#### **Thyroid and Parathyroid Imaging**



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#### **Nuclear Medicine Procedure**



Gamma Camera

- Patient injected with small amount of radioactive material.
- Radiopharmaceutical localizes in patient according to metabolic properties of that drug.
- Radioactivity decays, emitting gamma rays.
- *Gamma rays that exit the patient are imaged.*

# What are the nuclear medicine imaging methods?

#### **Conventional tumor imaging :**

- Planar : 2D
- > SPECT : 3D
- SPECT-CT : 3D (Function and anatomy)





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#### **PLANAR / SPECT**

**SPECT CT** 

# Onco PET : ▶ PET : 3D ▶ PET -CT : 3D (Function and anatomy)



#### **Physical Properties SPECT Radionuclides**

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Radionuclide	T/2	Type of	E(kev)
	physical	radiation	
Technitium 99m(Tc-99m)	6 hrs	Gamma	140
Iodine I131	8 days	Gamma/	364/606
		Beta	
Iodine I123	13.2 hrs	Gamma	159
Gallium Citrate (Ga-67)	78.3 hrs	Gamma	90,190,290
Thallium Chloride 201	73.1 hrs	X-ray	68-83
(Tl201)			
Indium 111 (In 111)	2.8 days	Gamma	173,247
Xenon 133	5.2 days	Gamma	81
Kripton 81m	13 secs.	Gamma	190

## Physical Properties of positron emitting (PET) Radionuclides



Radionuclide	T/2 phy-	positron	Productivity
	sical (min)	energy	
Carbon 11	20	0.96	accelerator
Nitrogen-13	10	1.19	accelerator
Oxygn-15	2	1.73	accelerator
Fluorine 18	110	0.635	accelerator
Gallium 68	68	1.9	generator
			(germanium 68)
Rubidium 82	1.3	3.15	generator
			(strontium-82)

## **Thyroid Scan : Procedure**



**Tc-99m Pertechnetate** 

<u>I-123</u>

Dose	0.5-4.0 mCi given IV	0.5 mCi orally	
Half Life	6 Hours	13 Hours	
Cost	Not Expensive (Generator)	Expensive (Cyclotrone)	
Time of imaging	20 min post injection	6 and 24 hours post ingestion	
Remarks	Trapped not organified	Trapped and organified	

## **Normal Values Of Thyroid Uptake**



I131 OR I-123 RAIU (4 & 24 hours) :

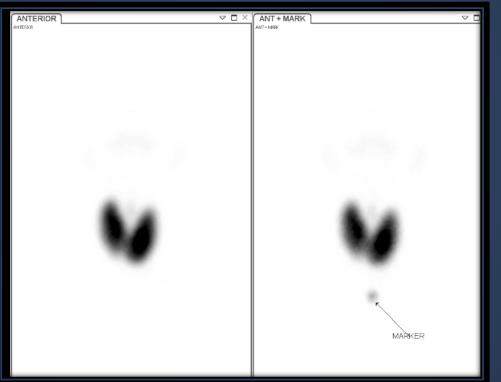
Normal 4 hour RAIU : 5 - 15%

Normal 24 hour RAIU : 8 - 35%

• Tc- 99m Uptake (20 min Uptake) : N (0.5 -4 .0%)

## **Causes of High Thyroid Uptake**





- Hyperthyroidism : Grave's
   Disease or TSH-secreting
   pituitary adenoma
- Autonomous toxic nodule
- Multinodular toxic goiter (Plumer's Disease)
- Enzyme defects : Dyshormonogenesis.
- **Iodine starvation (Iodine deficiency)**
- Lithium Therapy
- Recovery phase of thyroiditis.
- Rebound following abrupt withdrawal of antithyroid meds

#### **Causes of Low Thyroid Uptake**



#### Parenchymal Destruction:

- Acute, Subacute and Chronic Lymphocytic Thyroiditis

#### Hypothyroidism:

- Primary or Secondary (insufficient pituitary TSH secretion)
- Surgical/Radioiodine Ablation of Thyroid

#### Blocked Trapping:

- Iodine load (most common): Iodinated contrast material, Food rich in iodide: fish , cabbage ,...etc
   Exogenous thyroid hormone replacement depressing TSH levels (thyrotoxicosis factitia)
  - Ectopic thyroid: Struma Ovarii

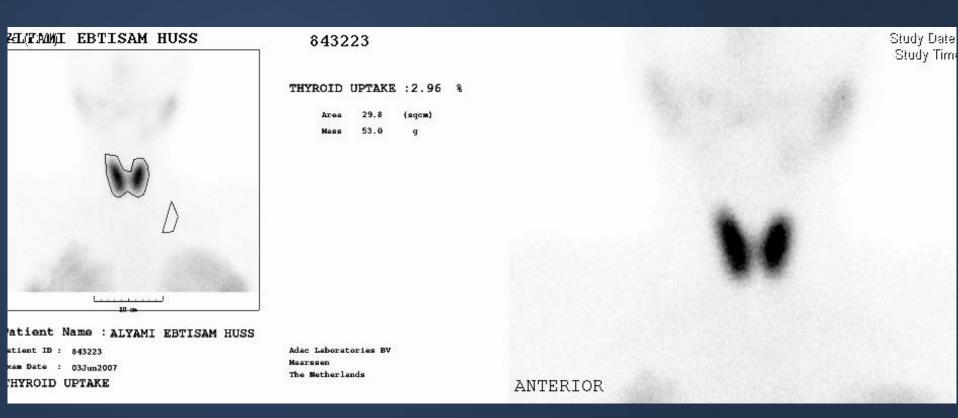
#### Blocked Organification:

- Antithyroid medication (PTU): Note- Tc-99m uptake should not be affected

## Tc-99m Thyroid scan and uptake Imaging plus uptake studies

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#### **THYROID METASTASES STUDY** (I-123 or I-131 as Sodium Iodide)



#### Indications

• Detection and localization of persistent or recurrent functioning thyroid cancer

#### **Patient Preparation**

- Stimulation of potentially functioning thyroid tissue: A. Inject recombinant human thyrotropin on 2 consecutive days and administer the radiopharmaceutical on the third day.
- B. Withdraw thyroid replacement hormones : 1. Thyroxine (T-4) for at least 4 weeks.
- 2. Triiodothyronine (T-3) for at least 10 days.

• The patient must not have had i.v iodinated contrast material (IVP, CT with contrast, myelogram, angiogram) for at least 3 weeks.

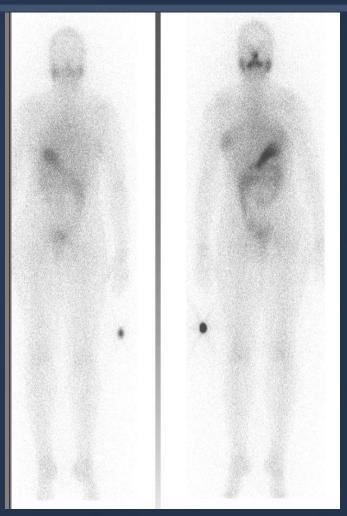
• The patient should be NPO for at least 4 hours prior to radiopharmaceutical administration and for at least 1 hour afterwards

#### Radiopharmaceutical, Dose, & Technique of Administration

• Radiopharmaceutical: Oral administration

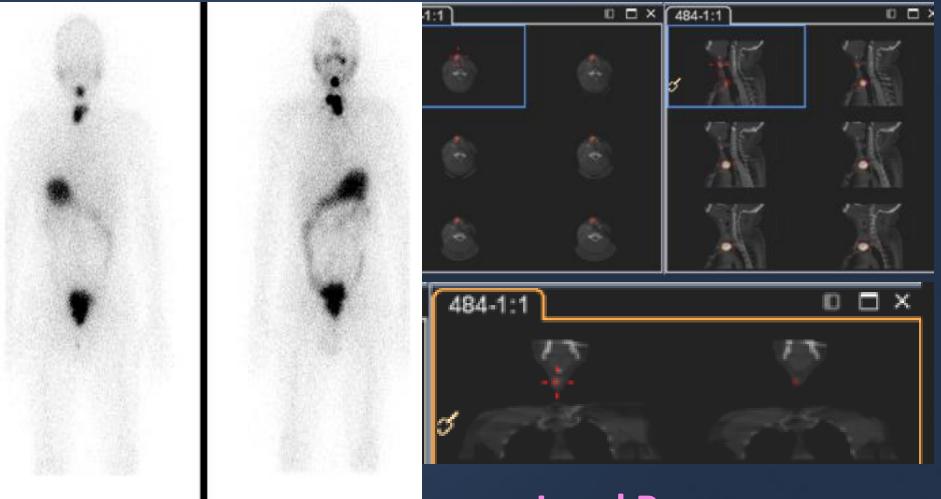
- a. I-123 as sodium iodide : 2 mCi
- b. I-131 as sodium iodide : 2-10 mCi

Imaging using Gamma camera : Whole body scan



#### Negative WBS

#### I-123 or I-131 Whole Body Scan(WBS) Planar Vs SPECT CT

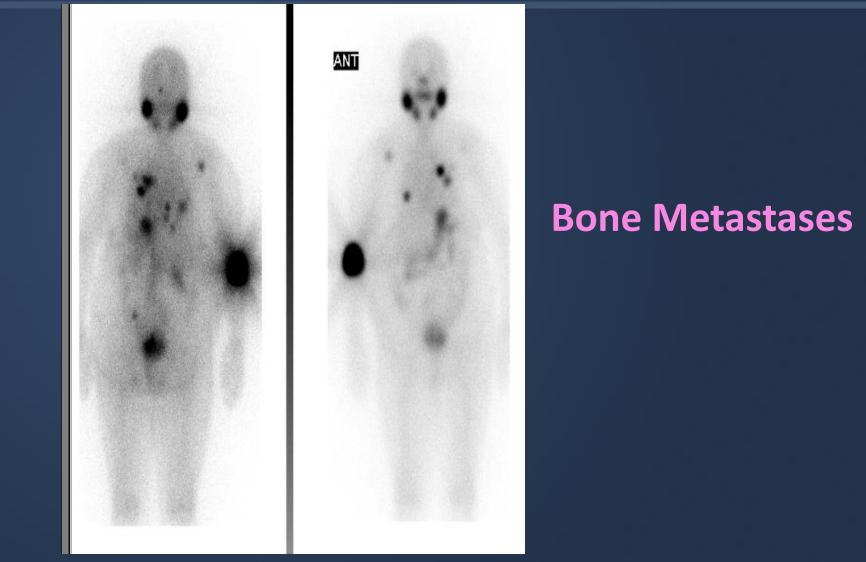


**Local Recurrence** 

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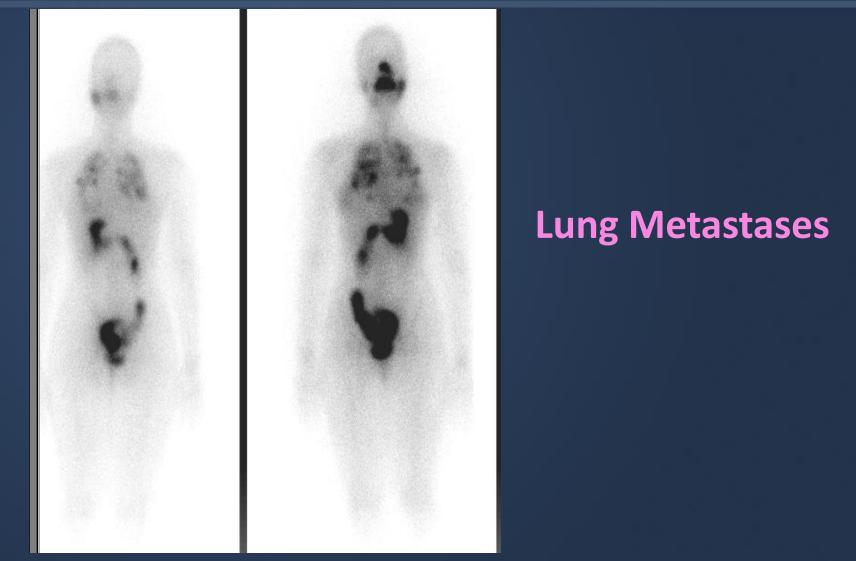
#### I-123 or I-131 Whole Body Scan(WBS)





#### I-123 or I-131 Whole Body Scan(WBS)





## When is thyroid scanning helpful? Indications for Thyroid Scan

- Evaluation of thyroid nodules : No. & type
- Evaluation of congenital hypothyroidism : Agenesis Vs. Dyshormonogenesis.
- Evaluation of neck masses : ectopic thyroid, thyroglobal cyst.
- Evaluation of thyrotoxicosis.

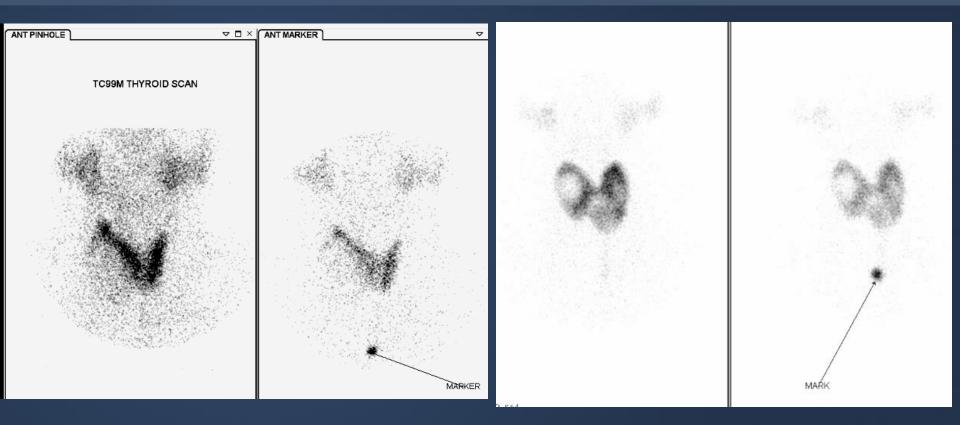
## **Evaluation of thyroid nodules** Single vs MNG

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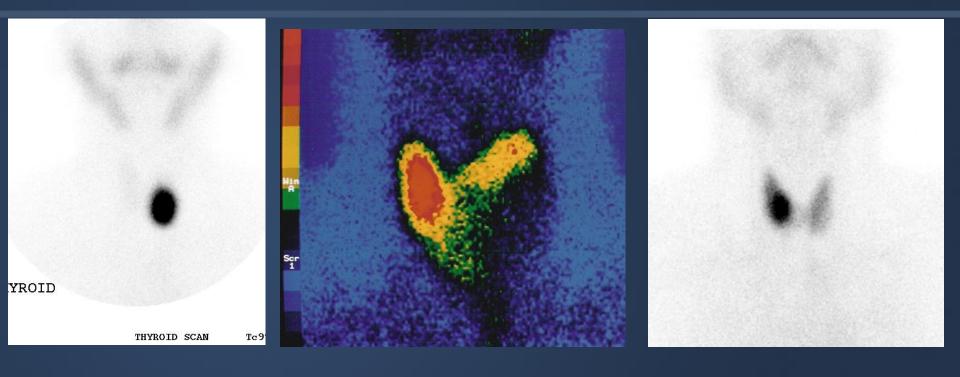
#### Solitary cold nodule Multinodular goiter The chance of malignancy is more in Solitary cold nodule than in MNG

## **Evaluation of thyroid nodules** Single vs MNG



Solitary cold nodule Multinodular goiter The chance of malignancy is more in Solitary cold nodule than in MNG

## **Evaluation of thyroid nodules** Hot vs Cold vs warm



Hot < 5% Malignant

15-20% Malignant

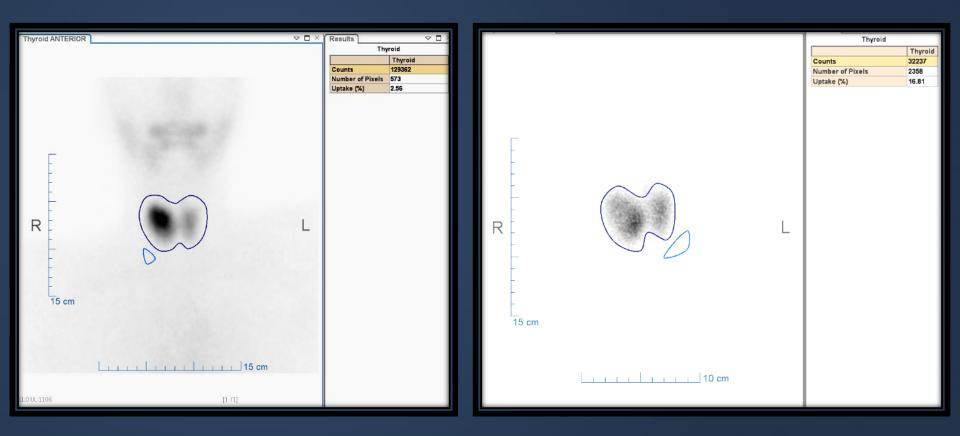
Cold

warm Suspicious

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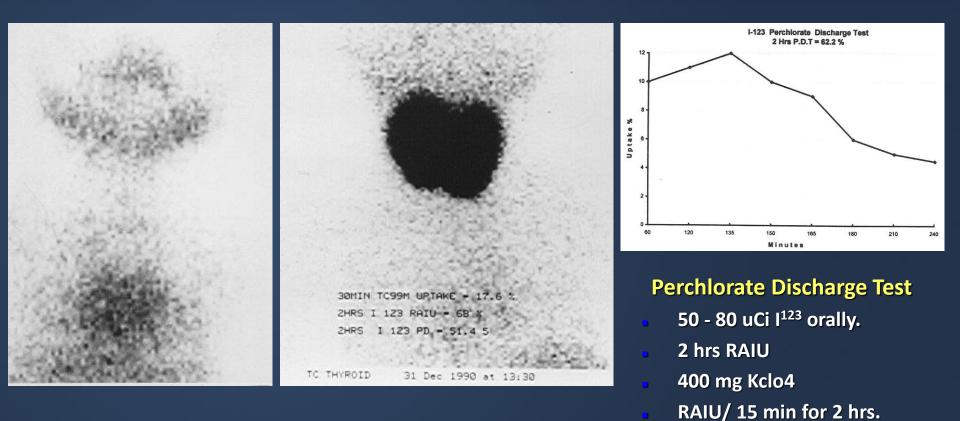


#### Discordance Tc –I123 Scan



#### The chance of malignancy of a discordant nodule about 20%

#### **Evaluation of congenital hypothyroidism** Agenesis vs Dyshormonogenesis



#### Agenesis

#### **Dyshormonogenesis**

Positive test : >= 15 fall of

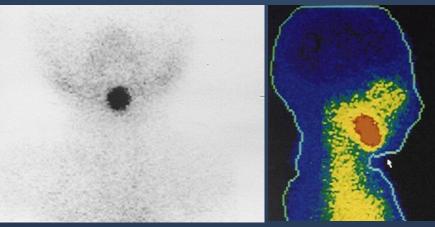
RAIU below 2 hrs. uptake.

## **Evaluation of neck masses** ectopic thyroid vs thyroglosal cyst

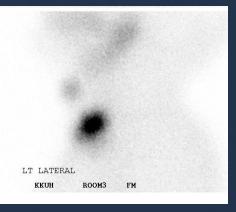












#### Thyroglosal cyst

#### Lingual thyroid

## **Evaluation of Thyrotoxicosis**

• Thyrotoxicosis IS NOT synonymous to Hyperthyroidism

 Thyrotoxicosis: Is a complex of signs and symptoms due to elevated thyroid hormones in the blood

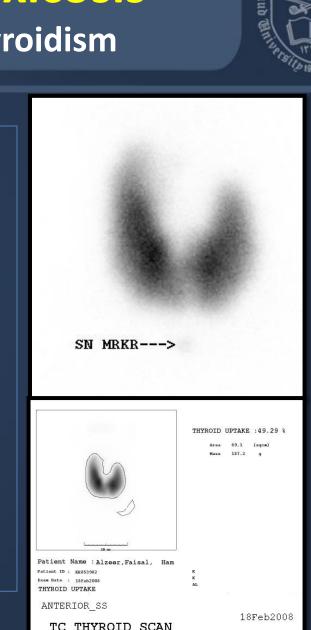
 Hyperthyroidism : Overproduction of thyroid hormones by the thyroid gland (hyperactive gland)

## **Evaluation of thyrotoxicosis** Thyrotoxicosis with hyperthyroidism

- Graves' Disease .
- Neonatal hyperthyroidism.
- Toxic nodular goiter : MNTG or Plummers disease ATN or toxic adenoma
- Iodine induced

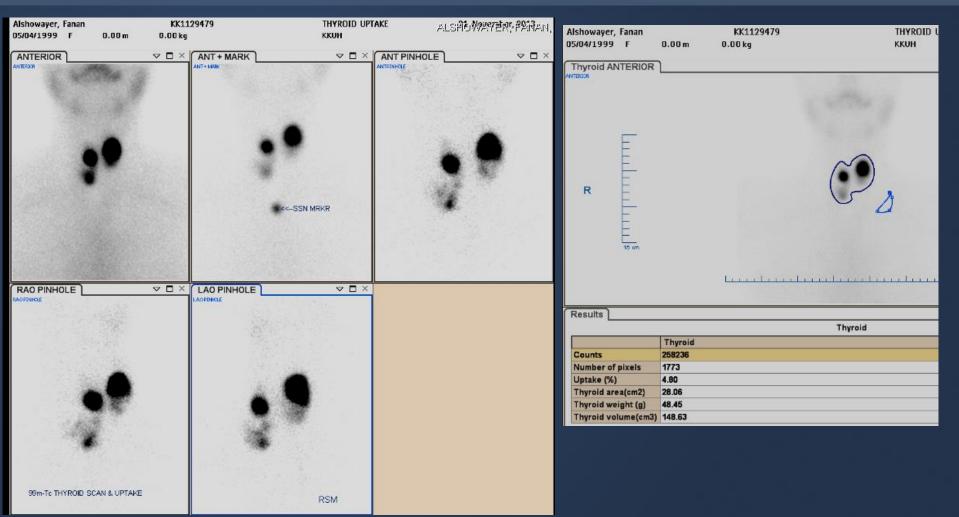
   (Jod-Basedow disease)
- Rare causes :

Execssive HCG by trophoblastic tumor Hypothalamic pituitary neoplasms (TSH induced)

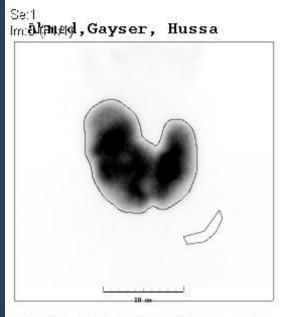




## **MNTG (Plummers Disease)**



### **Evaluation of thyrotoxicosis** Thyrotoxicosis with hyperthyroidism

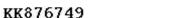


Patient Name : Ahmed, Gayser, Hussa Patient ID : KK876749 Exam Date : 20Apr2008 THYROID UPTAKE

Adac Laboratories BV Maarssen The Netherlands

ANTERIOR

#### <u>Graves' Disese on top of MNG</u> Nodular Graves Disease (Marine-Lenhart syndrome)



ANTERIOR SS SS

THYROID UPTAKE :47.46 %

Area 144.0 (sqcm) Mass 563.1 g A.GAYSER. ... MOHAM

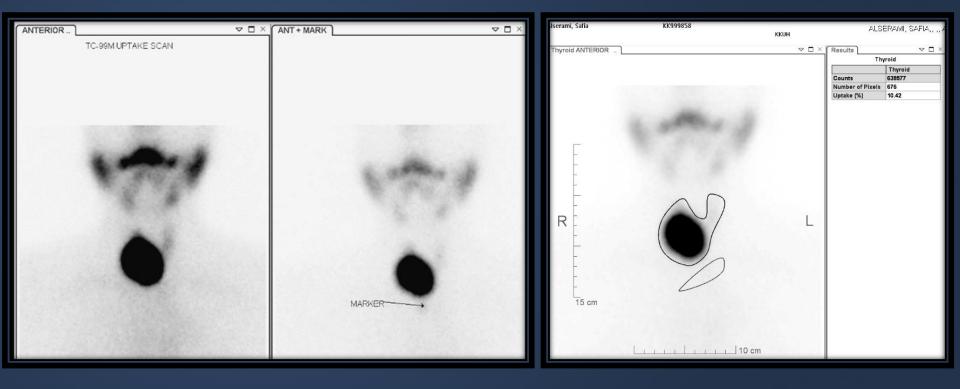
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## **Evaluation of thyrotoxicosis** Thyrotoxicosis with hyperthyroidism

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## **Evaluation of thyrotoxicosis** Thyrotoxicosis without hyperthyroidism

- Subacute thyroiditis.
- Chronic thyroiditis with transient thyrotoxicosis
- Thyrotoxicosis factitia (exogenous hormone).
- Thyroid extract (e.g.Hamburger thyrotoxicosis)
- Ectopic thyroid : Metastatic thyroid carcinoma Struma ovari



TC99M THYROID UPTAKE KKUH RM4 AA&MAY&A/RAHMAN

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## Radioactive Iodine Therapy for Hyperthyroidism



- Isotope used : I131
- Physical Properties: Solution or capsule
- Main side effect : Hypothyroidism
- Dose :
  - a. Calculated : Considering weight and uptake of the gland
  - b. Empirical :
    - Graves: 5-15 mCi
    - ATN : 15-20 mCi

Radioactive lodine Therapy for Thyroid Cancer



#### Isotope used : I131 Physical Properties: Solution or capsule

- Thyroid remnant : 80-100 mCi
- Lymph Node Mets : 100 mCi
- Local Recurrence : 100 mCi
- Lung Mets : 150 mCi
- Bone Mets : 200 mCi

# Parathyroid Scan





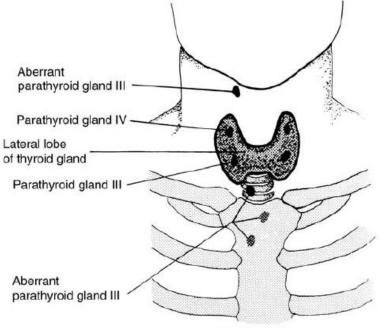


Figure 13.10. Normal and aberrant distribution of the parathyroid glands.

## Parathyroid Scan LEARNING OBJECTIVES...

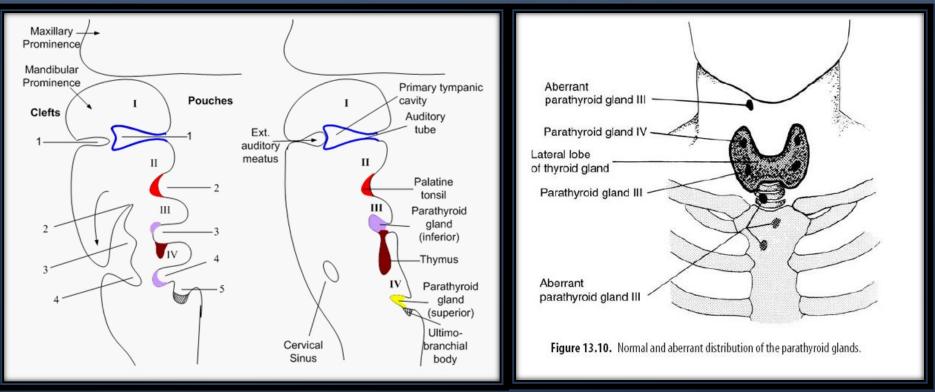


## At the end of the lecture you will be able to answer the following questions:

- Describe the physiologic principles of underlying Tc-99m parathyroid scintigraphy.
- Describe the various methods used for parathyroid scintigraphy with emphasis on SPECT and SPECT /CT .
- Identify the common imaging features of pathologic parathyroid glands.
- Discuss causes of false negative and false positive scans.



#### **Normal and Ectopic Parathyroid Glands**

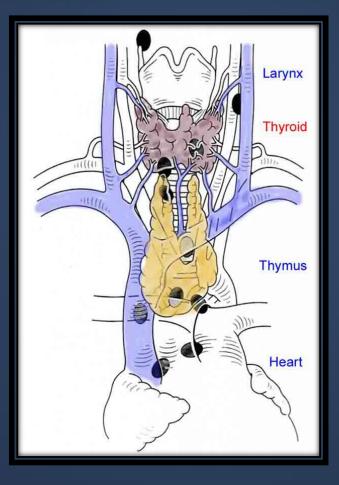


The third pair of pouches: proliferates into the inferior parathyroid glands and the thymus

The fourth pair of pouches: proliferates into the superior parathyroid glands and the lateral analge of the thyroid gland.

Because the inferior parathyroid glands undergo more extensive migration during embryogenesis, they are more likely to be found in ectopic locations.

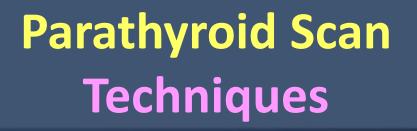
## **Ectopic Parathyroid Glands**



#### Location of an ectopic parathyroid glands

- Submandibular
- Retropharyngeal
- Retroesophageal
- Posterosuperior mediastinal
- Intrathyroidal
- Within the tracheoesophageal groove Carotid sheath
- Thyrothymic ligament
- Intrathymic

Antero-superior mediastinal.





- TL-201 \_ Tc-99m subtraction
- Tc-99m Sestamibi ( Dual Phase )
- Tc-99m Tetrofosmin ( Dual Phase )

## **Parathyroid imaging**



99mTc / 201Tl Subtraction Radiopharmaceutical 99mTc sestamibi Activity administered 80 MBg (2 mCi) 201Tl; 925 MBg (25 mCi) 370 MBg (10 mCi) 99mTc **Images acquired** Inject Tl .rst and acquire 15-min Anterior (and oblique) 100 000 count view of neck and views at 15 min and mediastinum. at 2–3 h; SPECT as needed Then acquire similar Tc images without moving patient. Subtract Tc data from Tl after normalization to equal count densities

## PARATHYROID IMAGING

#### Tc-99m-Sestamibi

The Parathyroid Study depicts hypertrophied parathyroid tissue, probably because of uptake of Tc-99m-sestamibi in the mitochondria of hyperactive cells.
 Indications : Detect and localize parathyroid adenomas .
 Patient Preparation :None.
 Radiopharmaceutical, Dose, & Technique of Administration

- Radiopharmaceutical: 25 mCi Tc-99m-sestamibi i.v.
- Patient position: Supine with head and neck extended and immobilized.
- Gamma camera Imaging field:
- 1. Neck.
- 2. Upper two thirds of the mediastinum.

Acquire images at 15 minutes and 2-3 hours post injection. SPECT/SPECT CT images improves localization.

• TI – Tc99m subtraction : Several protocols have been developed for routine subtraction of thyroid tissue from parathyroid tissue

#### **Parathyroid Scan** Dual phase MIBI Scan (Or Tetrofosmin)

MOHAMMED NAJI





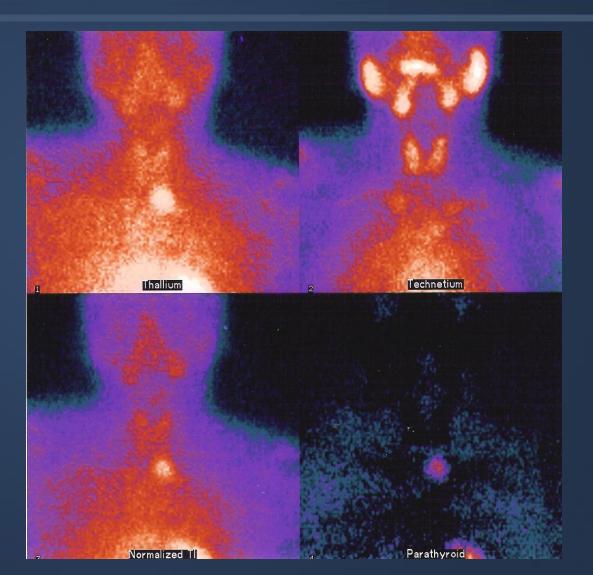


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Abnormal parathyroid glands could be visualized

Normal parathyroid glands are small and not visualized

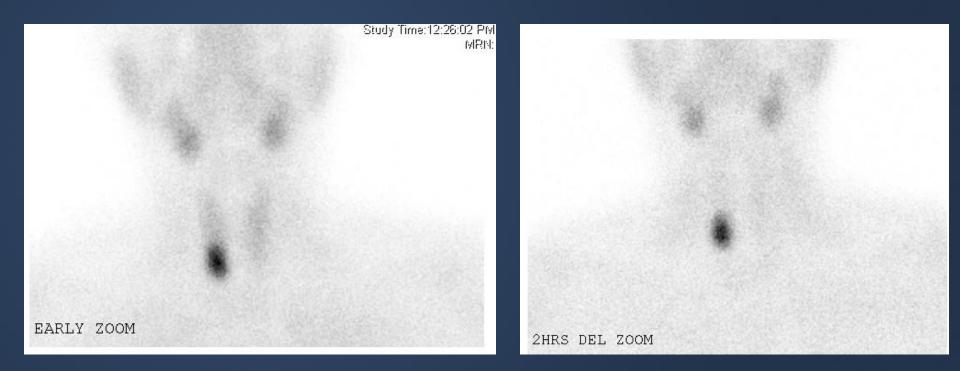
#### **Parathyroid Scan** Tc-Tl Subtraction Scan





## Parathyroid Scan Sestamibi dual phase

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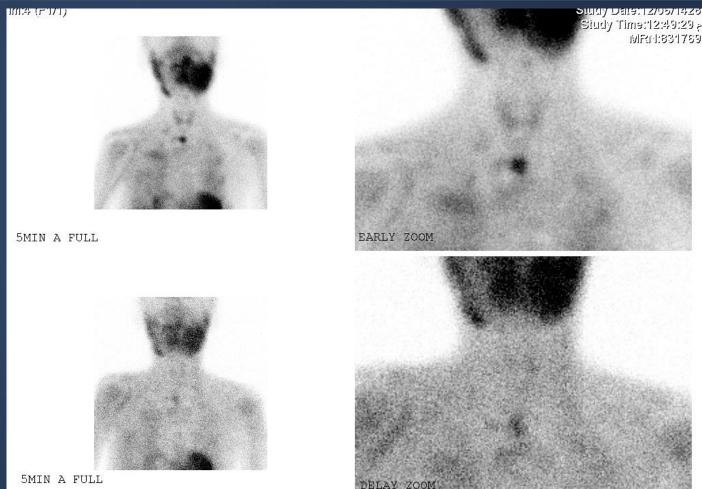
#### Sestamibi Dual Phase ( Planar vs SPECT CT)

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#### **Parathyroid Scan** Ectopic Parathyroid : 16% of total adenomas



KKUH

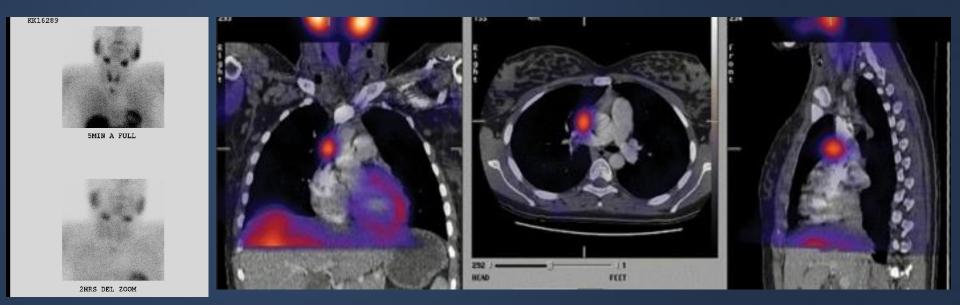
ROOM4

RSM

PARATHYROID MIBI

27Jun2007

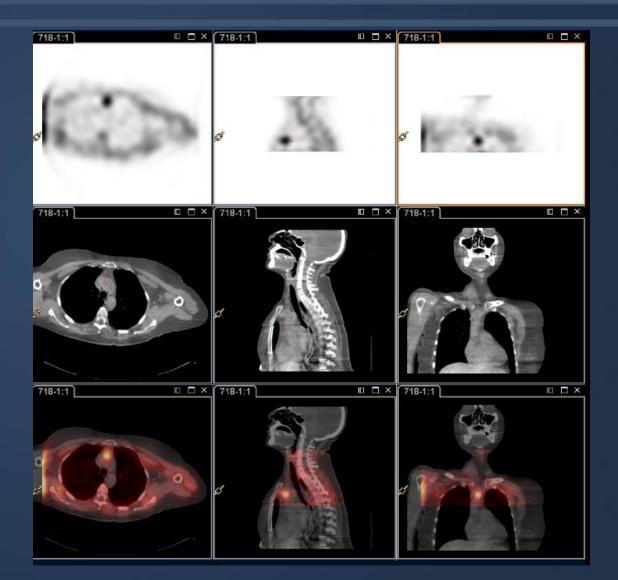
### Ectopic Parathyroid Adenoma PLANAR vs SPECT/CT



SPECT-CT images accurately localize the adenoma and guide the surgeon to the best surgical approach

#### **Ectopic parathyroid adenoma Antero-superior mediastinum**

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#### Sestamibi Parathyroid Scan Result



#### High PTH / High Ca

FN

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(H)



### Q:What is the cause of the FN result...?

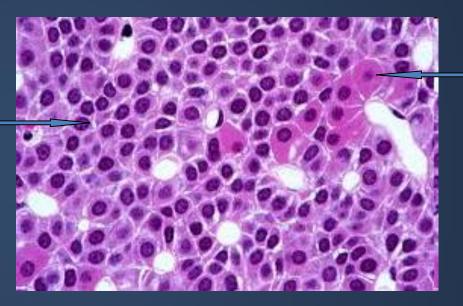


### A: Mechanism of sestamibi uptake



## **Parathyroid Cells**

**Chief cells** 



**Oxyphil cells** 

Normal parathyroid glands comprise 2 cell types:

 i. Chief cells: responsible for PTH production
 ii.Oxyphil cells: eosinophilic cells whose cytoplasm is composed almost entirely of mitochondria.

While the normal oxyphil cell does not synthesize and secrete PTH, the oxyphil cells of pathologic parathyroid glands do secrete the hormone.

### **Mechanism of Sestamibi uptake**

#### SESTAMIBI : METHOXYISOBUTYLISONITRILE

- Its parathyroid uptake was first reported by Coakley et al. in 1989
- Mechanism of MIBI uptake and retention is still unclear. Multifactors have been proposed:

#### a. Biochemical properties of the tracer :

Lipophilicity : The lipophilic sestamibi molecule is concentrated by mitochondria. This explains why adenomas with an abundance of mitochondrial-rich oxyphil cells retain the sestamibi Cationic charge

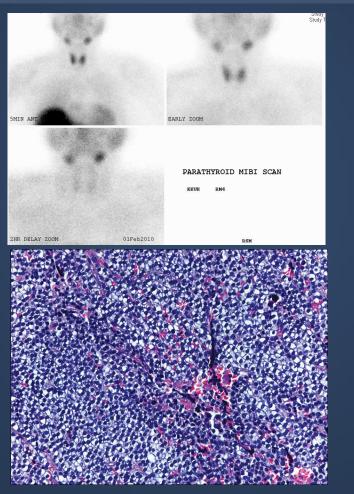
**b.** Cell Type : A predominance of oxyphil cells within an adenoma is more likely to lead to a positive scan.

**<u>c. Local factors:</u>** blood flow, trans-capillary exchange, interstitial transport and negative intracellular charge of both mitochondria and membranes.

J Nucl Med. 1990; 31: 1166-1167.

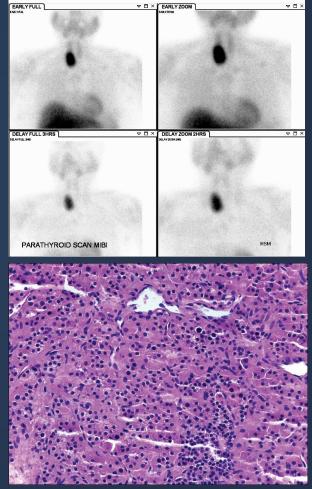
## **Cell Type and Scan Result**





Parathyroid adenoma composed entirely of

glycogen-rich chief cells



Parathyroid adenoma composed mainly of mitochondrial-rich oxyphil cells.



## What is the cause of the

## FN result...?

## "FN" Sestamibi Scan...?



- Histologic type : False-negative scans can occur with parathyroid glands containing predominantly clear cells.
- Size and Location: Smaller-volume parathyroid adenomas and those in the upper position are less likely to be localized with sestamibi scans.
- Number of adenomas: FN rate is increased with MGD compared with patients with a single adenoma .
- **Decreased tracer concentration : Possible association.**

a. P-glycoprotein expression

b. Multidrug resistance-related protein expression

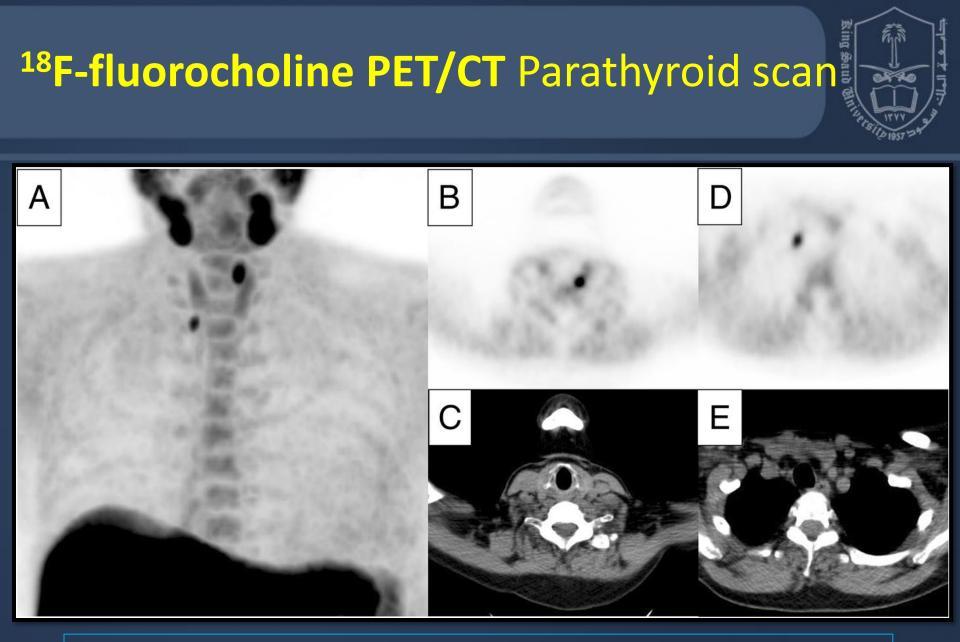
• Variability of radiotracer uptake in parathyroid adenomas: Related to differences in perfusion and metabolic activity

*Even with refinements in sestamibi scanning, the fact that all parathyroid adenomas are not created equal on a cellular level may inevitably lead to FN scans in a certain number of cases.* 

Arch Surg. 2007;142(4):381-386.



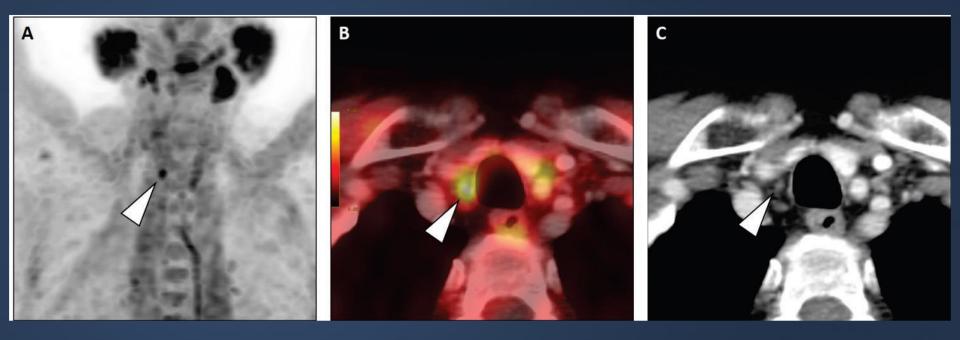
## <sup>18</sup>F-fluorocholine PET/CT



**Double parathyroid adenoma (SESTAMIBI scan was negative)** 



#### <sup>18</sup>F-fluorocholine PET/CT Parathyroid scan



Focal hyperactivity posterior to the right caudal pole of right thyroid lobe and, histologically confirmed as parathyroid adenoma.

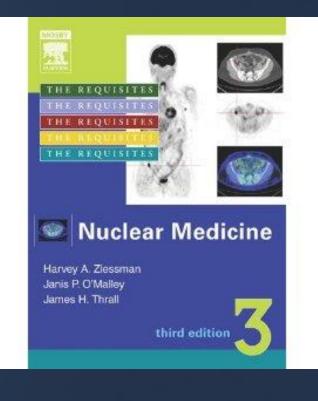
Previous imaging by neck US and MIBI was inconclusive

# Reference book and the relevant page numbers..

• Nuclear Medicine: The Requisites, Third Edition (Requisites in Radiology) [Hardcover]

Harvey A. Ziessman MD, Janis P. O'Malley MD, James H. Thrall MD

<u>Relevant Pages :</u> I- Thyroid and Parathyroid : 71-105 II- Oncology : 264-274 , 279 -283 ,302 -345 ,119-133 109 -112 ,296 -299



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