

# **74 Year Old male with H&N cancer and abnormal TFT**

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# HPI

- ❖ 74 year old M with PMH of HTN, recently Dx Sq.C.Ca of the tongue
- ❖ Was in his usual state of health until 8 months ago
- ❖ c/o Wt loss, difficulty swallowing, voice change, intermittent palpitation
- ❖ Found to have tongue lesion (Sq.c.ca on biopsy)
- ❖ Underwent hemi-glossectomy, elective bilateral neck dissection (3/10/14).
- ❖ During surgery found to have large Rt side thyroid nodule → Rt side thyroid Lobectomy.
- ❖ We were consulted for abnormal TFT

# ROS

**Constitutional:** Wt loss (40 pounds over 8 months)

**HENT:** No blurred vision, no double vision, no headache

**Neck:** no neck pain, no neck swelling, + **difficulty swallowing**, + **voice change**.

**Cardio/pulm:** No CP, + **intermittent palpitation**, no orthopnea or PND

**GI:** No N/V/D, no constipation, no melena or hematochezia

**GU:** Negative,

**Skin/MSK:** negative, no rash

**Neuro** denied any tremors or weakness.

## **PMH:**

- ✓ Colon cancer s/p surgical resection and chemo in 2006
- ✓ HTN
- ✓ BPH
- ✓ Recently Dx Sq.C.Ca (tongue)

## **Family History:**

- ✓ No FH of thyroid cancer
- ✓ No FH of any other thyroid disease

## **Surgical history:**

- ✓ Total colectomy 2006

## **Social history**

- ✓ Quit smoking 8 months ago, drink alcohol socially, no illicit drugs.

## **Home medications**

- ✓ Atenolol 50 mg po daily
- ✓ Doxazosin 2 mg po daily
- ✓ Multivitamins

# On examination

**Vitals:** BP 134/94, Pulse 71, no fever, RR 18, BMI 19

**General:** awake alert, comfortable, thin man

**HEENT:** normocephalic non traumatic, no pallor, no jaundice. No double vision, no increase insertion, no exophthalmos

**Neck:** supple, s/p Rt lobectomy and bilateral neck dissection

**CVS/Pulm:** clear equal air entry no added sounds, regular pulse, S1 + S2, no murmur.

**Abd:** soft lax, no organomegaly, no tenderness, audible bowel sounds.

**Skin:** normal, not diaphoretic

**Neuro:** alert, no tremor, CN intact, DTR normal

**Psych:** normal mood, and affect

# General labs

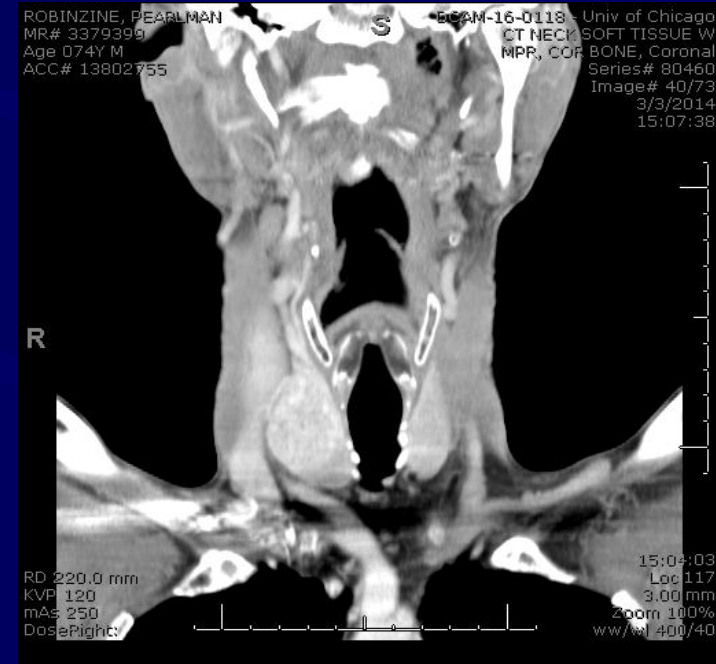
Test/date	3/10/2014
Na/K	138/4.0
BUN/Cr	6/0.6
eGFR	110
ALP	83
Ca	9.5
ALT/AST	9/12
Hb	11.6
WBC	7.6
Plt	322

# TFT

Test/date	3/3	3/13	3/17
TSH (0.4 – 4.5)	<0.01	<0.01	<0.01
FT4 (0.8 – 1.8)	1.86	1.0	0.50
T3 (20 – 195)	184	66	37
rT3 (160 – 353)		380	
TPO&Tg Abs	<0.4		



Surgery 3/10



Neck CT scan prior to the surgery



# Clinical Qs

- What is the pattern of recovery of hypothalamic-Pituitary-Thyroid axis following treatment of hyperthyroidism?
- What is the cause of central hypothyroidism (direct pituitary thyrotroph suppression or hypothalamic TRH deficiency)?
- Is the length of preexisting hyperthyroidism and baseline free T4 predict the duration of the central hypothyroid phase?

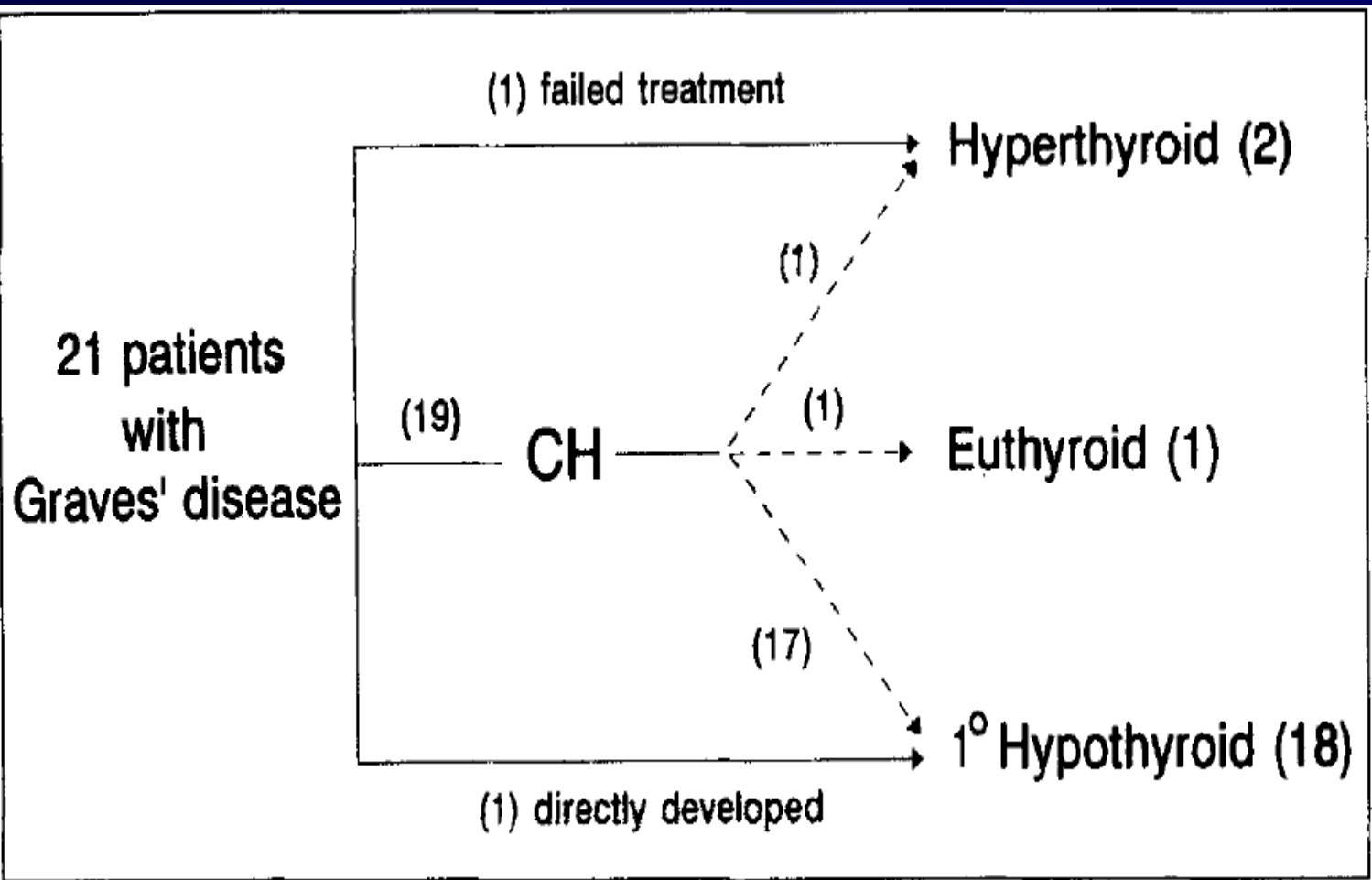




# **Pattern of Recovery of the Hypothalamic-Pituitary-Thyroid Axis Following Radioactive Iodine Therapy in Patients with Graves' disease**

Harry L. Uy, MD, Charles A. Reasner, MD, Mary H. Samuels, MD, San Antonio, Texas

- ✓ 21 subjects with Graves' disease (17 women and 4 men), aged 18 to 54 years entered the study.
- ✓ Patients who were pregnant, who had underlying psychiatric or cardiopulmonary illness, or who required medications known to affect TSH values were excluded from the study.
- ✓ None of the patients were treated with thionamides or iodides after receiving radioactive iodine. Beta blockers were continued as required to control hyperadrenergic symptoms.



TABLE

**Characteristics of 19 Patients With Graves' Disease  
Who Experienced a Central Hypothyroid Phase Following <sup>131</sup>I Therapy**

Patient No.†	<sup>131</sup> I Dose (mCi)	Onset of CH (Days)	Duration of CH (Days)	Thyroid Profile at Onset of CH		
				Free Thyroxine (0.8–2.7 ng/dL)	Triiodothyronine (90–190 ng/dL)	TSH (0.39–4.6 mU/L)
1†	17.7	29	28	0.5	60	0.09
2	29.9‡	50	18	0.4	70	0.01
3†	11.2	84	14	0.4	41	1.1
4†	9.8	119	14	NA§	NA	0.1
5†	15	68	16	0.4	47	1.6
6	19.8	55	28	0.6	43	0.01
7	16.3	55	47	0.7	60	0.01
8	20.4	56	15	0.4	74	0.08
9	15.5	27	28	0.7	59	0.02
10	13.7	71	19	0.3	52	0.01
11	12.1	85	14	0.5	80	0.4
12†	15.4	44	28	0.6	45	0.01
13†	12.1	65	35	0.5	44	0.01
14	10	73	14	0.4	56	3.5
15	15.8	27	14	0.5	64	0.7
16	20	69	43	0.8	71	0.02
17†	13.6	68	28	0.5	51	0.02
18†	12.3	72	31	0.6	42	0.03
19†	9.4	77	35	0.7	56	0.02

†Following treatment, patients 1–17 became hypothyroid; 18 became euthyroid; and 19 became subclinically hyperthyroid.

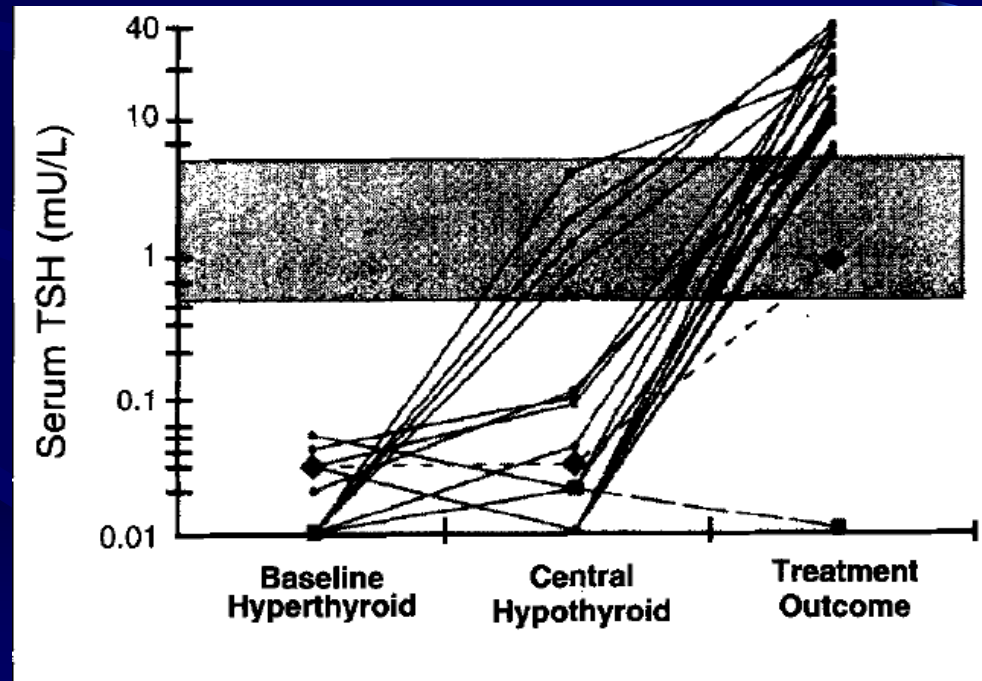
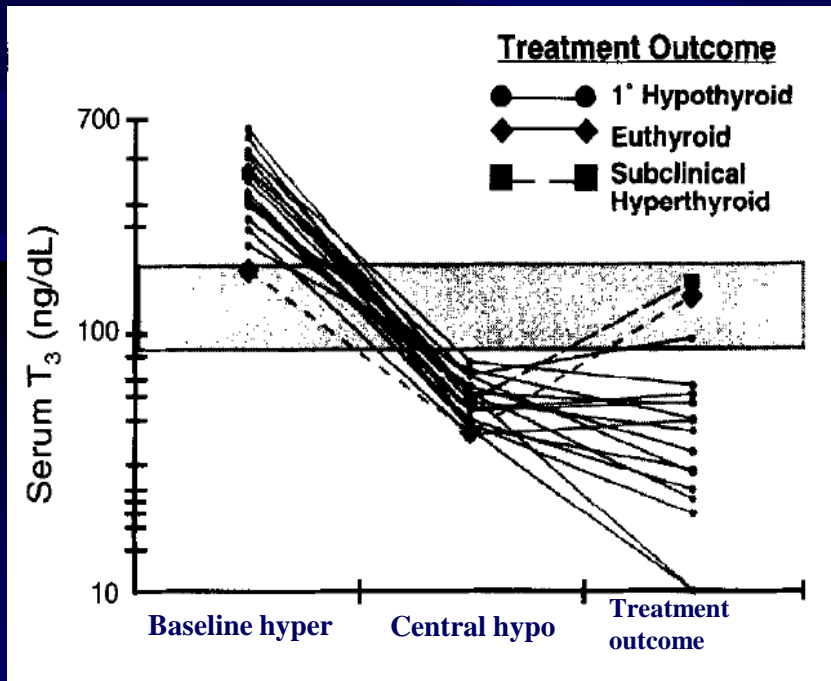
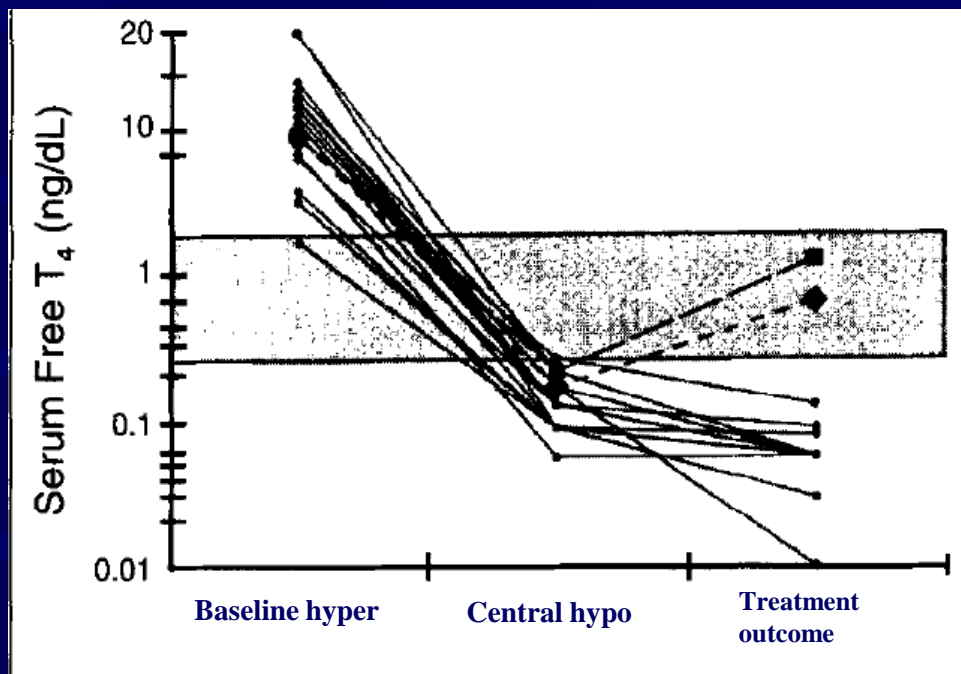
‡Treated with a short course of thionomides before <sup>131</sup>I therapy.

§Received maximum outpatient dose of <sup>131</sup>I as retreatment.

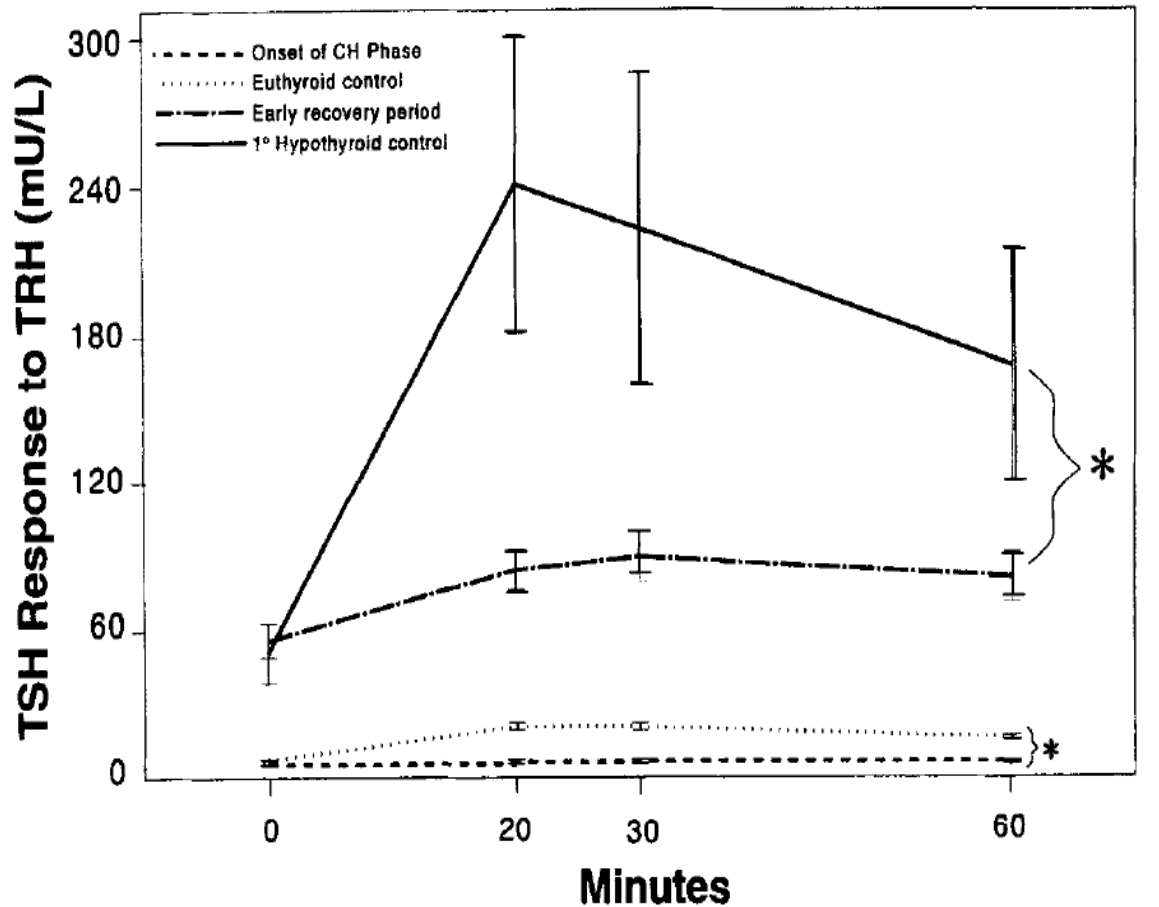
¶Total thyroxine = 3.9 µg/dL (normal range 4.5–12.0).

CH = central hypothyroid; TSH = thyroid-stimulating hormone; NA = not available.

## Pattern of recovery of HPT Axis



## Response to TRH stim test



**Figure 3.** Thyrotropin-releasing hormone (TRH) test results. Thyroid-stimulating hormone (TSH) response following 500  $\mu$ g intravenous TRH injection in 18 patients who experienced a central hypothyroid phase (CH), and in 29 euthyroid and 3 hypothyroid control patients. At onset of CH (---), patients exhibited a blunted TSH response (\* $P < 0.01$ , by repeated-measures analysis of variance using the natural logarithms of the TSH value) compared with euthyroid control patients ( $n = 29$ , ●●●). Similar blunted TSH response (\* $P < 0.01$ ) was observed during the early period after onset of 1° hypothyroidism ( $n = 12$ , -•-•-) in patients compared to control hypothyroid subjects with similar basal TSH elevations ( $n = 3$ , —).

# Results

- ✓ Nineteen (**90%**) of the patients with Graves' disease experienced a transient central hypothyroid phase.
- ✓ Recovery of the HPT axis occurred a mean of **24.7 +- 2.4** days (range 14 to 47) after the onset of central hypothyroidism.
- ✓ Central hypothyroid phase occurred a mean of **62.8 + 5.1** days following 131-I treatment.
- ✓ **Blunted TSH response** to TRH compared to 29 euthyroid control subjects, suggesting primary feedback suppression at the level of the pituitary thyrotrophs
- ✓ **\*\*\*\*** The length of preexisting hyperthyroidism, baseline free T4, and administered dose of I-131 failed to predict the duration of the central hypothyroid phase, although a higher dose of I-131 was associated with an earlier onset of central hypothyroidism ( $P < 0.05$ )

# Summary

- Clinicians should be aware of the delay in the recovery of the HPT axis that occurs after treatment of patients with hyperthyroidism and is manifested by a transient central hypothyroid phase.
- The blunted TSH response to TRH stimulation during this period suggests that suppression occurs primarily at the level of the **pituitary thyrotrophs**
- The length of preexisting hyperthyroidism, and baseline free T4 failed to predict the duration of the central hypothyroid phase



# Back to my patient

## Surgical pathology:

- Invasive squamous cell carcinoma (3.8 cm), moderately differentiated
- The **1.9 cm** microfollicular thyroid nodule, unencapsulated but well circumscribed and demonstrates microfollicular architecture with stromal hyalinization, calcification, and associated random endocrine atypia. These findings are not diagnostic of carcinoma
- papillary thyroid microcarcinoma measures <0.1 cm, is unencapsulated, and shows no evidence of lymphovascular invasion, perineural invasion, or extrathyroidal extension. Margins are uninvolved

- ✓ **Patient was complaining of worsening fatigue**
- ✓ **started on Levothyroxine 50 mcg po daily**
- ✓ **Will see me in 4 weeks in the clinic → repeat TFT**

# References

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**Thank you**

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