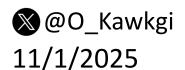


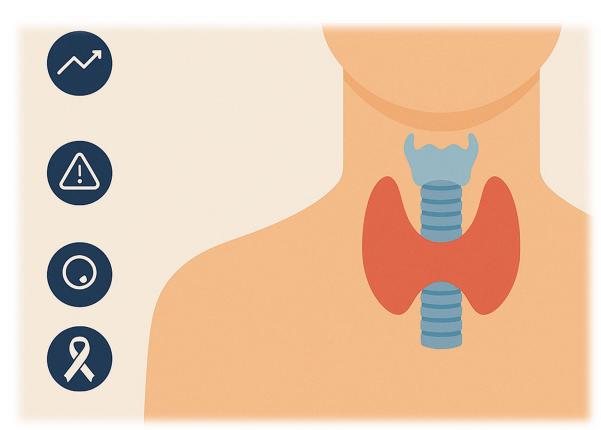
Overview of Thyroid Disorders

Fall Family Medicine Review Course

Omar M. El Kawkgi, MD **Assistant Professor of Medicine** Omar.elkawkgi@uky.edu







LEARNING OBJECTIVES



Evaluate and manage hyperthyroidism and hypothyroidism, including subclinical disease



Diagnose and manage thyroid medical emergencies



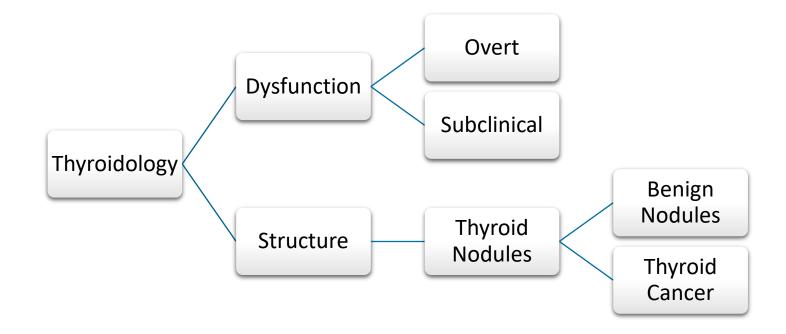
Evaluate and manage thyroid nodules



Describe and manage different types of thyroid cancer



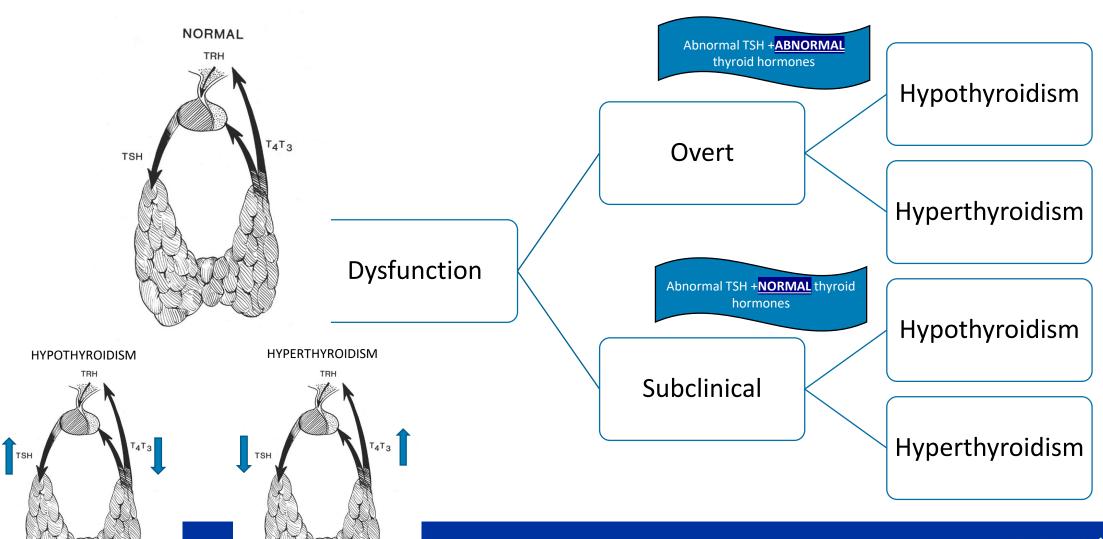
Schematic Overview





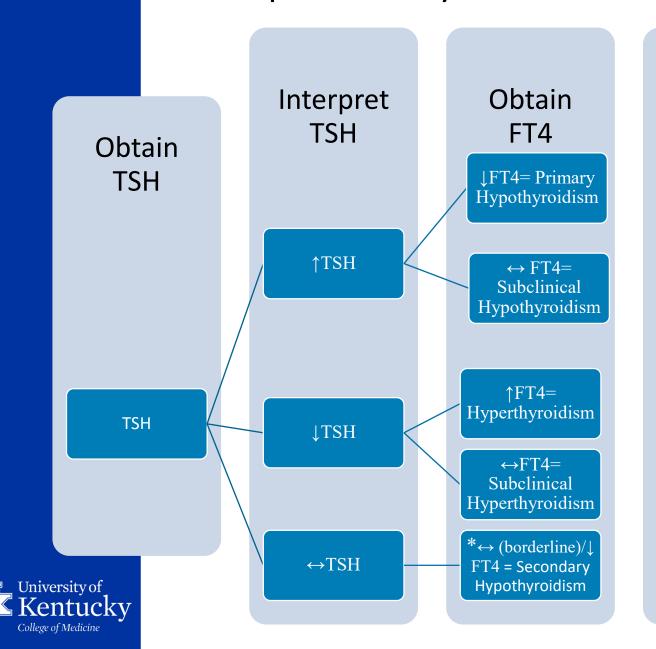
THYROID DYSFUNCTION: TOO MUCH OR TOO LITTLE

Thyroid Dysfunction





4 Steps to Thyroid Function Problems



Determine etiology

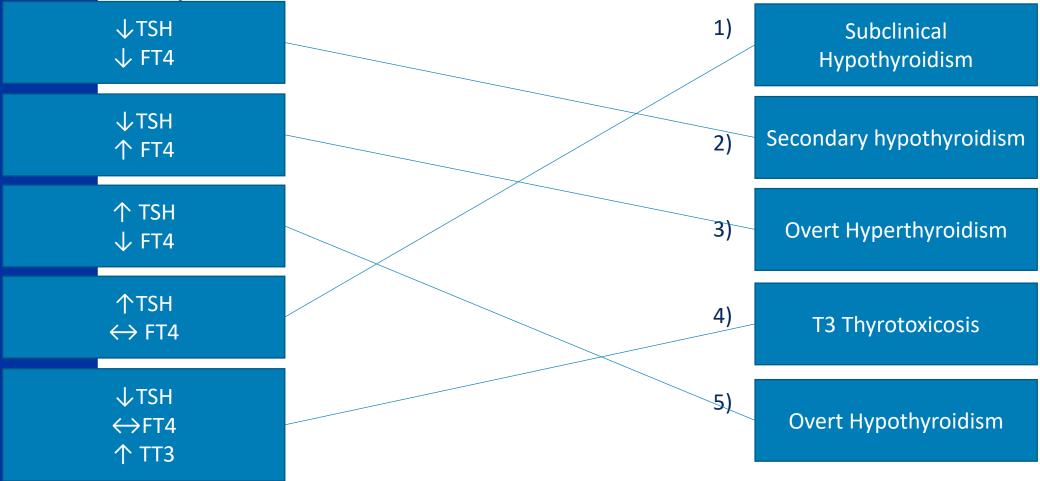
*If normal TSH: Only obtain FT4 if clinically suspect secondary hypothyroidism

O /			
Autoimmune (Hashimoto's) Thyroiditis	Usually associated with positive TPO antibody		
Postablative	Following RAI/ Surgery		
Transient thyroiditis	Hypothyroid phase of thyroiditis		
Drugs	Lithium, amiodarone, TKI's, Immune checkpoint inhibitors, Valproic acid, Interferon		
Infiltrative disease	Hemochromatosis, amyloidosis, sarcoidosis, scleroderma, etc.		
Genetic	Thyroid agenesis/ dysgenesis		
Head and Neck radiation	History of treated head and neck cancer		

Graves' Disease	Autoimmune activation of TSH receptor	
Transient thyroiditis	Short lived and usually self resolves + possible hypo phase	
Toxic Nodule/ Multiple toxic nodules	Autonomous function of one ore more thyroid nodule(s)	
Drugs	Lithium, amiodarone, Immune checkpoint inhibitors,	

Pituitary or hypothalamic disease	Lack of signal to thyroid
Drugs	Glucocorticoids

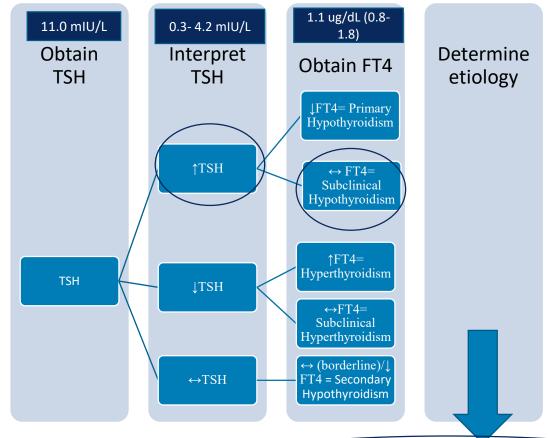
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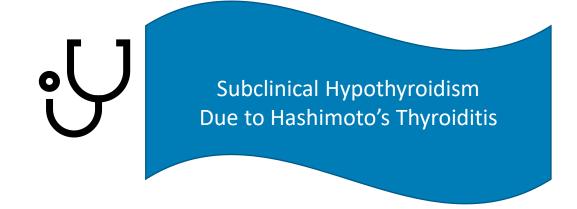




Case #1

- 55-year-old female
- Complains of mild fatigue
- Physical exam:
 - ➤ Diffuse goiter
- TPO antibody positive

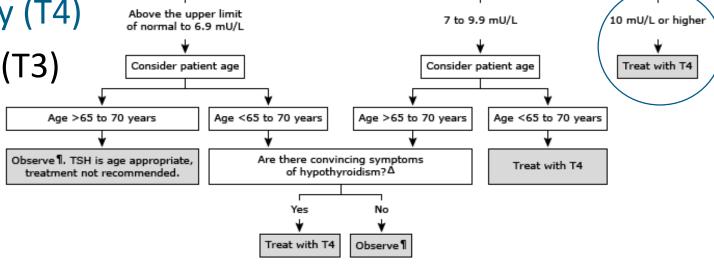




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Head and Neck radiation	History of treated head and neck cancer			

- A) Start levothyroxine therapy (T4)
- B) Start liothyronine therapy (T3)
- C) Recheck in 3 months
- D) Obtain thyroid ultrasound

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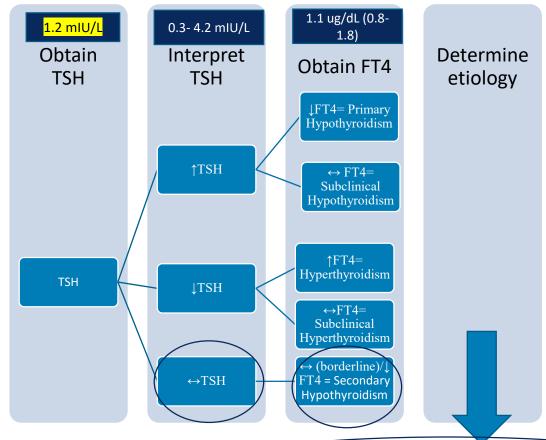
What is the TSH?

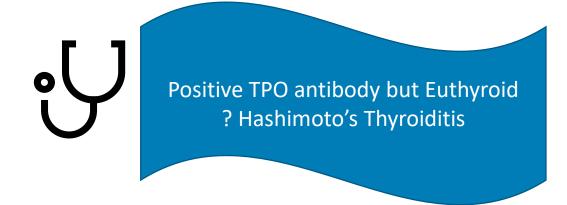
Teaching Point

Treat hypothyroidism with LT4
Start when TSH > 10 or if highly symptomatic*, TPO +'ve
High TSH may be normal at older age

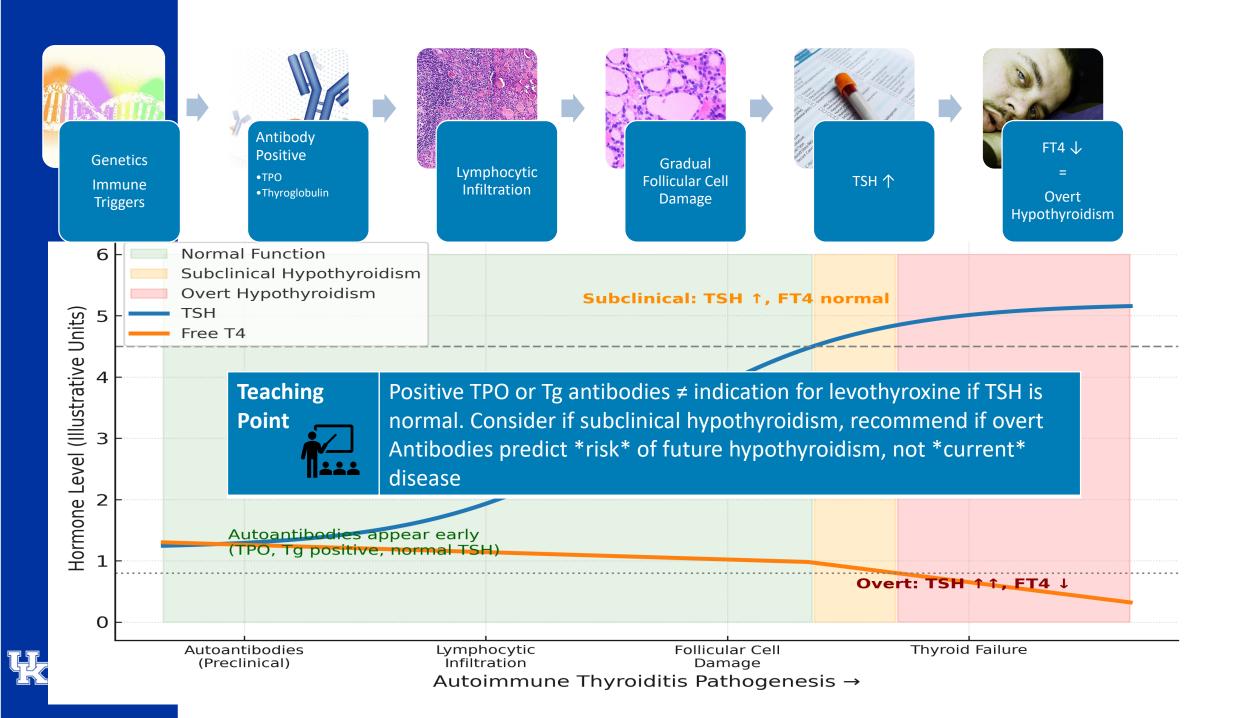
Don't Treat the Antibody Treat the Thyroid Function

- 55-year-old female
- Complains of mild fatigue
- Physical exam:
 - ➤ Diffuse goiter
- TPO antibody positive





Autoimmune (Hashimoto's) Thyroiditis	Usually associated with positive TPO antibody			
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Genetic	Thyroid agenesis/ dysgenesis			
Head and Neck radiation	History of treated head and neck cancer			



Case #1 Continued

Started on LT4

Her other medications include omeprazole 20 mg daily, calcium supplements, lisinopril 5 mg daily

She is taking her medications and supplements in the morning with breakfast

6 weeks later:

- TSH 10 mIU/mL (↑)
- FT4 1.2 (normal)



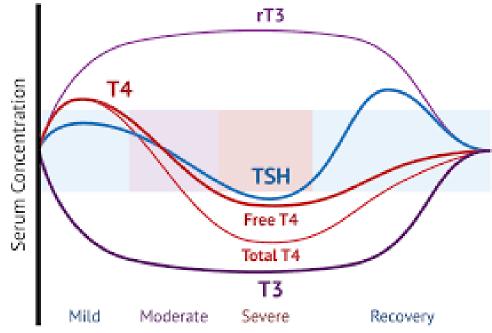
- A) Increase dosage of levothyroxine
- B) Continue current dose of levothyroxine and add T3 therapy
- C) Assess compliance and educate on how to take levothyroxine
- D) Obtain levothyroxine absorption test

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- B) Continue current dose of levothyroxine and add T3 therapy
- C) Assess compliance and educate on how to take levothyroxine
- D) Obtain levothyroxine absorption test



Mild Elevated TSH not Associated with Subclinical Hypothyroidism

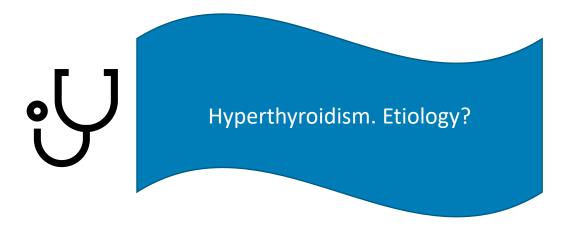
- Non-thyroid disease related etiologies for elevated TSH:
 - Non-thyroidal illnesses/critical illness
 - Age age appropriate rise in TSH
 - o Drugs: metoclopramide, domperidone
- Spuriously elevated TSH
 - Assay variability
 - Heterophile antibodies
- Pituitary-related rises in TSH
 - TSH-secreting adenoma rare (high TSH & normal or high T4/T3)
 - Resistance to TSH

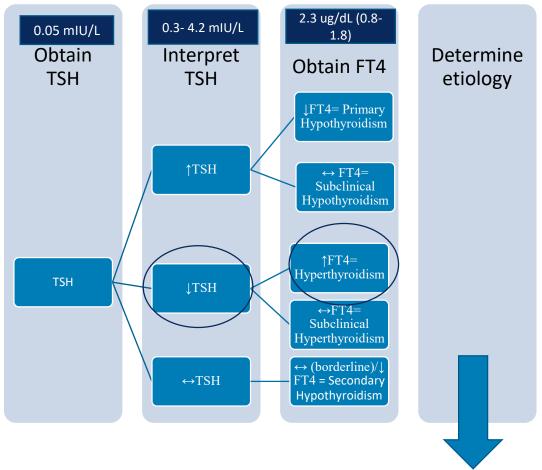




Case #2

- 23-year-old female
- Complains of weight loss and irregular menstrual cycles
- Physical exam:
 - ➤ Right sided thyroid nodule
 - **≻**Tremor





Graves' Disease	Autoimmune activation of TSH receptor	
Transient thyroiditis	Short lived and usually self resolves + possible hypo phase	
Toxic Nodule/ Multiple toxic nodules	Autonomous function of one ore more thyroid nodule(s)	
Drugs	Lithium, amiodarone, Immune checkpoint inhibitors,	

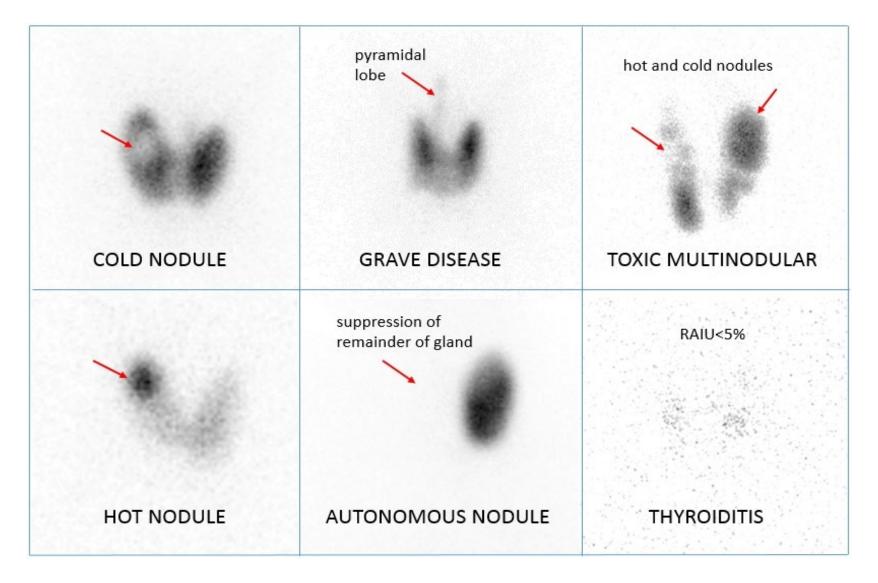
- A) Start methimazole 20 mg daily
- B) Recheck thyroid function in 3 months
- C) Obtain I-131 thyroid uptake and scan
- D) Start PTU 150 mg daily

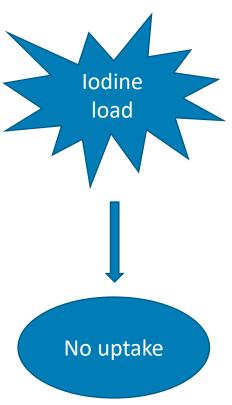
- A) Start methimazole 20 mg daily
- B) Recheck thyroid function in 3 months
- C) Obtain I-131 thyroid uptake and scan
- D) Start PTU 150 mg daily

If given option of TRAb testing vs. uptake and scan → chose TRAb first



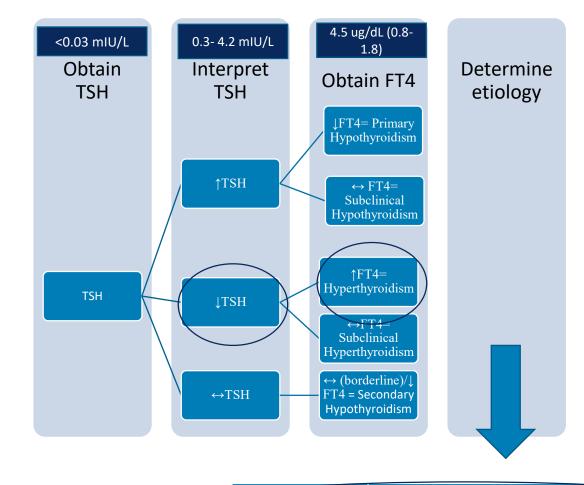
Interpreting Thyroid Uptake and Scan

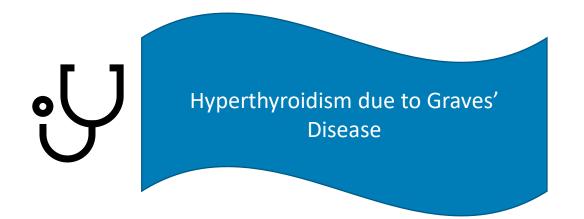




Case #3

- 33-year-old female
- Complains of palpitations
- Physical exam:
 - ➤ Diffuse goiter
 - ➤ Bilateral proptosis
 - ➤ Conjunctival erythema
- TRAb antibody positive





Graves' Disease	Autoimmune activation of TSH Receptor
Transient thyroiditis	Short lived and usually self resolves + possible hypo phase
Toxic Nodule/ Multiple toxic nodules	Autonomous function of one ore more thyroid nodule(s)
Drugs	Lithium, amiodarone, Immune checkpoint inhibitors,

- A) Obtain I-131 thyroid uptake and scan
- B) Start methimazole 20 mg daily
- C) Treat with 10 mCi of radioactive iodine
- D) Obtain TSI antibody level

- A) Obtain I-131 thyroid uptake and scan
- B) Start methimazole 20 mg daily
- C) Treat with 10 mCi of radioactive iodine
- D) Obtain TSI antibody level



Avoid RAI treatment in Graves' disease and active thyroid eye disease. TRAb is the most sensitive marker for diagnosis.

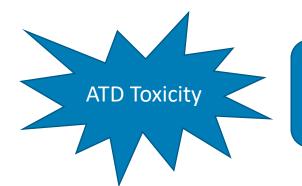
Graves' Disease

- Most common cause of hyperthyroidism
- Presents with classic hyperthyroid symptoms
- Goiter
- Specific P/E findings:
 - > Thyroid eye disease
 - Pre-tibial myxedema
 - > Thyroid acropachy
- TRAb/ TSI positive in most cases
- Diffuse uptake on uptake and scan
- Treated with:
 - ➤ Anti thyroid drugs- first line
 - > RAI
 - ➤ Surgery









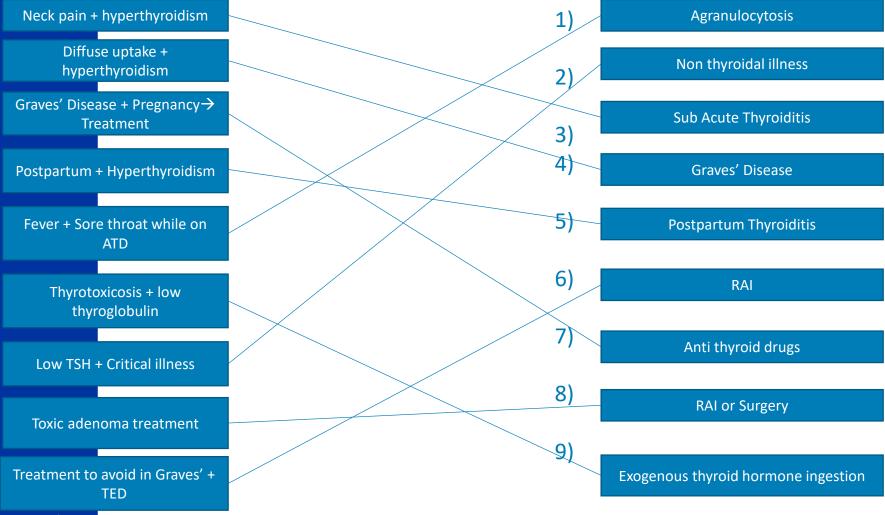
Agranulocytosis, rash, hepatoxicity

Other Causes of Abnormally Low Serum TSH

- Subclinical hyperthyroidism
- Recovery from hyperthyroidism (lag in TSH suppression)
- Pituitary / hypothalamic disease (adenoma)
- Euthyroid sick syndrome (non-thyroidal illness syndrome)
- Some medications such as corticosteroids, dopamine, dobutamine
- First trimester of pregnancy



If you see this Choose This.





Thyroid Dysfunction Emergencies

Thyroid Storm:

Trigger: acutely ill, iodine load, etc.

- Fever (thermoregulatory dysfunction)
- CNS (metabolic encephalopathy)
- Severe cardiovascular problems (marked sinus tachycardia, atrial arrhythmias, heart failure, hypotension)
- Gastrointestinal and hepatic (vomiting, diarrhea, abnormal LFTs)
- -Treatment:

ICU, beta blockers, anti-thyroid drugs (PTU) then SSKI, glucocorticoids

Myxedema Coma:

Trigger: acutely ill, dehydration, opiates, medication nonadherence

- Hypothermia (thermoregulatory dysfunction)
- CNS (metabolic encephalopathy)
- Severe cardiovascular problems (bradycardia, hypotension, heart failure)
- Respiratory (hypoventilation, hypoxia)
- Biochemical (hyponatremia, hypoglycemia)
- -Treatment:

ICU, hydrocortisone (until concomitant AI ruled out), levothyroxine +/- T3

Teaching Point

Clinical diagnosis. No thyroid hormone level is diagnostic. Give ATD **BEFORE** lodine in Thyroid storm.

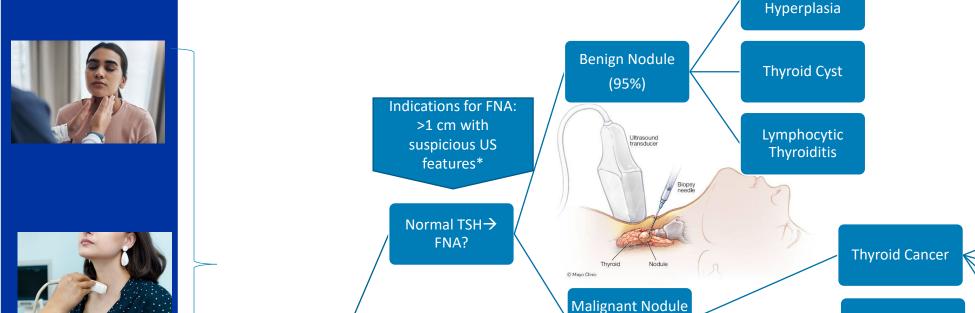
Thyroid Dysfunction High Yield Pearls



- 1) TSH is the single most sensitive test in detecting thyroid disease as it is affected BEFORE T4*
- 2) I- 131 Scan cannot be interpreted in patients who received a high iodine load (contrast, amiodarone, etc.) within the last 30-90 days
- 3) Treat thyroid storm with steroids, beta blockers, PTU, iodine
- 4) Treat Graves' disease with ATD, RAI or surgery. Never use RAI in pregnancy or in those with TED

THYROID STRUCTURE: LUMPS & BUMPS

Thyroid Structural Disorders





Thyroid Nodule





Low TSH → Benign "Hot" Thyroid Uptake Nodule and Scan

(5%)

Adenoma/

Lymphoma

Metastasis to thyroid

Follicular Thyroid Cancer

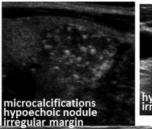
Papillary Thyroid Cancer

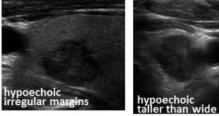
Medullary Thyroid Cancer

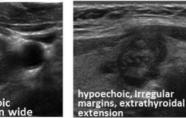
Anaplastic Thyroid Cancer

ATA Nodule Sonographic Pattern Risk of Malignancy

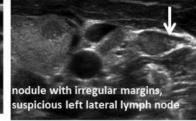
High Suspicion 70-90%











Intermediate Suspicion 10-20%

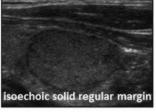
> Low Suspicion 5-10%

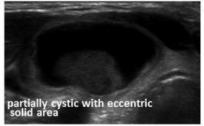














Very low Suspicion <3%

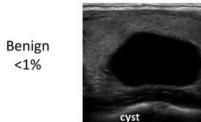
University of Kentucky

College of Medicine





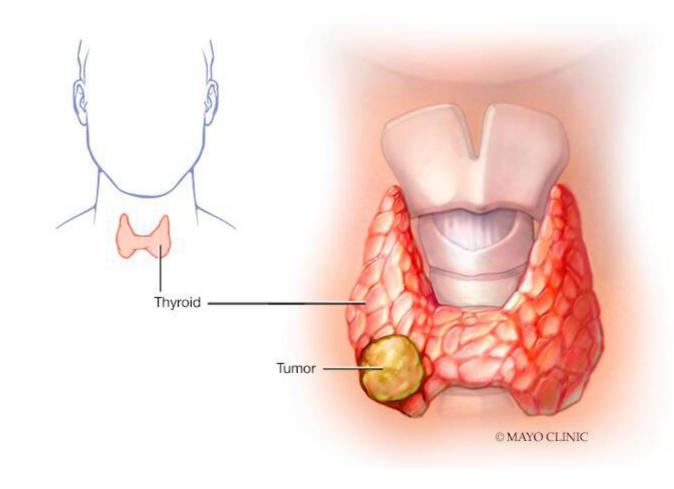






Risk Factors for Thyroid Cancer

- Age >70 or <14
- Head/ neck radiation history
- Family history
 - ➤ PTC: FAP, Cowden, Carney complex
 - ➤ Follicular: Werner Syndrome
 - ➤MTC: MEN2a, MEN 2b
- Hashimoto's → Lymphoma?

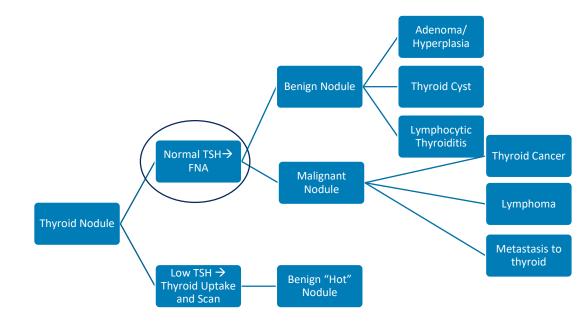


Types of Thyroid Cancer

Туре	Incidence	Survival Rate	Clinical Characteristics
Papillary	70-80%	95%	 Local spread Lymph node mets. Slow growing Biochemical marker: Thyroglobulin
Follicular	10-15%	91%	 Hematological spread Bone and lung mets. Slow growing Biochemical marker: Thyroglobulin
Medullary	5-10%	91%	 May be sporadic or A/W MEN 2A or MEN2B RET mutation Biochemical marker: Calcitonin
Anaplastic	1-2%	1-7%	Locoregional invasionBone and lung mets.Fast growing

Case #4

- 49-year-old female
- CT neck obtained in ED due to MVA
- Incidental thyroid nodule
- Thyroid US: 1.8 cm left thyroid nodule of intermediate suspicion
- Lab Results:
 - >TSH 0.8 (0.3-4.2 mIU/L)
- Physical exam:
 - ➤ Palpable left nodule
 - ➤ Mobile with swallowing
 - ➤ No cervical lymphadenopathy



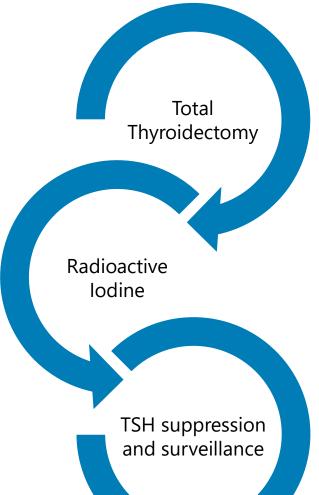
- A) Obtain thyroglobulin level
- B) Obtain calcitonin level
- C) Refer for surgery
- D) Perform fine needle aspiration biopsy

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- B) Obtain calcitonin level
- C) Refer for surgery
- D) Perform fine needle aspiration biopsy

Teaching Point

Thyroglobulin level is not helpful in determining malignant vs. benign nodules. Calcitonin is only used when medullary thyroid cancer is suspected.

Historical Approach To Thyroid Cancer Treatment



RADIO-IODINE HALTS ONE TYPE OF CANCER

Radioactive chemical brings about history-making recovery of patient dying from thyroid tumors

The man shown in the contrasting portraits at right is a Brooklyn shoe salesman named Bernard Brunstein who is destined to become one of the most famous patients in medical history. Brunstein is the first person known to be cured (insofar as a cure can be established by medical tests on a living patient) of metastatic cancer, a form of the disease in which the malignancy spreads through the body from an original tumor. Metastatic cancer has always been 100% fatal. But Brunstein's tumors were destroyed in a simple, almost miraculous way: by the drinking of four doses of radioactive iodine. When Brunstein was admitted to New

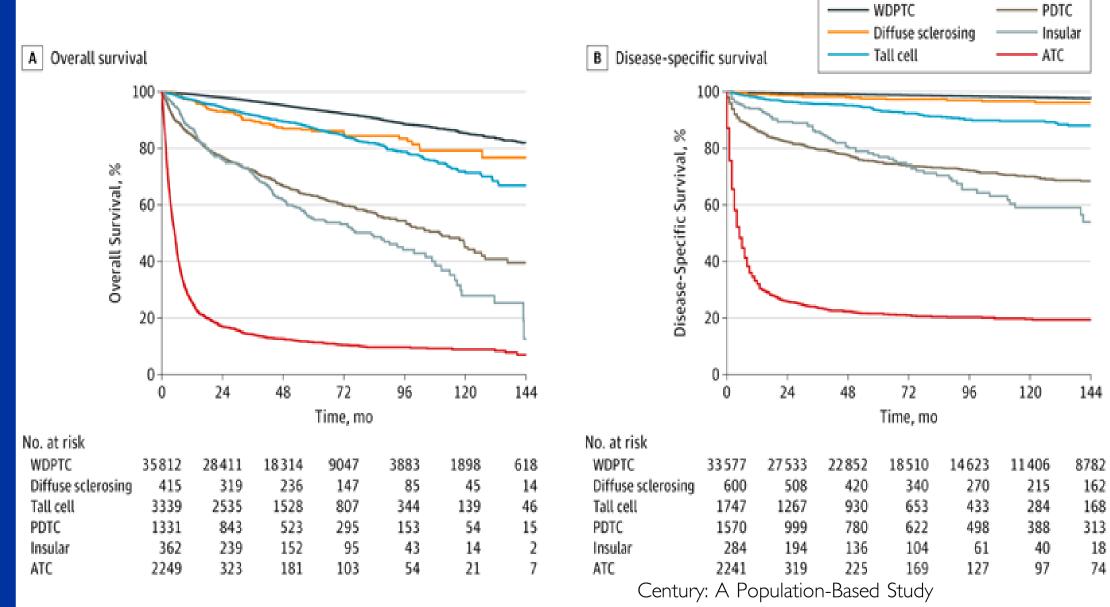




BERNARD BRUNSTEIN IN 1942 (LEFT): AS HE LOOKS TODAY

iodine is chemically identical with ordinary iodine, it gives off a powerful radiation that can kill any tissue that absorbs it in sufficient concentration. The chemical had never been effectively used as a treatment for cancer, but Brunstein agreed to try it in the hope that it might help. It did. Three months after he drank his first glassful of the tasteless, colorless liquid, his heart began to slow down and he started to put on weight. Geiger counters placed over the tumor sites revealed that there was a heavy concentration of radio-todine in these areas. After three additional doses the tumors slowly began to diminish in size







Natalia Genere, MD; Omar M. El Kawkgi, MB, BCh, BAO; Rachel E. Giblon, MS; Salvatore Vaccarella, PhD; John C. Morris, MD; Ian D. Hay, MD, PhD; and Juan P. Brito, MBBS

STRIKING A BALANCE

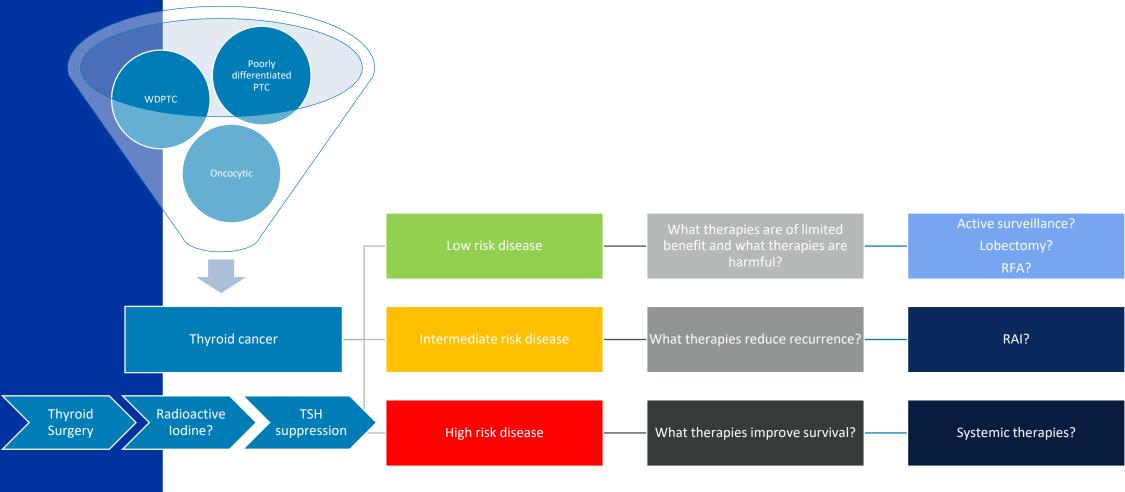
Under-Treatment of Intermediate or High Risk Disease



Over-Treatment of Low Risk Disease



Heterogenous Group of Diseases Need Individualized Approaches





Low Risk Disease-Extent of Surgery and Recurrence

ORIGINAL SCIENTIFIC REPORT

Thyroid Lobectomy for Low-Risk 1–4 CM Papillary Thyroid Cancer is not Associated with Increased Recurrence Rates in the Dutch Population with a Restricted Diagnostic Work-Up

J. F. Lin¹ · P. M. Rodriguez Schaap · M. J. H. Metman · E. J. M. Nieveen van Dijkum · C. Dickhoff · T. P. Links · S. Kruijff · A. F. Engelsman ·

901 patients diagnosed with low risk 1-4 cm PTC between 2005-2015 with median follow up of 7.7 years

2.6% recurrence rate in cohort

No SS difference in TT vs. TL



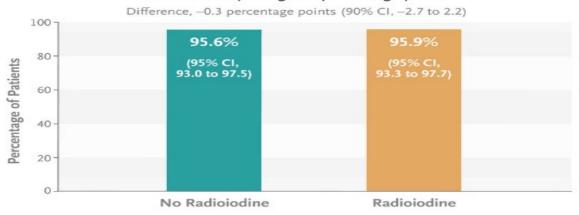
Lin JF, Rodriguez Schaap PM, Metman MJH, Nieveen van Dijkum EJM, Dickhoff C, Links TP, Kruijff S, Engelsman AF. Thyroid Lobectomy for Low-Risk 1-4 CM Papillary Thyroid Cancer is not Associated with Increased Recurrence Rates in the Dutch Population with a Restricted Diagnostic Work-Up. World J Surg. 2023 May;47(5):1211-1218.

Low Risk Disease- Role for RAI?

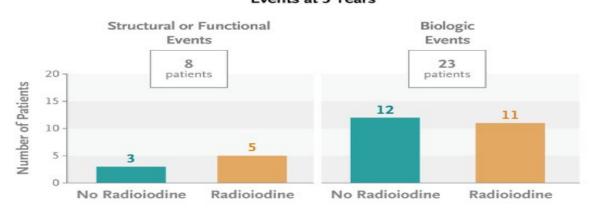


No Event at 3 Years

Noninferiority margin, 5 percentage points



Events at 3 Years





2025 ATA DTC Guideline Recommendations

Risk category	Typical RAI recommendation	Recommended ¹³¹ I activity level	Goals of therapy
Low	No	1.1-1.85 GBq (30-50 mCi)	None or remnant ablation
Intermediate-low and intermediate-high	Consider	1.1-3.7 GBq (30-100 mCi)	Remnant ablation +/- adjuvant therapy
High	Yes	3.7-5.55 GBq (100-150 mCi)	Remnant ablation and adjuvant therapy
Distant metastases	Yes	3.7-7.4 GBq (100-200 mCi) or consider dosimetry	Treatment of known disease, remnant ablation



Thyroid Structure High Yield Pearls



- 1) Incidence: PTC > Follicular > Medullary> Anaplastic
- 2) Do not FNA hot nodules (Low TSH, High localized uptake)
- 3) Do FNA cold nodules that are > 1 cm + US features
- 4) No role for pre-operative tumor markers
- 5) Not all thyroid cancers require Surgery, RAI and TSH
 Suppression- low risk PTC cancers can be managed with
 lobectomy without RAI

Questions?

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