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# Estimation of FT3, FT4 and TSH Levels Among Different age-sex Groups of Healthy Population in Rajshahi city

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Abstract: Background: Thyroid is an endocrine gland which is located below the larynx. The principal hormones of thyroid gland are thyroxine  $(T_4)$ , and tri-iodothyronine  $(T_3)$ . The activity of the thyroid gland depends on the age as well as gender. However, with advancing age the concentration of these hormones decreases in apparently healthy male and female. The current study was carried out to investigate the impact of age and gender on the level of free triiodothyronine (FT<sub>3</sub>), free thyroxine (FT<sub>4</sub>) and thyroid stimulating hormone (TSH) in apparently healthy individuals. Objective: This study was designed to observe the levels of FT<sub>3</sub>, FT<sub>4</sub> and TSH among different age-sex groups of normal healthy population residing in Rajshahi City. Methods: This cross-sectional descriptive study was carried out in the department of physiology in collaboration with institute of nuclear medicine and Allied Sciences, Rajshahi. 120 apparently healthy persons in age group of 8-65 years were studied. Random blood sample was taken to measure the level of free T<sub>3</sub>, free T<sub>4</sub>, TSH by Radioimmunoassay (RIA) and Immunoradiometric assay (IRMA). Data were analyzed by ANOVA Test and independent t test. Result: Nonsignificant drop of FT4, TSH level and significant drop of FT3 level with advancing age. Moreover, mean FT<sub>3</sub>, and FT<sub>4</sub> concentration is non-significantly higher in male in comparison to female. However mean TSH concentration is significantly higher in male in comparison to female. Conclusion: It can be concluded from the present study that non-significant drop of FT4, TSH level and significant drop of FT3 level with advancing age.

Keywords: Free Triiodothyronine (FT3), Free Thyroxine (FT4) and Thyroid Stimulating Hormones (TSH), Radioimmunoassay (RIA) and Immunoradiometric assay (IRMA).

### **Original Research Article**

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Article at a glance:

Study Purpose: To contribute to existing knowledge or propose new ideas.

Key findings: According to results obtained in this study, normal aging is characterized by gradual sex dependant decline in FT3 concentration without significant change in FT4 and TSH concentration. The mean FT3 and FT4 levels are non significantly higher in male than female and mean TSH levels are significantly higher in male than female.

Newer findings: Mean FT3 and FT4 concentration is non-significantly higher in male in comparison to female. However mean TSH concentration is significantly higher male in comparison to female but serum thyroid hormones and TSH levels were found to be higher in females than males in previous study.

Abbreviations: FT3 – Free Triiodothyronine, FT4 – Free Thyroxine, TSH – Thyroid Stimulating Hormone, RIA – Radioimmunoassay



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### **INTRODUCTION**

The Human thyroid gland secretes about 103 nmol of thyroxine T<sub>4</sub> and 7 nmol of triiodothyronine T3 per day. T3 is 3 to 5 times more active than T<sub>4</sub>, T<sub>3</sub> is also formed in the peripheral

tissue (liver, kidney, muscle) by deiodination of T<sub>4</sub>. Thyroid hormones mostly transport in the blood as bind form. Normally 99.98% of T<sub>4</sub> and 99.8% of T<sub>3</sub> in plasma are bound to plasma proteins and the rest are in free forms. 67% of T4 is bound to Thyroxine

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binding  $\alpha$ - globulin (TBG), 20% to Thyroxine binding pre-albumin (TBPA) and 13% to albumin. On the other hand, 46% of binding T<sub>3</sub> is loosely bound to TBG and 54% to albumin but bond not at all to TBPA. The free forms are the active form (FT<sub>3</sub> and FT<sub>4</sub>) and the free forms are in equilibrium with bindings form in plasma and tissue.<sup>1</sup>

The levels of serum free T<sub>3</sub>, free T<sub>4</sub> and TSH are remarkable different from infant to old age including both male and female.2 This variation of free T<sub>3</sub>, free T<sub>4</sub> and TSH levels in different decades of life also affected by sex, nutritional status, seasons, geographical region. Due to particular family setup in Bangladesh, female are more exposed to nutritional deficiencies that causes health problems including thyroid disorder. However, with increasing age, the concentration of these hormones dropped in both sexes. The drop was more in female than males. Higher frequencies of thyroid problems in females may be attribute to stress, multiple pregnancies and lactation.<sup>3</sup>Thyroid disease is common and its prevalence increases with age especially in the elderly and in often associated with greater morbidity than in younger person. The relation of the thyroid gland to the ageing process is of interest because of the importance of the organ in regulating the rates of various body functions. It was found that during a normal human life span, serum T3 remains high during adolescence, then it remains stable until late middle age and ultimately decrease with their increasing age.4 TSH values increased significantly in females over age 60. Throughout all decades, males had stable TSH levels that were slightly higher than the female levels before age 60 and lower there after.5 Estrogens cause increased secretion of thyroid binding globulin (TBG). On the other hand, TBG levels are depressed by androgens.6

Khan *et al.*<sup>7</sup> and Kaur *et al.*<sup>4</sup> showed that concentration of T<sub>3</sub> and T<sub>4</sub> was significantly higher in females than males, while TSH concentration was not significantly different in both sexes. But Meng *et al.*<sup>8</sup> found female with advancing age had high TSH and high FT<sub>3</sub> level than male. Mukhtar *et al.*<sup>9</sup> indicated that the level of T<sub>4</sub>, T<sub>3</sub> and TSH in Blood Serum of male and female were of insignificant difference. Rotha *et al.*<sup>10</sup> showed that level of T<sub>4</sub> was higher in female than male. They further concluded that T3 and TSH levels are not influenced by gender. But Ahmed et al.11 and Chaurasia et al.12 have observed serum T4 level slightly higher in male and TSH level higher in female. Alom et al.3 and Abbas et al.13 shows that the serum thyroid hormones and TSH levels were found to be higher in females than males. On comparing the different thyroid hormone levels in different age groups showed that serum T3 levels were higher in children and declined progressively with age while serum T<sub>4</sub> levels declined slightly only in elderly age group. On contrary mean TSH levels was found to be significantly changed with age. Suzuki et al.14. suggested that concentration of free thyroid hormone and resistance indices were decreased with aging in males. In female the free thyroid hormone concentration was not altered by aging but TSH level increased in an age dependent manner. So the present study has been designed to know the variation of normal level of free T<sub>3</sub>, free T<sub>4</sub> and TSH regarding age and sex of people in Rajshahi City. So, that normal level is easily detectable in respect of age to differentiated from the abnormal one.

# **METHODS**

This cross-sectional descriptive study was carried out in the department of physiology in collaboration with institute of nuclear medicine and Allied Sciences between the period of January 2016 to December 2016. Apparently 120 healthy persons aged 8-65 years were selected residing in Rajshahi City. Purposive sampling technique was applied to select each subject. After taking informed consent, complete history taking and physical examination were done and recorded in a preformed data sheet. After breakfast, 5ml of venous blood sample were drawn into the test tube (from the anticubital space of the forearm) by venipuncture after taking all aseptic precautions. After coagulation serum was separated by centrifugation at 3500 rpm for 2 minutes. Then serum were used for estimation of FT<sub>3</sub>, FT<sub>4</sub> and TSH level by Radioimmunoassay (RIA) and Immunoradiometric assay (IRMA). The results of TSH were expressed in µIU/ml and FT3 and FT4 were expressed in fmol/ml. Collected data were analyzed by using SPSS (statistical package social sciences) computer software for programmed (version-20) and the tests of significance were calculated by using ANOVA test and independent t test. P value at or below 0.05 was **RESULTS** taken as level of significance.

Table 1: Serum FT <sub>3</sub> level of male in different age groups (n=60)			
Age group in years including only male respondents	FT <sub>3</sub>	P-value	
	fmol/ml		
	(mean±SD)		
Group 1 (8-17 yrs)	8.36±3.10		
	(8.36-11.46)		
Group 2 (18-40 yrs)	8.23±1.90	.198 ns	
	(8.23-10.13)	.19813	
Group 3 (41-65 yrs)	7.16-±1.10		
	(7.16-8.26)		

## ns=Not significant (p=>0.05).

The test of significance was calculated using Anova test.

Table 1 shows FT<sub>3</sub> level of male respondents in different age group. The mean FT<sub>3</sub> level is almost same in adolescent group and in man

upto 40 years of age, after that slight decrease of FT3 level is seen.

Table 2: Serum FT₄ level of male in different age groups (n=60)			
Age group in years including only male	FT <sub>4</sub>	P-value	
respondents	fmol/ml		
	(mean±SD)		
Group 1 (8-17 yrs)	24.22±5.56		
	(24.22-29.78)		
Group 2 (18-40 yrs)	22.15±13.73	.695 ns	
	(22.15-35.88)	.090115	
Group 3 (41-65 yrs)	21.79±6.37		
	(21.79-28.16)		

ns=Not significant (p=>0.05).

The test of significance was calculated using Anova test.

Table 2 shows FT<sub>4</sub> level of male respondents in different age group. The mean FT<sub>4</sub> level is seen highest in adolescent group. A sharp fall is seen after that in the age group upto 40 years.

Further little decrease in the mean level of FT<sub>4</sub> is seen after 40 years.

Table 3: Serum TSH level of male in different age group	os (n=60)
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Age group in years including only male	TSH	P-value
respondents	µIU/ml	
	(mean±SD)	
Group 1 (8-17 yrs)	3.92±4.60	
	(3.92-8.52)	
Group 2 (18-40 yrs)	4.57±10.37	<b>711</b> ng
	(4.57-14.94)	.711 <sup>ns</sup>
Group 3 (41-65 yrs)	$2.68 \pm 4.32$	
	(2.68-7.0)	

ns=Not significant (p=>0.05)

The test of significance was calculated using Anova test.

Table 3 shows TSH level of male respondents in different age group. The TSH level increases from adolescent group to middle age group. But after 40 years there is a decrease in TSH level which comes down to lower than the adolescent level.

Age group in years including only female	FT3	P-value
respondents	fmol/ml	
	(mean±SD)	
Group 1 (8-17 yrs)	7.66±1.09	
	(7.66-8.75)	
Group 2 (18-40 yrs)	8.17±1.62	0.388 <sup>ns</sup>
	(8.17-9.79)	0.300
Group 3 (41-65 yrs)	7.65±1.32	
	(7.65-8.97)	

Table 4. Serum	ET <sub>2</sub> lovel of female	e in different age gro	n = (n = 60)
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ns=Not significant (p=>0.05).

The test of significance was calculated using Anova test.

Table 4 shows FT<sub>3</sub> levels of female respondents in different age group. The mean FT<sub>3</sub> level is almost same in adolescent group and in

female upto 40 years of age. After that slight decrease of FT<sub>3</sub> level is seen.

Age group in years including only female	FT <sub>4</sub>	P-value
respondents	fmol/ml	
	(mean±SD)	
Group 1 (8-17 yrs)	19.99±5.54	
	(19.99-25.53)	
Group 2 (18-40 yrs)	20.69±11.37	0.626 <sup>ns</sup>
	(20.69-32.06)	0.02013
Group 3 (41-65 yrs)	22.34±5.67	
	(22.34-28.01)	

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ns=Not significant (p=>0.05).

The test of significance was calculated using Anova test.

Table 5 shows FT<sub>4</sub> levels of female respondents in different age group. Mean FT4 level is seen highest after 40 years of age. Gradually increase is seen after adolescent age group upto 65 years.

Age group in years including only female	TSH	P-value
respondents	µIU/ml	
	(mean±SD)	
Group 1 (8-17 yrs)	1.33±0.66	
	(1.33-1.99)	
Group 2 (18-40 yrs)	1.63±0.99	0. 216 <sup>ns</sup>
	(1.63-2.62)	0. 210
Group 3 (41-65 yrs)	2.51±3.59	
	(2.51-6.1)	

ns=Not significant (p=>0.05).

The test of significance was calculated using Anova test.

Table 6 shows TSH levels of female respondents in different age group. The level of TSH increases from adolescent group to older age group. But after 40 years there is a increase in TSH level which comes higher than the adolescent group.

Table 7: Gender distribution of hormone level			
Hormone level	Male (n=60)	Female (n=60)	P-value
FT₃ (fmol/ml)	7.96±2.24	7.83±1.38	0.708 <sup>ns</sup>
FT4 (fmol/ml)	22.74±9.48	21.04±7.93	0.282 <sup>ns</sup>
TSH (µIU/ml)	3.79±7.15	1.81±2.21	$0.044^{s}$

ns=Not significant (p=>0.05).

s=Significant (p=<0.05).

The test of significance was calculated using independent t test.

Table 7 shows thyroid hormone levels among total male and total female respondents, mean value of all the hormone show higher value in male group but the difference is not significant in case of FT<sub>3</sub> and FT<sub>4</sub>. Statistically significant difference is seen in case of TSH, which showed much higher value in male group.

### **DISCUSSION**

The levels of FT<sub>3</sub>, FT<sub>4</sub> and TSH have significant effects among gender with age. Gender has a vital role in the level of thyroid hormones. However with increasing age, the concentration of these hormones dropped in both sexes. The drop is more in female than men. Women have more economic and family responsibility as compared to men. Present study found that, non-significant drop of FT4, TSH level and significant drop of FT3 level with advancing age. This findings are similar with the study of Suzuki et al. (2012)<sup>14</sup>, Abbas et al. (2014)<sup>13</sup>, Alom et al. (2016)<sup>3</sup> It occurs due to reduced secretion and concentration of FT3 levels and increaseturn over rate of FT<sub>3</sub>.<sup>14</sup> In this study, it is found that higher FT4, FT3 and high TSH level in male in comparison to female. This finding is consistent with Ahmed et al. (2009)11, Chaurasia et al. (2011)<sup>12</sup> and Dambal et al. (2013).<sup>15</sup> Lower thyroid hormone concentration in female may represent the fact that hypo-thalamo-pituitary-thyroid axis in female is set at lower regulatory level. Furthermore, the finding may be due to influence of female sex hormone estrogen on thyroid binding globulin (TBG) level. Estrogen stimulates the synthesis of TBG which increases total T4 and T3 but decreases FT<sub>4</sub> and FT<sub>3</sub> concentration.

In contrast, Rotha *et al.*  $(2015)^{10}$  state that higher T<sub>4</sub> in female than male. It may be due to the

fact that they measured total T<sub>4</sub> concentration instead of FT4 concentration. Moreover, Dika et al. (2010) found no influence of sex on thyroid hormone concentration.<sup>16</sup> It may be due to the fact that they selected very narrow age range (19-37) years. So influence of sex hormones were maximum in both gender to find out any significant difference of thyroid hormone concentration. Furthermore, Kaur et al. (2007)<sup>4</sup>, Khan et al. (2010)<sup>7</sup> and Alom et al. (2016)<sup>3</sup> showed higher concentration of T<sub>4</sub>, T<sub>3</sub> in female than male. Ahmed et al. (2009)11 state that T4, T<sub>3</sub> concentration decreases in female after 60 years of age. It represents that menopause causes abrupt reduction of estrogen level as well as TBG and total T<sub>4</sub>, T<sub>3</sub> concentration. This study showed higher FT<sub>4</sub> and TSH concentration in (8-17) years age group in comparison to all other groups. This finding is in agree with Kaur et al. (2007)<sup>4</sup>, Alom et al. (2016)<sup>3</sup> and Dambal et al. (2013).<sup>15</sup> This finding represents that marked changes occur in thyroid function during puberty as an adaptation to body and sexual development. Adaptation of hypothalamopituitary-thyroid axis during puberty in response to increased energy expenditure may be the reason. Moreover, we observed FT<sub>4</sub>, FT<sub>3</sub> and TSH concentration is higher in below forty years male in comparison to corresponding female group. It exhibits that hypothalamo-pituitary-thyroid axis is set at lower regulatory level in female Wllke et al. (1983).17 However, after forty years age, TSH concentration is non-significantly higher and FT4, FT3 concentration is non-significantly lower in male in comparison to female. It represents, that aging process affects the hypothalamo-pituitary-thyroid axis more in male in comparison to female.

### Declarations

I, hereby, declare that the submitted Research Paper is my original work and no part of it has been published anywhere else in the past.

## **Ethical approval**

Ethical clearance for the study was taken from the Instutional Review Board and concerned authority, Rajshahi Medical College & Hospital.

### **Conflict of interest**

None declared.

### Consent

Informed written consent was taken from each patient or patient's attendant.

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