ACKNOWLEDGEMENTS

I would like to express my sincere thanks and gratitude to my supervisor Dr. Badr Eldine Hassan Elabid for his close supervision, encouragement and sustained support.

I wish to emphasize my heartily thanks to Dr. Mohamed Abdel Rahim, Head department of Biochemistry, Faculty of Science, Sudan University of Science and technology, for his advice, and follow up as a co-ordinator of the MSc program in clinical biochemistry.

My thanks and appreciation to (AlFatih, Najwa) the Technical Staff of the Radiation and Isotopes Centre (RICK) Khartoum, for collaboration and assistance in this work.

Thanks are extended to all patients and controls who donate me the samples.

Abstract

It is evident that, in the last years there is an increase incidence of thyroid disorders in Sudanese people. This study was conducted essentially to know the effects of age, sex and residence on the incidence of thyroid disorders, and to know the influence of iodine status of a subject on thyroid function by measuring urinary iodine concentration.

A total of 100 Sudanese patients, with thyroid disorders visiting the Radiation and Isotope Centre (RICK) at Khartoum, during the period of February 2005 to July 2005, were selected randomly to contribute in this study. A total of 30 healthy subjects from the copatients were volunteered to participate in this study as a control group. Specimens of sera and urine samples were collected from all patients and controls to estimate thyroid hormones, T₃ & T₄ by (RIA) method and TSH by (IRMA) method.

Urine samples were used to measure urinary iodine concentration by Sandle Koltholt Reaction (using ammonium persulfate as a catalyst).

The patients were categorized as hyperthyroidism and hypothyroidism. There is an increased incidence of hyperthyroidism in the middle aged patients, while hypothyroidism is more common in the elderly, the results were found to be as follows:

- In the hyperthyroidism (n=80): there were 16 patients (20%) of age < 20 years, 43 patients (53.3%) aged between 20-40 years, and 21 patients (26.2%) of age > 40 years.

- In hypothyroidism (n=20): there were 2 patients (10%) of age < 20 years, 6 patients (30%) aged between 20-40 years and 12 patients (60%) of age > 40 years.

Females were more susceptible to thyroid disorders than males for both hyperthyroid and hypothyroidism:

- In hyperthyroidism females were 65 out of 80 (81,3%).
- In hypothyroidism females were 14 out of 20 (70%).

The disease was found to be distributed in all regions of Sudan, with increased incidence of hyperthyroidism in Central-Sudan (Khartoum State & Gazera area) (47.5%), North (18.8%), East (5%), West (13.7%) and South (5%) while for hypothyroidism there was increased incidence in the West (45%), Central-Sudan (26%), North (25%), East (5%), South (0%).

In this study, urinary iodine concentration was found to be highly significantly raised in patients with hyperthyroidism compared to the control (P<0.01), while highly significantly reduced in patients with hypothyroidism (P<0.01).

From this study it is recommended that urinary iodine could be used as a screening and diagnostic test for both hyper and hypothyroidism.

ملخص البحث

تلاحظ ان هنالك ازديالاً مضطرلاً في اختلال نشاط الغدة الدرقية لدى السودانيين من حيث زيادة او انخفاض في نشاط الغدة. و قد اجريت هذه الدراسة اساسا لمعرفة مدى تأثير العمر ، الجنس ، مكان السكن الاصلى للفرد على نشاط الغدة الدرقية، مع قياس نسبة اليود لدى الفرد و معرفة مدى تأثيره على وظائف الغدة الدرقية و ذلك بقياس نسبة تركيز اليود في البول.

تم اختيار مجموعة 100 مريض مصاب باختلال نشاط الغدة الدرقية، منهم 80 مريضا مصاب بفرط نشاط الغدة، اضافة مريضا مصاب بفرط نشاط الغدة، اضافة الدرقية و 20 مريضا مصاب بغانون من اى مرض كعينة مرجعية للمقارنة.

و قد شملت الفحوصات كل المجموعات (130) شخص حيث تم قياس الثايرونين ثلاثى اليود و رباعى اليود (T₃, T₄) عن طريقة (RIA) و قياس محفز الغدة الدرقية ثلاثى اليود و رباعى اليود (IRMA) اما نسبة تركيز اليود فى البول وقيست بطريقة تفاعل ساندل كوتولت Sandle (Kottlot Sandle و قد خلصت الدراسة بأن اعلى نسبة من المرضى المصابين بفرط نشاط الغدة الدرقية تتراوح اعمارهم ما بين 20 و 40 سنة. (43 مريضا من بين 80 مريض) بنسبة 53.8% . اما اعلى نسبة من المرضى المصابين بانخفاض نشاط الغدة الدرقية اعمارهم اكثر من 40 سنة (12 مريضاً من بين 20 مريضاً) بنسبة 60% . ان النساء اكثر عرضة للاصابة باختلال الغدة الدرقية حيث هنالك 65 امرأة من بين 80 مصابا بانخفاض نشاط القدة بازدياد نشاط الغدة (بنسبة 81.8%) و 14 امرأة من بين 20 مصابا بانخفاض نشاط الغدة (بنسبة 70%). كذلك تلاحظ بان اعلى نسبة من المرضى المصابين بازدياد نشاط الغدة الدرقية يقطنون وسط السودان (ولاية الخرطوم + ولاية الجزيرة) بنسبة 45% و ان اعلى نسبة من المرصابين بانخفاض الغدة الدرقية يقطنون غرب السودان بنسبة 45%.

تلاحظ من هذه الدراسة بان هنالك ارتباط قها بين مستوى تركيز اليود في البول و اختلال نشاط الغدة الدرقية زيادة كان ام نقصالاً باحتمال احصائي (0.01) ، و هذا يمكننا من ان نستخدم قياس مستوى تركيزليود في البول لمعرفة نشاط الغدة الدرقية از ديادا او نقصانا أ.

List of contents

		Page
	Contents	i
	Acknowledgements	V
	Abstract	vi
	Abstract in Arabic	viii
	List of tables	ix
	List of figures	X
	List of Abbreviations	xi
	Chapter One	
1.1	Introduction	1
1.2	Objectives	2
	Chapter Two	
2	Literature Review	3
2.1	The endocrine glands	3
2.1.1	The thyroid gland	7
2.1.1.1	Embryonic development	7
2.1.1.2	Internal structure of thyroid gland	10
2.1.1.3	Hormones of the thyroid gland	10
2.1.1.3.1	Formation and secretion of thyroid	
	hormones	10
2.1.1.3.2	Synthesis and release of thyroid hormones	11
2.1.1.3.3	Transport and degradation of T ₃ & T ₄	12
2.1.1.3.4	Actions of thyroid hormones	13

2.1.1.3.4.1	Effects on general metabolism	13
		Page
2.1.1.3.4.2	Effect on growth and development	14
2.1.1.3.4.3	Effects on nervous system	14
2.1.1.3.4.4	Effects on carbohydrates metabolism	14
2.1.1.3.4.5	Effects on lipid metabolism	14
2.1.1.3.4.6	Effects on vitamin requirement	15
2.1.1.3.4.7	Effects on water and minerals metabolism	15
2.1.1.3.4.8	Effects on the cardiovascular system	15
2.1.1.3.4.9	Effects on the elementary tracts	15
2.1.1.3.4.10	Effects on gonads	15
2.1.1.3.4.11	Effects on mammary glands	15
2.1.1.3.5	Control of thyroid hormones	16
2.1.1.4	Disorders of the thyroid gland	18
2.1.1.4.1	Goiter	18
2.1.1.4.1.1.	Cause of goiter	18
2.1.1.4.2	Hyperthyroidism (thyrotoxicosis)	18
2.1.1.4.2.1	Clinical symptoms of hyperthyroidism	19
2.1.1.4.3	Hypothyroidism	21
2.1.1.4.3.1	Clinical symptoms of hypothyroidism	21
2.1.1.5	Thyroid function tests	23
2.1.1.5.1	Measurement of basal metabolic rate	
	(BMR)	23
2.1.1.5.2	Radioactive iodine uptake and thyroid	
	scanning	23

		Page
2.1.1.5.3	Estimation of T_3 , T_3 and TSH	24
2.1.1.5.4	Measurement of protein bound (PBI) and	
	thyroxine binding globulin (TBG)	24
2.2.	Iodine	24
2.2.1	The role of iodine in thyroid function	24
2.2.2	The amount of iodine needed daily for	
	normal thyroid function	25
2.2.3	Measurement of iodine intake	26
2.2.3.1	Urinary iodine measurement	28
2.2.4	Iodine induced problems	28
2.2.4.1	Disorders of excessive iodine intake	28
2.2.4.1.1.	Iodine deficiency disorders (IDD)	29
	Chapter three	
	Materials and methods	
3.1	Materials	31
3.1.1.	The study group	31
3.1.2	Samples collection	31
3.1.2.1	Blood samples	31
3.1.2.2	Urine sample	32
3.2	Methods	32
3.2.1	Estimation of total thyroxine (T ₄)	32
3.2.2	Estimation of total triiodothyronine	35
3.2.3	Estimation of TSH	37

		Page
3.2.5	Estimation of Creatinine	41
3.2.6	Urinary iodine measurement	42
3.3	Statistical analysis	45
	Chapter Four	
	Results	46
	Chapter Five	
	Discussion	60
	Chapter Six	
	Conclusion and Recommendation	63
	References	65
	List of Tables	
Table. No.		Page
2.1	Scale used to relate iodine nutrition to	
	urinary iodine concentrations	27
4.1	Frequency of thyroid disorders (hypo -	
	thyroidism & hyperthyroidism) according	
	to age	49
4.2	Frequency of thyroid disorders (hypo &	
	hyperthyroidism) according to gender	50
4.3	Distribution of patients with thyroid	

Estimation of blood urea.....

39

3.2.4

	disorders (hypo & hyperthyroidism)	
	according to original area	51
4.4	The means \pm SD of T3, T4, TSH &	
	urinary iodine concentration in	
	hyperthyroidism compared with control	
	group	52
4.5	The means \pm SD of T3, T4, TSH &	
	urinary iodine concentration in	
	hypothyroidism compared with control	
	group	53

List of Figures

Fig.	Figure Title	Page
2.1	The pituitary gland and its connection with	
	hypothalamus	4
2.2	The hypothalamus and the pituitary gland showing major	
	tracts and the nuclei closely related to the pituitary	
	glands	5
2.3	Feedback control of the endocrine gland	

	activity	6
2.4	Ventral view of the thyroid gland	8
2.5	Photomicrograph of the thyroid gland	9
2.6	Feedback control between the thyroid gland, the anterior	
	pituitary and hypothalamus	17
2.7	Patients with hyperthyroidism and goiter	20
2.8	Patients with hypothyroidism	22
4.1	Frequency of hyperthyroidism according to age	54
4.2	Frequency of hypothyroidism according to age	55
4.3	Frequency of hyperthyroidism according to gender	56
4.4	Frequency of hypothyroidism according to gender	57
4.5	Distribution of hyperthyroidism according to original	
	area	58
4.6	Distribution of hypothyroidism according to original	
	area	59

ABBREVIATIONS

ACTH Adrenocorticotrophin hormone

ANS Aniline-1- naphthalene sulfonic acid

BMR Basal metabolic rate

Ca⁺⁺ Calcium

DIT Dilodotyrosine

FSH Follicle stimulating hormone

GFR Glomerular filteration rate

I₂ Iodine

ICCID International Council for the Control of Iodine Deficiency

IDD Iodine deficiency disorders

IH Inhibitory hormone

IP Iodine pump

IR Endoplasmic reticulum

IRMA Immunoradiometric assay

LDL Low-density lipoprotein

LH Leutinizing hormone

Mc Monoclonal

MIT Monoiodotyrosine

NSB Non-specific binding

PBI Protein bound iodine

Pc Polyclonal

RH Releasing hormone

RIA Radioimmunoassay

rT3 Reverse T₃

T₃ Tri-iodothyronin

T₄ Thyroxin (tetraiodothyronin)

TBA Thyroxin-binding albumin

TBG Thyroxin-binding globulin

TBPA Thyroxin-binding prealbumin

TRH Thyrotrophin releasing hormone

TSH Thyroid stimulating hormone (thyrotrophin)

U.S.A United State of America

UNICIEF United Nation International Children Education Fund

WHO World Health Organization