



## A Study of Renal Manifestations in Hypothyroid Patients Found During Pre Anesthetic Check Up

### KEYWORDS

hypothyroidism , renal functions , anesthesia .

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**ABSTRACT** Introduction: Thyroid hormone influences renal development , kidney structure ,

renal hemodynamics ,GFR , the functions of many transport system along nephron , sodium and water homeostasis . Disorder of thyroid function can result from any abnormality that leads to insufficient synthesis of thyroid hormone .The most common primary hypothyroidism is linked to immune mediated glomerular injury and alteration in the production of thyroid hormone .The presence of hypometabolic state thus necessitates careful perioperative renal ,cardio-vascular monitoring and judicious use of anaesthetic drugs.

**Aim :**The present study was aimed to analyse renal parameters – blood urea , serum and urinary creatinine , serum and urinary electrolytes in patients with history of hypothyroidism posted for elective surgery during the year 2015 at decan college of medical sciences .

**Material and method :** The following biochemical parameters derived were estimated glomerular filtration rate (eGFR) and fractional excretion of sodium (FeNa). 50 people were taken as controls that had no medical illness and 50 cases of incidental subclinical hypothyroidism were taken as case study group . Mean and Standard deviation was assessed and p value  $\leq 0.05$  was considered significant.

**Result:** The mean and SD of blood urea (mg/dl) in controls is  $28.2 \pm 4.75$  as compared to  $27.29 \pm 8.1$  in cases . The P value is not significant . The mean and SD of serum creatinine (mg/dl) in controls was  $0.7 \pm 0.15$  as compared to  $1.10 \pm 0.24$  in cases .The P value was  $< 0.01$  which is significant . The mean and SD of electrolytes sodium , potassium and chlorides in controls were  $140.4 \pm 2.15$  ,  $4.06 \pm 0.25$  and  $98.3 \pm 2.25$  as compared to case study group of  $147.93 \pm 6.98$  ,  $4.15 \pm 0.42$  and  $107.81 \pm 10.35$  The P value of serum sodium and chloride is  $< 0.01$  which is significant . The p value of potassium is not significant .

**Introduction :** Hypothyroidism can result from any of a variety of abnormalities that lead to insufficient synthesis of thyroid hormones. Overall primary hypothyroidism accounts for approximately 95% of cases and only 5% or less being supratheroid origin <sup>(1)</sup>. The most common primary hypothyroidism is often auto-immune in nature, leading to myxedema in adults. Women are more affected than males <sup>(2)</sup>.

Thyroid hormone deficiency affects every tissue in the body so the symptoms are multiple. Pathologically, the most characteristic finding is the accumulation of glycosaminoglycans – mostly hyaluronic acid in the interstitial tissues, which increases the capillary permeability to albumin that accounts for interstitial edema. This accumulation is not due to excessive synthesis but due to decreased degradation of glycosaminoglycans <sup>(3)</sup>.

Hypothyroidism is associated with gain in weight, partly due to water retention in tissues and partly to fat storage; which is evident in particular sites like head, neck, trunk but spares limbs<sup>(4)</sup>. Disease affecting the kidneys can often be detected even in asymptomatic patients from clues derived from routine clinical and laboratory examination. Hypothyroidism decreases cardiac output leading to decreased renal blood flow and decreased glomerular filtration rate and thereby decreases reabsorptive and secretory maxima of the kidneys. The ability to concentrate urine is slightly impaired and mild proteinuria may occur as a result of effect of hypothyroidism on kidneys<sup>(5)</sup> . Many case reports

document increased levels of serum creatinine with hypothyroidism in humans<sup>(6-11)</sup>. A person with impaired renal functions can live a normal life depending upon the severity of impairment of renal function. Hypothyroidism may also result in depression of myocardial function, decreased spontaneous ventilation, abnormal baroreceptor function, reduced plasma volume, anaemia, hypoglycaemia, hyponatraemia and impaired hepatic drug metabolism . The presence of hypometabolic state necessitates careful perioperative cardiovascular monitoring and judicious use of anaesthetic drugs.

### AIM OF THE STUDY

The aim of the present study was to analyze the renal functions in subclinical hypothyroidism to possibly detect abnormal biochemical renal parameters at the earliest and thereby help in prevention of anesthesia complications and permanent damage to renals. The biochemical parameters to be assessed in this study were: Estimated glomerular filtration rate (eGFR) , fractional excretion of sodium (FeNa), blood urea, serum and urinary creatinine and electrolytes(sodium, potassium and chloride) .

### MATERIALS AND METHODS

The study was carried out in Department of Anesthesia DCMS at pre anesthesia check up clinics . The diagnosed cases of subclinical hypothyroidism were selected from the department of surgery and obstetrics coming for elective surgery . 50 people were taken as controls that had no medical illness and 50 cases of incidental subclinical hypothyroid were taken as case study group .The bio-chemical parameters of the above were compared to know the sta-

tistical difference.

**Collection of Sample (Blood and Urine):**

- About 5 ml of blood is collected from cubital vein by vene puncture under aseptic conditions into a sterile bottle and allowed to clot. The serum is separated and used for estimation of the above parameters.
- A spot urine sample is collected from the hypothyroid patients and this urine is used for the estimation of the above urinary parameters.
- Semi-auto analyzer and Electrolyte analyzer were used to asses the above parameters.
- Levey et al in 1999 proposed a formula called Modification of diet in renal disease(MDRD) to asses the GFR which included plasma creatinine, age, sex and race.

**MDRD formula is as follows:-**

$$GFR (ml / min / 1.73 m^2) = \frac{186}{X [Plasma creatinine (mg/dL)] \times X [Age]^{-0.203} \times X [0.742 \text{ if patient is female}] \times X [1.210 \text{ if patient is black}]}$$

**Fractional excretion of sodium:-<sup>(1)</sup>**

Estimation of the fractional excretion of sodium (FeNa) relates sodium clearance to creatinine clearance which differentiates pre-renal failure from post-renal failure.

$$FeNa = \frac{\text{Urinary sodium} \times \text{Plasma creatinine}}{\text{Plasma sodium} \times \text{Urinary creatinine}} \times 100$$

**Estimation of Urea and Creatinine**

Blood Urea and Serum Creatinine were estimated using the standard kit by Berthelot, End Point Assay and Jaffe's method

Data are expressed as mean±standard deviation (SD). The significant difference between the test group and the control group were analyzed using Anova. Data were analyzed using the SPSS software. A value of p<0.05 was set as the level of significance.

**ESTIMATION OF ELECTROLYTE (SODIUM, POTASSIUM AND CHLORIDE) BY ELECTROLYTE ANALYZER: PRINCIPLE:**

The AVL 9180 analyzer methodology is based on the ion-selective electrode (ISE). They are three different electrodes used in AVL 9180 analyzer (Sodium, Potassium and Chloride). Each electrode has an ion-selective membrane that undergoes a specific reaction with the corresponding ions contained in the sample being analyzed.

The ion concentration in the sample is then determined by using a calibration curve determined by measured points of standard solutions with precisely known ion concentration.

**Statistical analysis:** Mean and standard deviation (S.D) of all variables were calculated and compared with those of controls. Statistical significance was assessed and P-value <0.05 were considered significant.

**EXCLUSION CRITERIA**

1. Diagnosed cases of hypothyroidism on treatment
2. Emergency cases
3. Known cases of electrolyte imbalances
4. Known cases of diabetes and hypertension

**Results**

Sl.No.	Investigation		Control Subjects	Test subjects
1	Blood urea(mg/dL)	Mean	28.2	27.29
		SD	4.75	8.1
		SEM	2.82	
		t-test	0.31	
		P	NS	
2	Serum creatinine(mg/dL)	Mean	0.75	1.10
		SD	0.15	0.24
		SEM	0.08	
		t-test	3.35	
		P	<0.01	
3	Serum sodium(mEq/L)	Mean	140.4	147.93
		SD	2.15	6.98
		SEM	1.99	
		t-test	3.4	
		P	< 0.01	
4	Serum potassium(mEq/L)	Mean	4.06	4.15
		SD	0.25	0.42
		SEM	0.04	
		t-test	1.15	
		P	NS	
5	Serum chloride(mEq/L)	Mean	98.3	107.81
		SD	2.25	10.35
		SEM	3.35	
		t-test	2.82	
		P	< 0.01	
6	Urine creatinine(mg/dL)	Mean	74	76.5
		SD	14.18	15.28
		SEM	5.95	
		t-test	0.75	
		P	NS	
				(Cont.d)
7	Urine sodium(mmol/L)	Mean	164.4	120.78
		SD	27.7	62.85
		SEM	21.17	
		t-test	2.03	
		P	<0.05	

8	Urine potassium(mmol/L)	Mean	22.9	21.86
		SD	5.78	5.53
		SEM	2.28	
		t-test	0.45	
		P	NS	
9	Urine chloride(mmol/L)	Mean	116.2	106.94
		SD	39.7	44.75
		SEM	17.31	
		t-test	0.70	
		P	NS	
10	Estimated glomerular rate(eGFR)1.73m <sup>2</sup> /ml/min	Mean	134.5	81.38
		SD	14.01	21.05
		SEM	7.26	
		t-test	7.18	
		P	<0.01	
		SD :-	Standard Deviation	
		SEM:-	Standard Error of Mean	

The results of table 1 shows the P-value of eGFR, Serum Creatinine, sodium and chloride are highly significant and p-value of Urinary sodium is significant and p-value of the remaining biochemical parameters were considered as statistically not significant.

**DISCUSSION:** Hypothyroidism characterized by decreased production of thyroid hormones ,affects the functions of various organs like muscle, kidney, reproductive system, central nervous system and hemopoetic system due to its effect on various metabolic pathways in the body.

The reports of few studies conducted in this direction indicates, there is decreased renal blood flow, glomerular filtration rate and tubular reabsorptive and secretory maxima is reduced, and urine flow is reduced. Delay in water excretion appears to be due to decreased volume delivering to the distal diluting segment of nephron as a result of decreased renal blood flow. The impaired renal excretion of water and the retention of water in the interstitial ground substance results in increased total body water and reduced plasma volume. Occasionally hyponatremia is also reported which is associated with severe hypothyroidism. The amount of exchangeable potassium is usually nor-

mal in relation to lean body mass. It is also reported no change in the levels of non – protein nitrogen substances like urea, creatinine and uric acid. In present study, in addition to estimation of blood urea and serum creatinine, two important renal function tests i.e eGFR and FeNa are done to find out the renal hemodynamics and sodium handling capacity of the kidney in hypothyroid state.

The use of fractional excretion of sodium has become popular test as a more accurate means to differentiate pre-renal failure from renal and post renal failure. The fractional excretion of sodium represents the fraction of filtered sodium that is ultimately excreted in the urine. If the value is less than 1%, it indicates pre-renal, and greater than 2% indicates 2% renal causes. The result of the two tests indicates that out of 50 cases of hypothyroid patients investigated, 32 cases have reduction in eGFR values than control group(64%) and in 50% cases fractional clearance of sodium being less. The mean serum sodium levels in hypothyroid patients are also high as compared to control group, where as K<sup>+</sup>, urea and creatinine values do not differ from control group. Similarly urinary potassium values are same as that of control value.

### CONCLUSION

From this results it can be concluded that hypothyroid state affects renal functions in significant percentage of affected subjects as indicated by decreased glomerular filtration rate, decreased urinary sodium excretion and decreased fractional clearance of sodium resulting in water retention in the body. As these changes in kidney functions are reversible, it is necessary to identify hypothyroid state in person at the earliest and institute treatment immediately.

### BIBLIOGRAPHY

1. Milliers Textbook of Anesthesia -8 edition Ronald D. Miller, MD, Lars I. Eriksson, Lee A. Fleisher, MD, Jeanine P. Wiener-Kronish, MD and William L. Young
2. Clinical Anesthesia 7 th edition by Paul G Barash , Bruce F Cullen Robert K Stoelting
3. *Anesthesia and hypothyroidism: a review of thyroxine physiology, pharmacology, and anesthetic implications.* *Anesth Analg* 1982 Apr;61(4):371-83.
4. *Harrison's principles of internal medicine; Vol I & II, 19<sup>th</sup> edition.*
5. *D.M. Vasudevan, Sreekumari's; text book of biochemistry 7<sup>th</sup> edition.*
6. *Basic and Clinical Endocrinology, 9<sup>th</sup> Edition, Francis G Greenspan, David G Gardener*
7. *Potential Mechanism of hypothyroidism – induced Hyponatremia ; internal medicine Vol 39, No 12 (December 2000).*
8. *Hypothyroidism – Padma S. Menon, Indian journal of clinical practice, Vol. 10, No.7.*
9. *The Renal Manifestations of Thyroid Disease Laura H. Mariani and Jeffrey S. Berns J Am Soc Nephrol 23: 22–26, 2012*
10. *Iglesias P, Díez JJ: Thyroid dysfunction and kidney disease. Eur J Endocrinol 160: 503–515, 2009J Am Soc Nephrol 23: 22–26, 2012Thyroid Hormone and the Kidney 25www.jasn.org SCIENCE IN RENAL MEDICINE*
11. *Mooraki A, Broumand B, Neekdoost F,Amirmokri P, Bastani B: Reversible acute renal failure associated with hypothyroidism: Report of four cases with a brief review of literature. Nephrology (Carlton) 8: 57–60,2003*
12. *Montenegro J, González O, Saracho R,Aguirre R, González O, Martínez I: Changes in renal function in primary hypothyroidism. Am J Kidney Dis 27: 195–198, 1996*
13. *Kreisman SH, Hennessey JV: Consistent reversible elevations of serum creatinine levels in severe hypothyroidism. Arch Intern Med159: 79–82, 1999*
14. *den Hollander JG, Wulkan RW, Mantel MJ,Berghout A: Correlation between severity of thyroid dysfunction and renal function. ClinEndocrinol (Oxf) 62: 423–427, 2005.*