. The majority of patients, 69 (71.88%), had Type II DM, as shown in Fig. 2. The mean duration of disease in our study was 5.86 ± 2.65 years, with the majority of patients, i.e., 57 (59.38%), were between >2-5 years duration as shown in (Table 1). similarly, the frequency and percentage of patients according to BMI are shown in (Table 1).

Table 1. Demographic characterizations of the patients (Mean Age \pm SD = 42.97 \pm 10.29 years, Mean Duration of disease \pm SD = 5.86 \pm 2.65 years)

Variables	Frequencies (n)	Percentages (%)
Age (in years)		
20-30	12	12.50
31-40	27	28.13
41-50	36	37.50
51-60	21	21.87
Duration of disease (in years)		
>2-5 years	57	59.38
>5 years	39	40.32
BMI (in kg/m²)		
<27	37	38.54
≥27	59	61.48

Arshad et al.; AJMAH, 19(10): 1-7, 2021; Article no.AJMAH.74081



Fig. 1. Frequencies and percentages of patients according to gender



Fig. 2. Frequency and percentage of patients according to Type of Diabetes Mellitus

In our study, the overall frequency of thyroid dysfunction found in diabetic patients was 30 (31.25%) with hypothyroidism in 19 (19.79%), and hyperthyroidism in 11 (11.46%) patients and 66 (68.75%) patients were euthyroid in our study Table 2.

Stratification of thyroid dysfunction with respect to age groups and gender is shown in Table 3. Table 3 has also shown the stratification of thyroid dysfunction with respect to the type of diabetes mellitus and BMI, respectively. Stratification of thyroid dysfunction with respect to the duration of the disease is shown in Table 3.

3.2 Discussion

This paper demonstrates the importance of recognizing this interdependent relationship between thyroid disease and diabetes, which will help guide clinicians on the optimal screening and management of these conditions. We have conducted this study to determine the frequency of thyroid dysfunction in patients with DM.

Table 2. Frequency	y and percentage of
patients with the	vroid dysfunction

Thyroid dysfunction	Frequency (%)	
	yes	no
Hyperthyroidism	11 (11.46%)	83 (88.54%)
Hypothyroidism	19 (19.79%)	74 (80.21%)
Total	30 (31.25%)	66 (68.75%)

The age range in our study was from 20 to 60 years, with a mean age of 42.97 ± 10.29 years. The majority of the patients, 37.50%, were between 41 to 50 years of age. In our study, out of 96 patients, 61 (63.54%) were male, and 35 (36.46%) were females with a male to female ratio of 1.7:1. The majority of patients, 69 (71.88%), had Type II DM. In our study, the overall frequency of thyroid dysfunction found in diabetic patients was 30 (31.25%) with hypothyroidism in 19 (19.79%) and hyperthyroidism in 11 (11.46%) patients that is higher than the usual prevalence of thyroid dysfunction in the diabetic population. According to a study by Perros et al., who has

Variables	Categories	Hypothyroidism		Hyperthyroidism		Total	
	-	Yes	No	Yes	No	Yes	No
	20-30	01 (8.33%)	11 (91.67%)	01 (8.338%)	11 (91.67%)	02 (16.67%)	10 (83.33%)
Age groups	31-40	04 (14.81%)	23 (85.19%)	04 (14.81%)	23 (85.19%)	08 (29.63%)	19 (70.37%)
	41-50	08 (22.22%)	28 (77.78%)	04 (11.11%)	32 (88.89%)	12 (33.33%)	24 (66.67%)
	51-60	06 (28.57%)	15 (71.43%)	02 (9.52%)	19 (90.48%)	08 (38.10%)	13 (61.90%)
	P-value	0.463	· · · · · ·	0.920		0.626	· · · · ·
	Male	12 (19.67%)	49 (80.33%)	08 (13.11%)	53 (86.89%)	20 (32.79%)	41 (67.21%)
Gender	Female	07 (20.0%)	28 (80.0%)	03 (8.57%)	32 (91.43%)	10 (28.57%)	25 (71.43%)
	P-value	0.969	, , , , , , , , , , , , , , , , , , ,	0.501		0.668	· · · · ·
	I	03 (11.11%)	24 (88.89%)	06 (22.22%)	21 (77.78%)	09 (33.33%)	18 (66.67%)
Type of Diabetes Mellitus	II	16 (23.19%)	53 (76.81%)	05 (7.25%)	64 (92.75%)	21 (30.43%)	48 (6.57%)
	P-value	0.182	· · · · · ·	0.038		0.783	, , , , , , , , , , , , , , , , , , ,
Body Mass Index (BMI)	<27 kg/m ²	02 (5.41%)	35 (94.59%)	05 (13.51%)	32 (86.49%)	07 (18.92%)	30 (81.08%)
, , ,	≥27 kg/m²	17 (28.81%)	42 (71.19%)	06 (10.17%)	53 (89.83%)	23 (38.98%)	36 (61.02%)
	P-value	0.005	· · · · · ·	0.617		0.039	· · · · ·
	>2-5 years	10 (17.54%)	47 (82.46%)	07 (12.28%)	50 (87.72%)	17 (29.82%)	40 (70.18%)
Duration of Diabetes Mellitus	>5 years	09 (23.08%)	30 (76.92%)	04 (10.26%)	35 (89.74%)	13 (33.33%)	26 (66.67%)
	P-value	0.504	. , ,	0.760	. ,	0.716	. , ,

Table 3. Stratification of thyroid dysfunction (hypothyroidism, hypothyroidism) and overall thyroid dysfunction with respect to age groups,gender, thyroid dysfunction, Type of DM, BMI, and duration of disease

demonstrated an overall prevalence of 13.4% of thyroid diseases in diabetics with the highest prevalence in Type 1 female diabetics (31.4%) and lowest prevalence in Type 2 male diabetics (6.9%). However, these findings are contrary to our study findings, where the frequency of thyroid dysfunction was high (31.25%), and more significantly, this frequency was high in patients with Type II DM [12]. Various studies have also reported the prevalence of thyroid dysfunction with diabetes as a study in Greek diabetic patients also showed a prevalence of 12.3% [13], but this is much lower than our study findings. Similarly, another study shows the prevalence of 16% of thyroid dysfunction among Saudi patients with type 2 diabetes, validating our study findings of the increasing prevalence of thyroid dysfunction with DM and, more specifically, Type II DM [14].

In our study, hypothyroidism was the main thyroid dysfunction present in 19 (19.79%) patients with DM compared to hyperthyroidism that was present only in 11(11.46%) patients. These study findings are inconsistent with the study results from CEJ et al., which was also showing a higher frequency of hypothyroidism (26.6%) in patients with DM as compared to hyperthyroidism (19.8%) [11]. In the same vein, the study by Subekti et al. is also validating our study findings by showing the high frequency of hypothyroidism among 7 (7.59%) patients with coexisting DM as compared to hyperthyroidism in 7 (2.31%) subjects with coexisting DM [15]. Another significant finding of our study was the association of thyroid dysfunction, particularly hypothyroidism in patients with BMI \geq 27; this finding of high BMI and hypothyroidism association is in concordance with a study by Demitrost et al. in which the patients with higher BMI ≥ 25 were at increased risk of having hypothyroidism [16]. Another interesting finding in our study was an association of DM duration with thyroid dysfunction showing that patients with DM for greater than five years were at increased risk of developing thyroid dysfunction as validated by a study by Ogbonna et al. in which patients with DM greater than five years were at increased risk of developing thyroid dysfunction (OR= 3.3, p= 0.012) [17].

The limitation of this study is the small sample size and the single centered conduction of the study in the population of Rawalpindi and associated areas, which could have limited objectivity. The geographical and individual variation could impact the study findings if conducted in different hospitals of Pakistan. We recommend more studies in this subject to get more insight into these two (DM and thyroid dysfunction) interlinked diseases so that better patient's management could be carried out.

4. CONCLUSION

This study concluded that there is a high frequency of thyroid dysfunction in diabetic patients. In the same vein, hypothyroidism is the most common thyroid dysfunction associated with Type II DM. So, we recommend that there should be a routine screening of thyroid functions in every diabetic patient; thus, proper management can be done in these patients for good glycemic control and to reduce morbidity.

CONSENT

All authors declare that informed consent was obtained from each patient for publication of this research article and purpose of this study was thoroughly explained while they participated in our study.

ETHICAL APPROVAL

All authors hereby declare that all experiments have been examined and approved by the appropriate ethics committee and have therefore been performed in accordance with the ethical standards laid down in the 1964 Declaration of Helsinki."

FUNDING SOURCE

Funding was done by Clarity BioSciences, one of the nation's fastest growing companies in the pharmaceutical industry that deals in all kind of Anti Cancerous medicines.

COMPETING INTERESTS

Authors have declared that no competing interests exist.

REFERENCES

1. Schmidt AM. Highlighting Diabetes Mellitus: The Epidemic Continues. Arterioscler Thromb Vasc Biol. 2018;38:1-8.

DOI:10.1161/ATVBAHA.117.310221.

2. Petersmann A, Müller-Wieland D, Müller UA, Landgraf R, Nauck M, Freckmann G,

et al. Definition, Classification and Diagnosis of Diabetes Mellitus. Exp Clin Endocrinol Diabetes. 2019;127:1-7. DOI:10.1055/a-1018-9078.

- Knapp M, Tu X, Wu R. Vascular endothelial dysfunction, a major mediator in diabetic cardiomyopathy. Acta Pharmacol Sin. 2019;40:1-8. DOI:10.1038/s41401-018-0042-6.
- Forbes JM, Cooper ME. Mechanisms of diabetic complications. Physiol Rev. 2013, 93:137-188. 10.1152/physrev.00045.2011.
- Ogbonna SU, Ezeani IU, Okafor CI, Chinenye S. Association between glycemic status and thyroid dysfunction in patients with type 2 diabetes mellitus. Diabetes Metab Syndr Obes. 2019; 12: 1113-1122. DOI:10.2147/DMSO.S204836.
- Yang N, Zhang DL, Hao JY, Wang G. Serum levels of thyroid hormones and thyroid-stimulating hormone in patients with biogenic and hyperlipidaemic acute pancreatitis: Difference and value in predicting disease severity. J Int Med Res. 2016;44:267-277.
 - DOI:10.1177/0300060515618052.
- Gierach M, Gierach J, Junik R. Insulin resistance and thyroid disorders. Endokrynol Pol. 2014, 65:70-76. DOI:10.5603/EP.2014.0010.
- Chen L, Zhang M, Xiang S, Jiang X, Gu H, Sha Q, et al. Association Between Thyroid Function and Body Composition in Type 2 Diabetes Mellitus (T2DM) Patients: Does Sex Have a Role? Med Sci Monit. 2021;27:927440. DOI:10.12659/MSM.927440.
- Gutiérrez-Vega S, Armella A, Mennickent D, Loyola M, Covarrubias A, Ortega-Contreras B, et al. High levels of maternal total tri-iodothyronine, and low levels of fetal free L-thyroxine and total triiodothyronine, are associated with altered deiodinase expression and activity in placenta with gestational diabetes mellitus. PLoS One. 2020;15:0242743.

DOI:10.1371/journal.pone.0242743.

- Jali MV, Kambar S, Jali SM, Pawar N, Nalawade P. Prevalence of thyroid dysfunction among type 2 diabetes mellitus patients. Diabetes Metab Syndr. 2017;11:105-108. DOI:10.1016/j.dsx.2016.12.017.
- Udiong CE, Udoh AE, Etukudoh ME. Evaluation of thyroid function in diabetes mellitus in Calabar, Nigeria. Indian J Clin Biochem. 2007;22:74-78. DOI:10.1007/BF02913318.
- Perros P, McCrimmon RJ, Shaw G, Frier BM. Frequency of thyroid dysfunction in diabetic patients: value of annual screening. Diabetic medicine. 1995;12 :622-627. DOI:10.1111/j.1464-5491.1995 tb00553.x.
- Papazafiropoulou A, Sotiropoulos A, Kokolaki A, Kardara M, Stamataki P, Pappas S. Prevalence of thyroid dysfunction among greek type 2 diabetic patients attending an outpatient clinic. Journal of clinical medicine research. 2010;2:75-78.

DOI:10.4021/jocmr2010.03.281w.

- 14. Akbar DH, Ahmed MM, Al-Mughales J. Thyroid dysfunction and thyroid autoimmunity in Saudi Type 2 diabetics. Acta Diabetol. 2006;43:14-8. DOI:10.1007/s00592-006-0204-8.
- 15. Subekti I, Pramono LA, Dewiasty E, Harbuwono DS. Thyroid Dysfunction in Type 2 Diabetes Mellitus Patients. Acta Med Indones. 2017;49:314-323.
- 16. Demitrost L, Ranabir S. Thyroid dysfunction type 2 diabetes in mellitus: A retrospective study. Indian J Endocrinol Metab. 2012:16:334-335.

DOI:10.4103/2230-8210.104080.

 Ogbonna SU, Ezeani IU. Risk Factors of Thyroid Dysfunction in Patients with Type 2 Diabetes Mellitus. Front Endocrinol (Lausanne). 2019;10:440. DOI:10.3389/fendo.2019.00440.

© 2021 Arshad et al.; This is an Open Access article distributed under the terms of the Creative Commons Attribution License (http://creativecommons.org/licenses/by/4.0), which permits unrestricted use, distribution, and reproduction in any medium, provided the original work is properly cited.

Peer-review history: The peer review history for this paper can be accessed here: https://www.sdiarticle4.com/review-history/74081