

Assessment of Psychological Manifestations and Prevalence of Diabetes Mellitus in Patients with Thyroid Dysfunction

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Abstract

Diabetes mellitus and thyroid dysfunction are quite regularly associated. In diabetic patients the chronicity of thyroid dysfunction is two to three times greater than in patients with no diabetes mellitus, it increases with age and is significantly impacted by autoimmune diabetes and by female gender.

Material and Methods: Patients of all age groups who are known to have thyroid dysfunction were included in our study and also who are known to have thyroid dysfunction together with diabetes mellitus were included in our study.

Results: An entire of 291 cases were collected for assessment of psychological symptoms and a total of 62 cases were collected for determining the prevalence of diabetes mellitus in thyroid patients. Research work was accomplished in patients who encountered the inclusion criteria and found that the utmost frequent manifestations were uniform in patients with subclinical and undisguised thyroid disease and also identified that the prevalence of thyroid dysfunction in patients with diabetes mellitus was greater than the prevalence of diabetes mellitus in patients with thyroid dysfunction.

Conclusion: In conclusion, the levels of depression and stress were found to be little higher in case of hyperthyroidism patients when compared to hypothyroidism patients where the levels of anxiety were found to be almost similar in both cases. Instance of thyroid dysfunction in patients with diabetes mellitus was more than the occurrence of diabetes mellitus in patients with thyroid dysfunction.

Keywords: Thyroid dysfunction, Anxiety, Depression, Stress and Diabetes mellitus.

Introduction

Iodide uptake into the thyroid, oxidation of iodide, iodination and coupling of tyrosyl residues together with cellular uptake, proteolysis of thyroglobulin are the steps involved in synthesis and storage of thyroid hormones whereupon followed by their secretion [1]. Sulfhydryl groups are required for peripheral transformation of T₄ to T₃ [2]. L-type amino acid transporters and organic anion transporters are involved in uptake and transport of thyroid hormones [3]. The receptors of thyroid hormones belong to nuclear receptor super family that functions by regulating transcription [4]. The principal regulator of serum TSH levels at pituitary is thyroid hormone negative feedback [5]. T₄ and T₃ play a vital role during pregnancy, neural ballooning and demarcation [6]. The major physiologic effect of these hormones is that they stimulate the metabolism as well as metabolic rate of lipids, proteins and carbohydrates [7]. Mental symptoms like anxiety and depression gets developed in many

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patients with thyroid dysfunction. Psychiatric symptoms are mainly due to adrenergic hyperactivity and in vulnerable patients these may be due to thyroid autoimmunity process itself [8]. High rate of depression has been observed in patients with primary thyroid disease [9]. Typical elevated cortisol levels and impaired dexamethasone suppression are because of a functional disconnection of the hypothalamus with impairment of inhibitory glucocorticoid feedback pathway from the hippocampus to the hypothalamus [10]. Thyroid function gets dismal in stressful conditions. Thyroid-stimulating hormone secretion is inhibited by stress by the action of glucocorticoids on central nervous system [11]. So there is a need to remark the changes in these variables. Untreated thyroid disorders influence the control of diabetes in many patients [12, 13]. The most likely mechanism that paramount to T_2DM in thyroid dysfunction could be ascribed to perturbed genetic expression of redundancy of genes along with physiological deformity leading to plunged glucose usage and jettisoning in muscles, deluge of hepatic glucose output and boomed absorption of splanchnic glucose. These factors lead to insulin resistance. Insulin resistance is also kindred with thyroid dysfunction. Hypothyroidism and hyperthyroidism have been correlated with insulin intransigence which has been appeared to be the crucial cause of impeded glucose metabolism in T_2DM [14]. The pathophysiological mechanisms shrouding this linked regulation are increasingly being untangled. They are epitomized in the proclamation of 5' Adenosine Monophosphate-Activated Protein Kinase (AMPK), a central target not only for the intonation of insulin sensitivity but also for the evaluation of thyroid hormones on appetite and energy disbursement [15]. Subclinical hypothyroidism is linked with an enhanced risk of symptomatic hypoglycaemia [16]. There is a compelling confirmation from independent research that T_3 directly elevates islet β -cell mass via thyroid hormone receptor α -dependent pathways [17]. Furthermore, insulin emanation from β -cells is likely controlled by the curtailed mitochondrial T_3 receptor p43 [18]. These changes have been kindred with a decline in specific glucose transporters, namely GLUT2 and Kir6.2 and may thus be more broadly conciliated by the control of intracellular glucose availability which may have indications for other actions of T_3 [19]. FT_3 assists expression of important proteins involved in both glucose and lipid metabolism that may impact insulin secretion [20]. Declined FT_3 and FT_3/FT_4 ratios and elevated FT_4 levels were independently related to the prevalence of T_2DM among the adult population [21]. Autoimmune thyroid disorders frequently exist in children and adolescents with type 1 diabetes mellitus. The generality of these disorders was 17.6% and of those, the most common was chronic autoimmune thyroiditis. Microsomal auto antibodies are more accurately interconnected with the presence of chronic autoimmune thyroiditis when compared to thyroglobulin auto antibodies. The largest prevalent immunological mechanisms influencing diabetic patients were autoimmune thyroid disorders. The thyroid status of many patients with positive markers was subclinical hyperthyroidism (3%), overt hyperthyroidism (6%), subclinical hypothyroidism (11%), overt hypothyroidism (3%) and euthyroidism (77%). No strong amalgamations of thyroid autoimmunity

and other auto immunological disorders like celiac disease or existence of other autoimmune antibodies were identified [22]. The residence of inhibitor of extra thyroidal conversion of thyroxine (T_4) to triiodothyronine (T_3), dysfunction of the hypothalamo-hypophysial-thyroid axis and the existence of thyroid hormone binding inhibitor are contemplated to be tangled in abnormal thyroid function in patients with diabetes mellitus [23]. So, there is a necessity to remark the association between diabetes mellitus and thyroid dysfunction.

Material and Methods

The research work was overseen in patients addressing the outpatient department of endocrinology. A total of 238 subjects for assessment of psychiatric symptoms and 45 subjects to determine the prevalence of diabetes mellitus in thyroid patients were encompassed in the study. Patients with different semblance of thyroid dysfunction who were determined by an endocrinologist were grouped as overt hypothyroidism (n=20(male=03; female=17)), subclinical hypothyroidism (n=190(male=07; female=183)), overt hyperthyroidism (n=22(male=01; female=21)) and subclinical hyperthyroidism (n=06(male=0; female=06)). IHEC approval was obtained after submission of protocol and IHEC No. is MGM/VCOP/PHARMD/017/2018. Patients were explained about the study and informed consent forms were sought by explaining them in their local language. The study period is 6 months.

Inclusion criteria: Males and females of all age groups diagnosed to have thyroid dysfunction and thyroid dysfunction together with diabetes mellitus were included in our study.

Exclusion criteria: Pregnant women and patients who are already known cases of anxiety and depression were excluded.

Study design: It is a prospective, observational study conducted in outpatient department of endocrinology at Samraksha Diabetes, Thyroid and Endocrine Superspeciality Hospital and Research Centre, Warangal.

Clinical response assessment: The levels of anxiety and depression were assessed by Hamilton Rating Scale and stress was assessed by Perceived Stress Scale. Patients with parameters of metabolic control such as $HbA_{1c} > 6\%$, high fasting and post lunch blood glucose levels, TSH $> 4.20 \mu UI/L$, $FT_4 < 0.9 ng/dl$ (in case of hypothyroidism) and TSH $< 0.27 \mu UI/L$, $FT_4 > 1.7 ng/dl$ (in case of hyperthyroidism) were contemplated. Autoimmunity was undaunted where anti-TPO levels were higher than 34 IU/ml.

Statistical analysis: Data scrutinization was executed using the excel sheet 2007. All variables were demonstrated as mean \pm SD. Assessment values were obtained during the period of treatment. The figure 1 shows the scales and clinical parameters used while assessing the patients.

Results

The values in the table 1 indicate the mean and standard deviation of scores of anxiety, depression and stress in

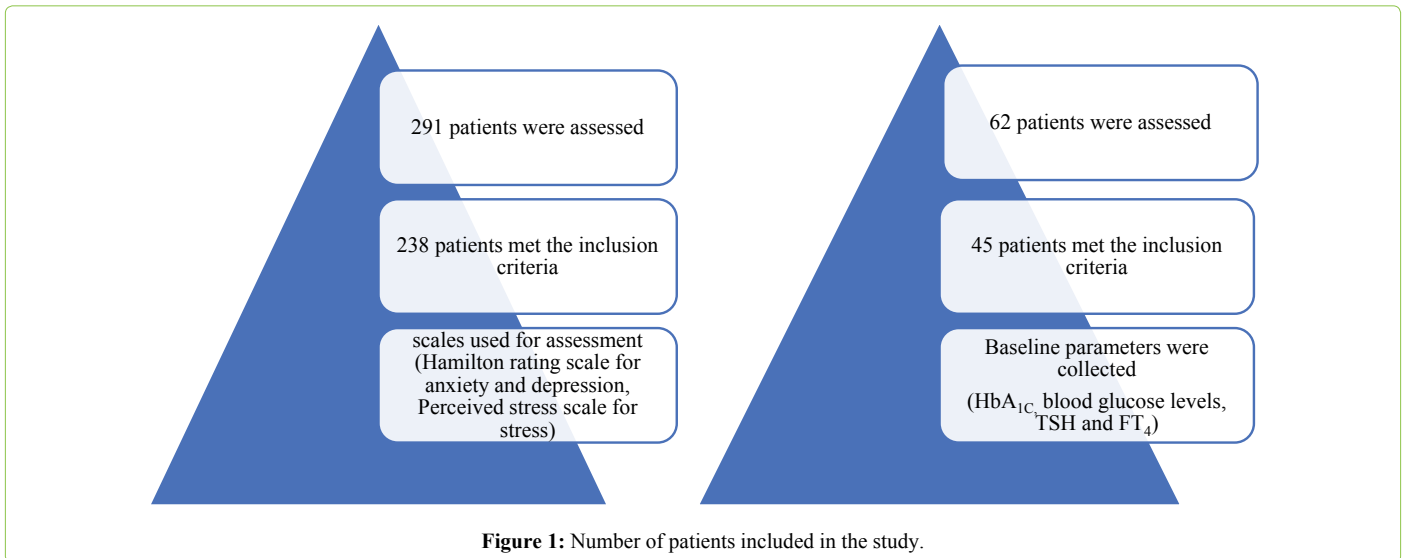


Figure 1: Number of patients included in the study.

Table 1: Scores of anxiety, depression and stress.

	Hypothyroidism (mean ± SD)	Hyperthyroidism (mean ± SD)	Control (mean ± SD)
Scores of anxiety (HARS)	7.7±3.9	7.5±4.3	5.0±1.9
Scores of depression (HDRS)	7.8±4.6	8.3±5.1	5.3±2.2
Scores of stress (PSS)	8.2±3.0	8.9±2.6	6.8±2.3

*HARS: Hamilton Anxiety Rating Scale; HDRS: Hamilton Depression Rating Scale; PSS: Perceived Stress Scale. Higher scores in HARS, HDRS and PSS reveal more anxiety, depression and stress.

males and females together in case of hypothyroidism, hyperthyroidism and control after receiving the treatment.

In figure 2 both subclinical and overt hypothyroidism was identified to be more common in females than in males. On terms of subclinical hypothyroidism females engross 96% and males 4%. On other hand in case of overt hypothyroidism females beguile 85% and males 15%.

From figure 3 it is clear that hyperthyroidism is typical in female's juxtaposition to males. Subject to subclinical hyperthyroidism females occupy 100% whereas males 0%. In case of overt hyperthyroidism females hold 95% and males 5%.

As per figure 4 most patients of hypothyroidism and hyperthyroidism experienced mild anxiety. 205 hypothyroidism patients and 27 hyperthyroidism patients are suffering from mild anxiety. Severe anxiety was not observed in hyperthyroidism patients.

As per figure 5 higher percentage of patient population does not have any depression while minute proportion has mild depression. The fraction of people experiencing moderate and severe depression is found to be equal in case of both hypothyroidism and hyperthyroidism.

As per figure 6 low stress was identified in significant proportion of patient population of hypothyroidism and hyperthyroidism where as high perceived stress was not seen in both cases. Moderate stress was identified only in case of hypothyroidism.

As per figure 7 a total of 45 patients were identified to have thyroid dysfunction plus diabetes mellitus out of which 4 patients were male and 41 patients were female.

Among all the cases, 93% were type 2 diabetes mellitus plus hypothyroidism, 5% were type 1 diabetes mellitus together with hypothyroidism and 2% were type 2 diabetes mellitus plus hyperthyroidism which can be seen in figure 8 beneath.

In a total of 45 patients, 27 patients initially have diabetes mellitus and then developed thyroid dysfunction and 18 patients initially have thyroid dysfunction and then developed diabetes mellitus as shown in figure 9.

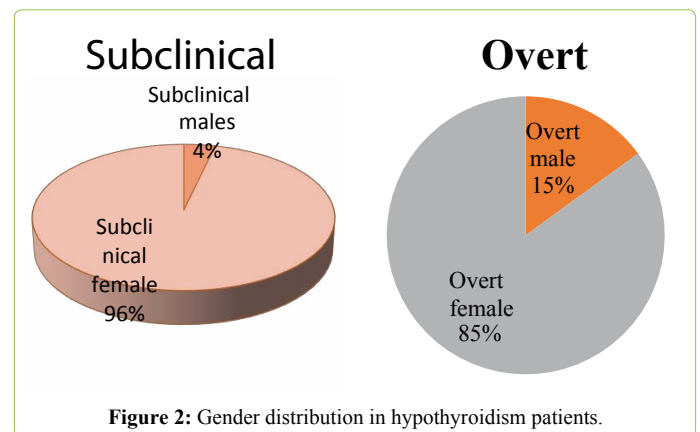


Figure 2: Gender distribution in hypothyroidism patients.

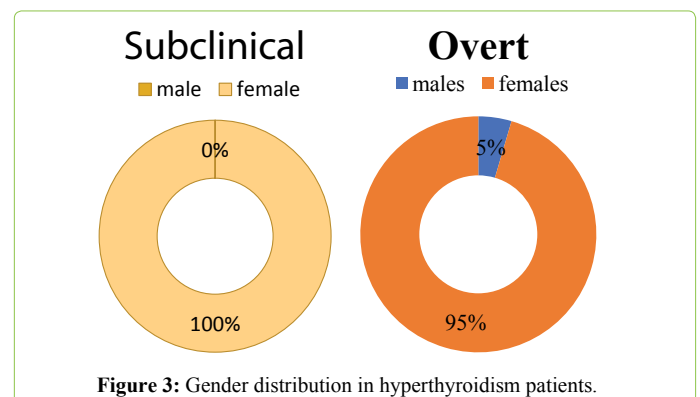


Figure 3: Gender distribution in hyperthyroidism patients.

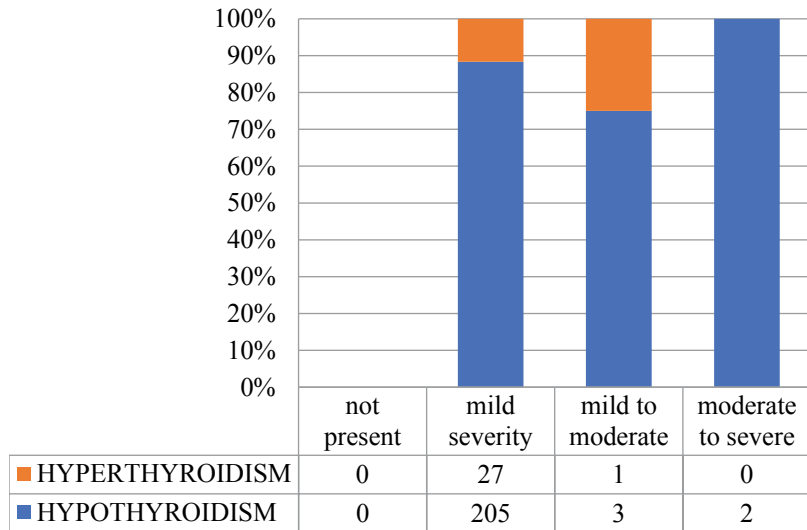


Figure 4. Prevalence of anxiety in hypothyroidism and hyperthyroidism.

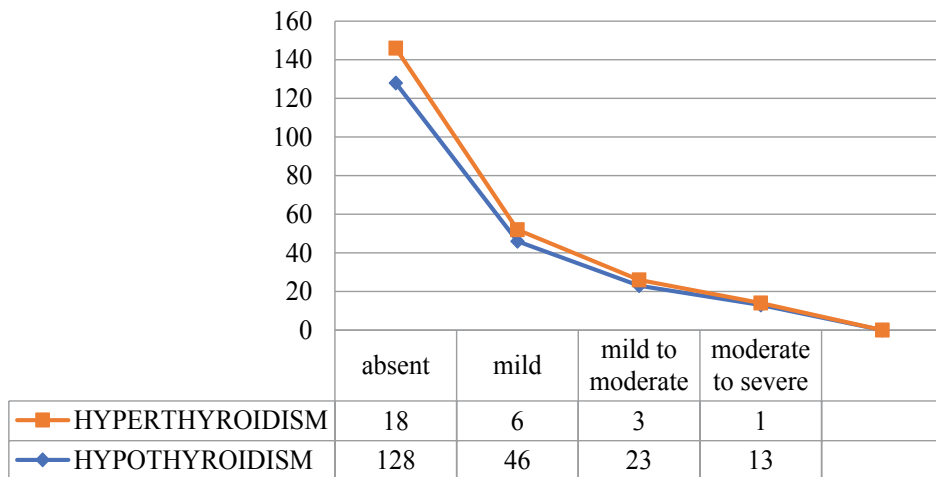


Figure 5: Prevalence of depression in hypothyroidism and hyperthyroidism

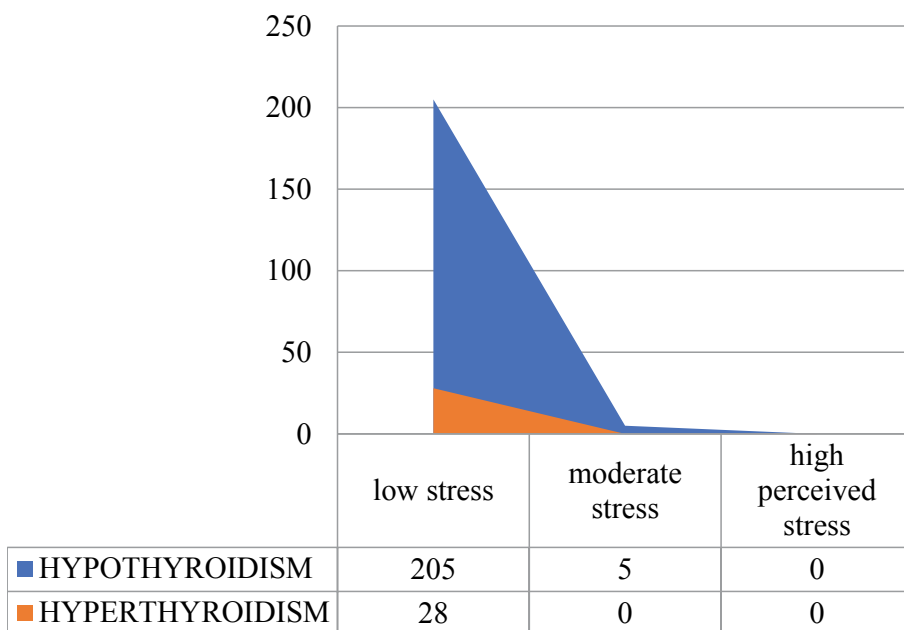
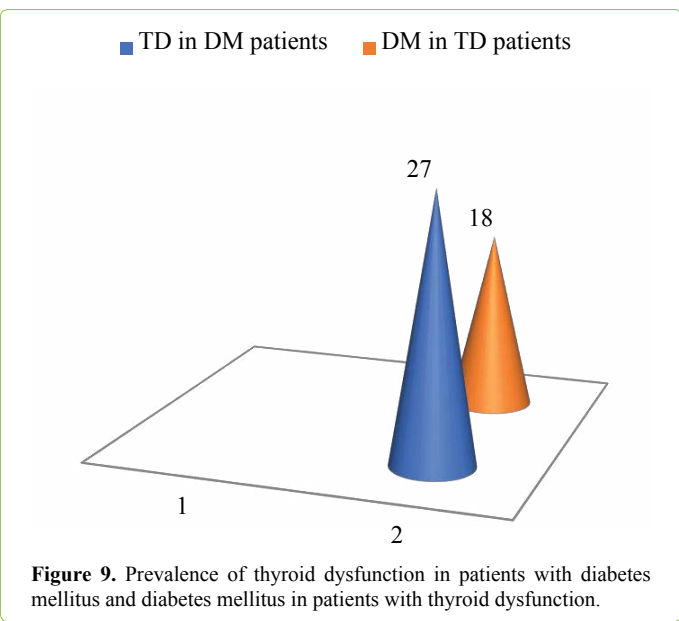
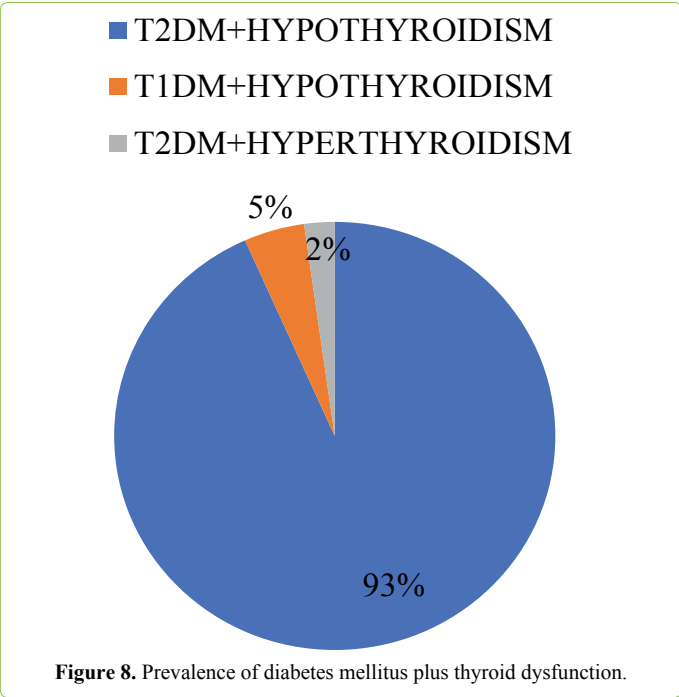
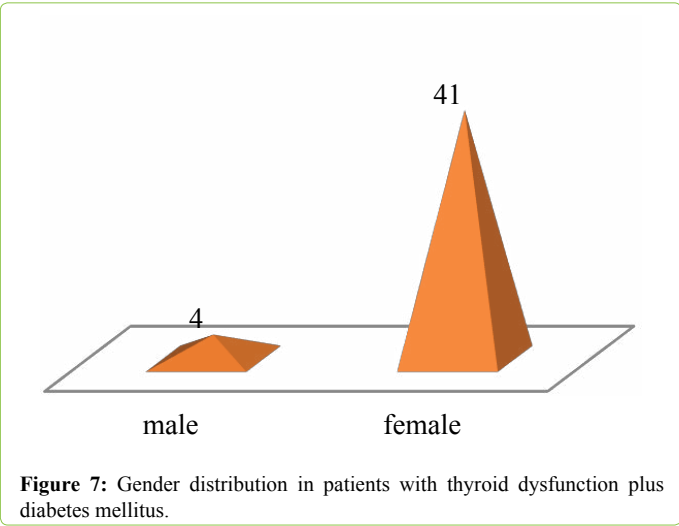


Figure 6. Prevalence of stress in hypothyroidism and hyperthyroidism.



Discussion

In this research work, patients with thyroid dysfunction were scrutinized in terms of depression, anxiety and stress. In previous studies, it was reported that these variables were mostly found in subclinical hyperthyroidism patients [24]. In our study, it is notable that the levels of these variables are almost similar in both hypothyroidism and hyperthyroidism patients. It is postulated that manifestations of thyroid dysfunction, anxiety and depression contemplate central nervous system β -adrenergic impairment [25, 26]. It is delineated that in utmost cases of hypothyroidism and hyperthyroidism, psychiatric symptoms retained with treatment of the thyroid dysfunction [27]. Stress levels were found to be indistinguishable in case of both hypothyroidism and hyperthyroidism. In our study, it is remarkable that levels of psychological symptoms decline significantly as thyroid function returned to normal, despite the actuality that no psychiatric treatment was applied. The symptoms may be abolished by L-thyroxine therapy [28, 29, 30]. In our study, there were outstanding improvements in levels of anxiety, depression and stress in contrast to the levels at the inception of the treatment. Also, the prevalence of thyroid dysfunction in patients with diabetes mellitus and diabetes mellitus in patients with thyroid dysfunction was scrutinized. It is noticeable from the existing literature that insulin resistance bears a crucial role in associating diabetes mellitus and thyroid dysfunction. Few studies have embellished the relationship between TH, TSH and T₂DM and signified that T₂DM is negatively related to FT₃ and positively interrelated to FT₄ in males as well as females and women who are with type 1 diabetes mellitus are at greater risk of symptomatic postpartum thyroid dysfunction and accordingly may advantage from routine thyroid function screening at postpartum visits [31, 32]. Our outcomes divulged homogeneous relationship between TSH, T₂DM and TH in both females and males.

Conclusion

In conclusion, the thyroid dysfunction (hypothyroidism and hyperthyroidism) was more habitually observed in females than in males. The extent of anxiety was identified to be almost similar in two cases but the levels of depression and stress were little higher in case of hyperthyroidism patients when compared to hypothyroidism patients. Prevalence of thyroid dysfunction accompanying diabetes mellitus was observed more in females than in males. Instance of thyroid dysfunction in patients with diabetes mellitus was more than the prevalence of diabetes mellitus in patients with thyroid dysfunction. So there is a need for early diagnosis of psychological symptoms and prevalence of diabetes mellitus in patients with thyroid dysfunction.

The authors declare that there's no conflict of interest concerning the publication of paper.

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