FREQUENCY OF THYROID DYSFUNCTION IN CHRONIC KIDNEY DISEASE PATIENTS

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ABSTRACT

Objective: To determine the frequency of thyroid dysfunction in chronic kidney disease patients.

Methods: A Cross-sectional Study conducted at Department of Medicine, Bahawal Victoria Hospital, Bahawalpur. Duration of the study was 20th October 2018 to 19th April 2019. Seventy-two patients with chronic kidney disease, 20 to 60 years of age and of both genders were included. Patients with known thyroid disorders or taking medication that can affect thyroid function (dopamine antagonists, antiepileptic, oral contraceptives, lithium, and glucocorticoids) were excluded. Venous blood sample was taken and sent to the laboratory for thyroid function tests.

Results: Age of the patients included in this study was from 20 to 60 year and mean age was 41.97 ± 9.76 years. Most of them 42 (58.33%) were between 41 to 60 years of age. Out of 72 patients, 44 (61.11%) were male and 28 (38.89%) were female with male to female ratio of 1.6:1. Mean duration of disease in our study was 5.71 ± 2.50 years. Mean BMI was 27.75 ± 3.01 kg/m2.Thyroid dysfunction was present in 23 (31.94%) patients of chronic kidney disease.

Conclusion: Thyroid dysfunction is frequently found in chronic kidney disease patients.

Keywords: chronic kidney disease, thyroid dysfunction, hypothyroidism.

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INTRODUCTION

Chronic kidney disease (CKD) is characterized by irreversible and progressive decline in kidney function secondary to a group of pathophysiologic processes. Kidney has excretory metabolic and synthetic functions. When kidney stops working many unwanted toxic substances accumulate in body leading to different signs and symptoms.¹ Thyroid hormone is also metabolized and excreted by kidneys so when kidneys stop working levels of thyroid hormones are also disturbed. ²⁻³

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Hypothyroidism, both overt and subclinical, is more prevalent in CKD as compared to hyperthyroidism, ⁴ with subclinical hypothyroidism being the most common ⁵. In a study almost half patients of CKD were found to have thyroid dysfunction⁶. Whereas in dialysis population the prevalence of subclinical and overt hypothyroidism was 26.6%^{7.} While in another study done in India subclinical hypothyroidism was found in 24.8% of the dialysis population.⁸ A study done in peritoneal dialysis (PD) patients reported prevalence of subclinical hypothyroidism and subclinical hyperthyroidism 15.6% and 4.1% respectively⁹.

As the chronic kidney disease goes on increasing in our population and thyroid dysfunction is associated with CKD, so the purposed significance of my study was to evaluate the frequency of thyroid dysfunction in chronic kidney disease patients. The above-mentioned studies have shown variability in frequency of thyroid dysfunction in CKD patients among different ethnic groups, so there is a need of a study in our geographical area which will provide the local magnitude of the problem. As routinely thyroid functions in CKD patients are not assessed in our general practice and failure to recognize the presence of abnormal thyroid hormone level in CKD may be a cause of poor prognosis, so the results of my study will help the clinicians to design a proper management protocol for these patients in order to reduce the morbidity of our population.

METHODS

It was a descriptive, Cross-sectional survey. All patients with chronic kidney disease as per-operational definition, between ages of 20-60 years, of both genders who were admitted in Department of Medicine, Bahawal Victoria Hospital, Bahawalpur were included. Study was conducted over a six-month period from 20th October 2018 to 19th April 2019. Patients were excluded if were taking medication that can affect thyroid functions (dopamine antagonists, contraceptives, antiepileptic, oral lithium, glucocorticoids) as assessed on history and medical record. Pregnant patients and Patients with known thyroid disorders evaluated on history and medical record were excluded as well. All patients participated in this study after an Informed consent. Sampling technique was non-probability. consecutive sampling. Sample size of 72 was calculated by using $n=z^2pq/d^2$ where Z=1.96, p=24.8%⁸, q=100-p, d=10%.

Approval was taken from the hospital committee of faculty members. Venous blood sample was taken and sent for thyroid function tests to the laboratory and thyroid dysfunction (as per-operational definition) was noted. All the data including the demographic data (age, gender, BMI, diabetes mellitus and hypertension) obtained from patients was recorded on the predesigned questionnaire. Chronic Kidney Disease was labeled if serum creatinine was above 1.3mg/dl on laboratory examination for last 12 weeks. Any of these Chronic Kidney Disease Stages was included:

- Stage 1: GFR > 90 ml/min.
- Stage 2: GFR between 60-90 ml/min.
- Stage 3: GFR between 30-59 ml/min.
- Stage 4: GFR between 15-29 ml/min.
- Stage 5: GFR < 15 ml/min

Patient was considered hypothyroid if TSH >5.2 mIU/L, FT₃<1.5 pg/ml and FT₄<0.8 pg/ml and was considered hyperthyroid if TSH <0.2 mIU/L and FT₃>4.2 pg/ml, FT₄>1.68 pg/ml. SCH was labelled if serum

thyroid stimulating hormone (TSH) was above the normal limit with normal level of free thyroxine (T4).

Formula used for BMI calculation was;

BMI = weight in kilograms / height in meters² (as measured by stadiometer)

>27 was taken as obese and \leq 27 as non-obese.

Data entry and analysis was done using SPSS version 22.0. Mean and standard deviation for the age of the patients, duration of CKD and BMI were calculated by applying descriptive statistics. We calculated percentages and Frequencies for categorical variables such as gender, stage of CKD (I/II/III/IV/V), diabetes mellitus (yes/no), hypertension (yes/no) and on hemodialysis (yes/no). Effect modifiers like age, gender, duration of CKD, BMI, stage of CKD (I/II/III/IV/V), diabetes mellitus (yes/no) and on hemodialysis (yes/no). Effect modifiers like age, gender, duration of CKD, BMI, stage of CKD (I/II/III/IV/V), diabetes mellitus (yes/no) and on hemodialysis (yes/no) were controlled by chi square test & fisher's exact test and if p-value was ≤ 0.05 , it was considered as significant.

RESULTS

Mean age of patients in our study was 41.97 ± 9.76 years ranging from 20 to 60 years. Most of them were between 41 to 60 years of age. Out of 72 patients, 44 (61.11%) were male and 28 (38.89%) were females with male to female ratio 1.6:1. Distribution of patients according to stage of CKD is shown in Figure I.

Mean duration of disease in our study was 5.71 \pm 2.50 years. Mean BMI was 27.75 \pm 3.01 kg/m².out of 72 patients, 35 patients (48.61 %) were diabetic and 44 patients (61.11%) were hypertensive. In our study, frequency of thyroid dysfunction in chronic kidney disease patients was found in 23 (31.94%) patients.





Table 1 has shown the stratification of thyroid dysfunction with respect to stage of CKD and

duration of CKD. Stratification of thyroid dysfunction with respect to BMI, DM & HTN is shown in Table 2.

Table I: Stratification of thyroid dysfunction with respect to stage and duration of CKD.

Thyroid					
Stage	dysfunction		p-value		
	Yes	No			
Ι	03	02			
II	09	20			
III	08	11	0.195		
IV	02	14			
V	01	02			
Duration					
≤5	13	33	0.373		
>5	10	16			

Table II: Stratification of thyroid dysfunction with respect to DM, HTN & BMI.

	Thyroid		
_	dysfunction		p-value
	Yes	No	
DM			
Yes	09	26	0.270
No	14	23	0.270
Hypertension			
Yes	16	28	0.212
No	07	21	0.515
BMI (kg/m ²)			
≤27	09	25	0.246
>27	14	24	0.340

DISCUSSION

Thyroid hormone has very important role in maintaining the normal metabolism of body. It also affects functions of other hormones. As the kidney function is compromised metabolism of thyroid hormone is also affected leading to thyroid abnormalities. Inversely thyroid hormone abnormalities, if left untreated, also increase the chance of developing kidney dysfunction.so proper diagnosis and treatment of thyroid dysfunctions are very important in the setting of kidney disease¹⁰.

We conducted this study to determine the frequency of thyroid dysfunction in chronic kidney disease patients. Age of the patients included in this study was from 20 to 60 year and mean age was 41.97 ± 9.76 years. Most of them 42 (58.33%) were between 41 to 60 years of age. 44 patients (61.11%) were male and 28 patients (38.89%) were females with male to female ratio 1.6:1. In our study, frequency of thyroid dysfunction in chronic kidney disease patients was found in 23 (31.94%) patients.

A Survey concluded the data of almost fifteen thousand adults which shows the inverse relation of GFR with hypothyroidism i-e with progressive reduction in GFR chance of hypothyroidism increases¹¹. Chonchol M et al concluded the same in his study¹².

In CKD altered metabolism of thyroid hormones affects the activity and blood levels of hormones¹³. Overt biochemical hypothyroidism, subclinical hypothyroidism (SCH), and low triiodothyronine (T3) syndrome have been reported in CKD population with subclinical hypothyroidism being the most common¹⁴⁻¹⁶. Hypothyroidism increases the risk of cardiovascular diseases as well making it even more important to diagnose and manage in order to reduce the overall mortality in CKD population¹⁷. As far as hyperthyroidism is concerned many studies done in CKD patients found no difference in its prevalence from general population¹⁸.

Many studies have shown that goiter and thyroid abnormalities are common in end stage renal disease¹⁹⁻²³. Some studies suggested thyroid abnormalities as an independent predictor of mortality in ESRD patients²⁰⁻²² probably because of chronic inflammation²³. There are several proposed hypothesis to explain the mechanism connecting the kidney failure with thyroid malfunction but none of them proved to be the rite one.in one study hyponatremia has been observed due to abnormal retention of water in distal nephron in patients of myxedema.²⁴. Moreover, cardiac dysfunction secondary to hypothyroidism may also cause changes in intrarenal hemodynamics leading to kidney dysfunction.

In patients with early CKD the prevalence of thyroid abnormalities is very less studied.in a nationally representative cohort of U.S. adults it was observed hypothyroidism ,both subclinical and clinical, has a direct correlation with decline in GFR with prevalence of more than 20 % in late CKD.²¹

CONCLUSION

This study concluded that frequency of thyroid dysfunction in chronic kidney disease patients is quite high. So, we recommend that thyroid functions in CKD patients should be assessed routinely for its early recognition and management in in order to reduce the morbidity of these particular patients.

CONFLICT OF INTEREST

None

ETHICAL APPROVAL

The study was approved from Institutional Ethical Review Board of Quaid-e-Azam Medical College, Bahawalpur, Pakistan, vide Reference No. 449/DME/ QAMC Bahawalpur dated November 30, 2020.

REFERENCES

- Inker LA, Astor BC, Fox CH, Isakova T, Lash JP, Peralta CA, et al. KDOQI US commentary on the 2012 KDIGO clinical practice guideline for the evaluation and management of CKD. Am J Kidney Dis. 2014;63(5):713–735.
- 2. Rajagopalan B, Dolia PB, Arumalla VK. Renal function markers and thyroid hormone status in undialyzed chronic kidney disease. Al Ameen J Med Sci. 2013;6(1):70–74.
- 3. Jeena EJ, Malathi M, Sudeep K. A hospital-based study of anti-TPO titer in patients with thyroid disease. Muller J Med Sci Res. 2013;4(2):74–77.
- 4. Sekhar H. Thyroid dysfunction in chronic kidney disease. Indian J Appl Res. 2016;6(10):348-349.
- Khatiwada S, Rajendra KC, Gautam S, Lamsal M, Baral N. Thyroid dysfunction and dyslipidemia in chronic kidney disease patients. BMC Endocr Disord. 2015;15:65.
- Sinha V, Kumar A, Kachhawa P, Agrawal S. Thyroid dysfunction and dyslipidemia in patients with chronic kidney diseases. Int J Med Sci Public Health. 2016;5:2597-2603.
- Paudel K. Prevalence and clinical characteristics of hypothyroidism in a population undergoing maintenance hemodialysis. J Clin Diagn Res. 2014; 8(4):MC01–4.
- Shantha GPS, Kumar AA, Bhise V, Khanna R, Sivagnanam K, Subramanian KK. Prevalence of subclinical hypothyroidism in patients with end-stage renal disease and the role of serum albumin: a crosssectional study from South India. Cardiorenal Med. 2011;1(4):255–260.
- 9. Ng YY, Wu SC, Lin HD, Hu FH, Hou CC, Chou YY, et al. Prevalence of clinical and subclinical thyroid disease in a peritoneal dialysis population. Perit Dial Int. 2012;32(1):86–93.
- Hollowell JG, Staehling NW, Dana Flanders W. Serum TSH, T4, and thyroid antibodies in the United States population (1988 to 1994): National Health and Nutrition Examination Survey (NHANES III) Journal of Clinical Endocrinology and Metabolism. 2002;87(2):489–499.
- 11. Lo JC, Chertow GM, Go AS, Hsu CY. Increased prevalence of subclinical and clinical hypothyroidism in persons with chronic kidney disease. Kidney Int 2005;67:1047-1052.

- Chonchol M, Lippi G, Salvagno G, Zoppini G, Muggeo M, Targher G. Prevalence of subclinical hypothyroidism in patients with chronic kidney disease. Clin J Am Soc Nephrol 2008;3:1296-1300.
- 13. Den Hollander JG, Wulken RW, Mantel MJ, Berghout A. Correlation between severity of thyroid dysfunction and renal function. Clin Endocricnol 2005;62:423-427.
- Chonchol M, Lippi G, Salvagno G, Zoppini G, Muggeo M, Targher G. Prevalence of subclinical hypothyroidism in patients with chronic kidney disease. Clin J Am Soc Nephrol 2008;3:1296-1300.
- 15. Fan J, Yan P, Wang Y, Shen B, Ding F, Liu Y. Prevalence and clinical significance of low T3 syndrome in non-dialysis patients with chronic kidney disease. Med Sci Monit 2016;22:1171-1179.
- Chandra A. Prevalence of hypothyroidism in patients with chronic kidney disease: A cross-sectional study from North India. Kidney Res Clin Pract 2016;35:165-168.
- 17. Duntas LH, Wartofsky L. Cardiovascular risk and subclinical hypothyroidism: Focus on lipids and new emerging risk factors. What is the evidence? Thyroid 2007;17:1075-1087.
- Mohamedali M, Maddika SR, Vyas A, Iyer V, Cheriyath P. Thyroid disorders and chronic kidney disease. Int J Nephrol 2014; 2014: 520281.
- 19. Lim VS: Thyroid function in patients with chronic renal failure. Am J Kidney Dis. 2001;38[Suppl 1]:80–84.
- Zoccali C, Tripepi G, Cutrupi S, Pizzini P, Mallamaci F. Low triiodothyronine: a new facet of inflammation in end-stage renal disease. J Am Soc Nephrol. 2005;16 :2789–2795.
- 21. Zoccali C, Benedetto F, Mallamaci F, Tripepi G, Cutrupi S, Pizzini P, et al. Low triiodothyronine and cardiomyopathy in patients with end-stage renal disease. J Hypertens. 2006;24 :2039–2046.
- Zoccali C, Mallamaci F, Tripepi G, Cutrupi S, Pizzini P. Low triiodothyronine and survival in end-stage renal disease. Kidney Int. 2006;70:523–528.
- 23. Carrero JJ, Qureshi AR, Axelsson J, Yilmaz MI, Rehnmark S, Witt MR, et al. Clinical and biochemical implications of low thyroid hormone levels (total and free forms) in euthyroid patients with chronic kidney disease. J Intern Med. 2007;262 :690–701.
- 24. Bradley SE, Stéphan F, Coelho JB, Réville P. The thyroid and the kidney. Kidney Int. 1974;6:346–365.

AUTHOURS CONTRIBUTIONS

- AM: Data Collection, Literature Review
- **RA:** Manuscript Writing, Statistical Analysis
- MS: Manuscript Writing
- RH, SN: Data Collection
- **ZF:** Data Collection, Topic Selection