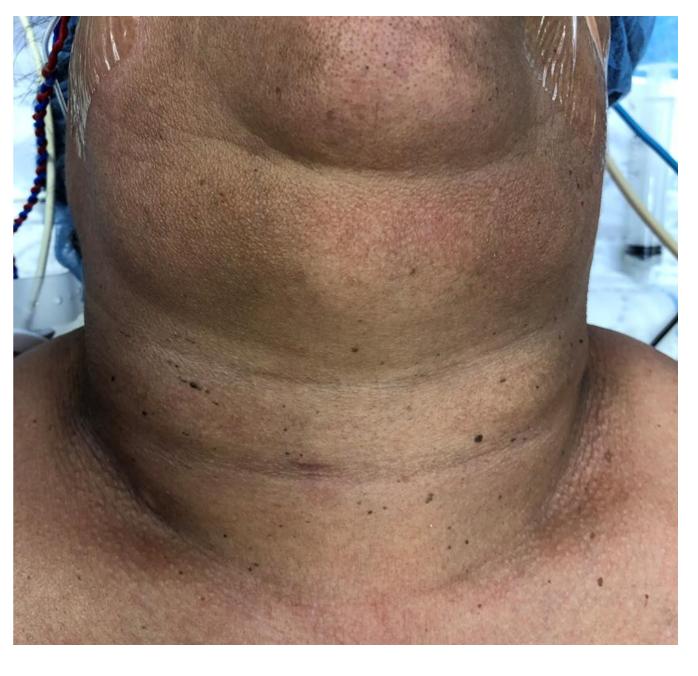
Asymptomatic Thyroid Nodules

John C. McAuliffe, M.D., Ph.D., F.A.C.S. (surgical oncology and endocrine) Montefiore Medical Center Albert Einstein College of Medicine December 13, 2019











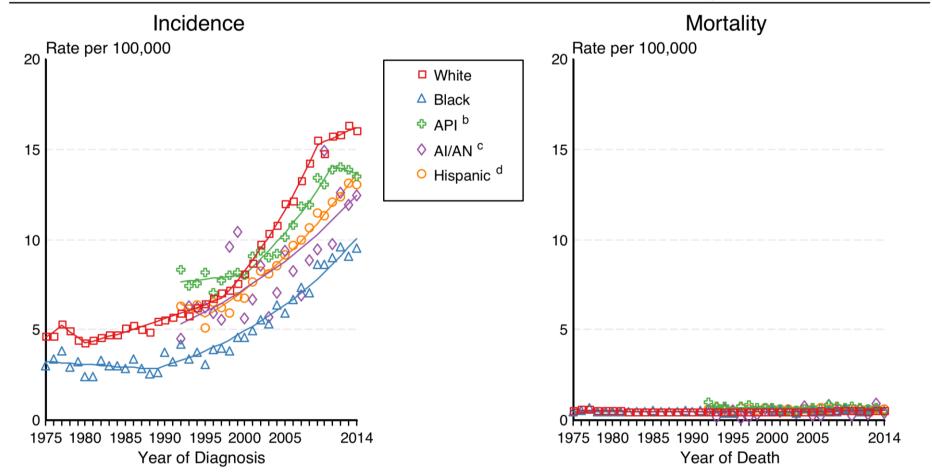






SEER Incidence and US Death Rates^a Cancer of the Thyroid, Both Sexes

Joinpoint Analyses for Whites and Blacks from 1975-2014 and for Asian/Pacific Islanders, American Indians/Alaska Natives and Hispanics from 1992-2014







Scope

- Incidence of Thyroid Cancer Rising due to Detection
- 2015 ATA Guidelines
- Bethesda and TIRADS
- Molecular Testing
- Surgical Interventions





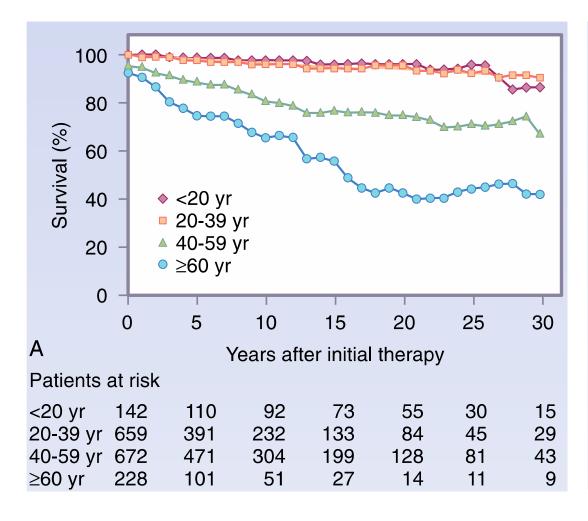
Incidence

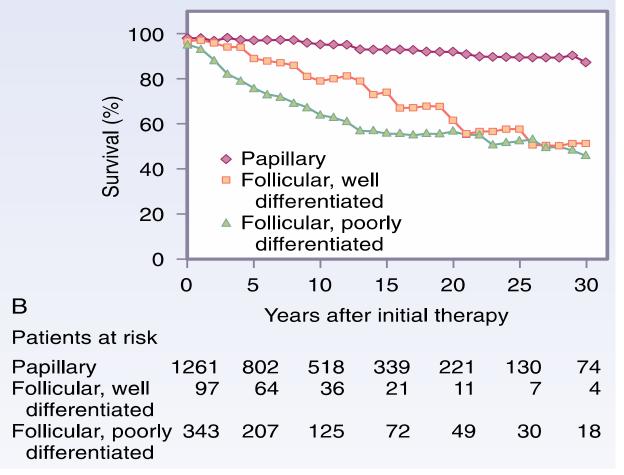
- Increased Detection:
 - Palpable Nodules:
 - 5% in women, 1% in men
 - » iodine-sufficient parts of world
 - Sonographic Nodules:
 - 19-68% of randomly selected individuals
 - Thyroid Cancers <1cm:
 - 25% of new thyroid cancers 1988-1989
 - 39% of new thyroid cancers 2008-2009
 - Thyroid cancer in autopsy specimens approx 11%
- Tripled from 1975 to 2009:
 - -4.9 to 14.3 per 100,000¹
 - 37,200 new cases of thyroid cancer in 2009
 - 63,000 new cases of thyroid cancer 2014





Thyroid Cancer Outcomes



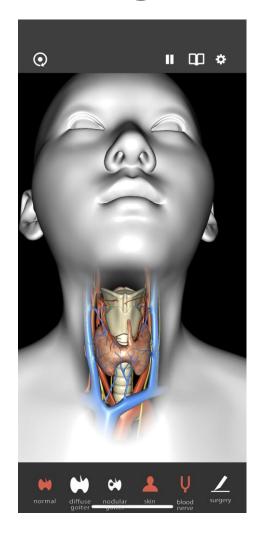


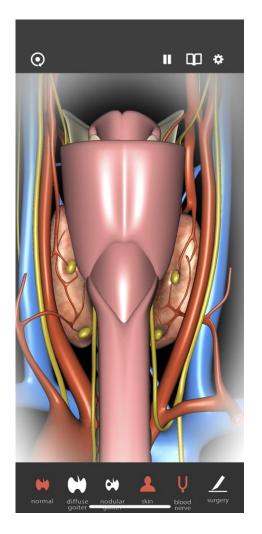
Schlumberger ML. NEJM 1998

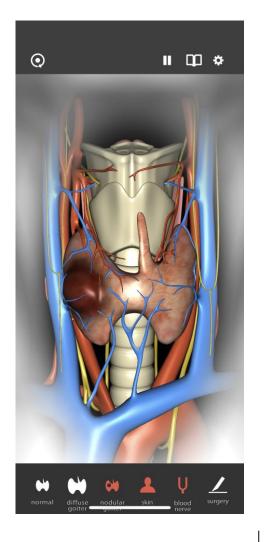




Thyroid Navigator App





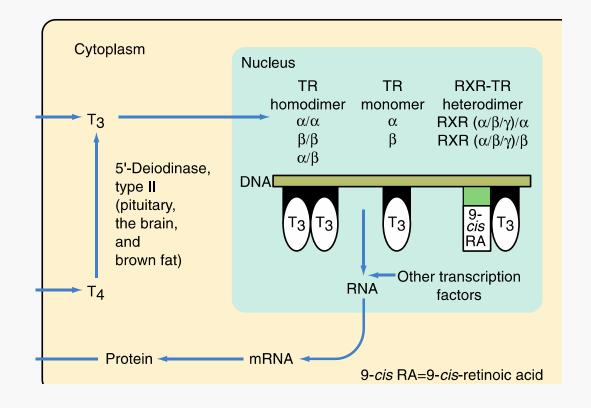


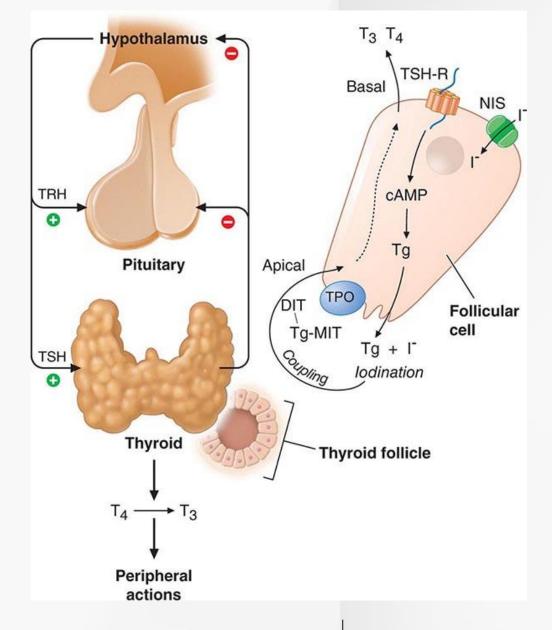




Work Up of Thyroid Nodules

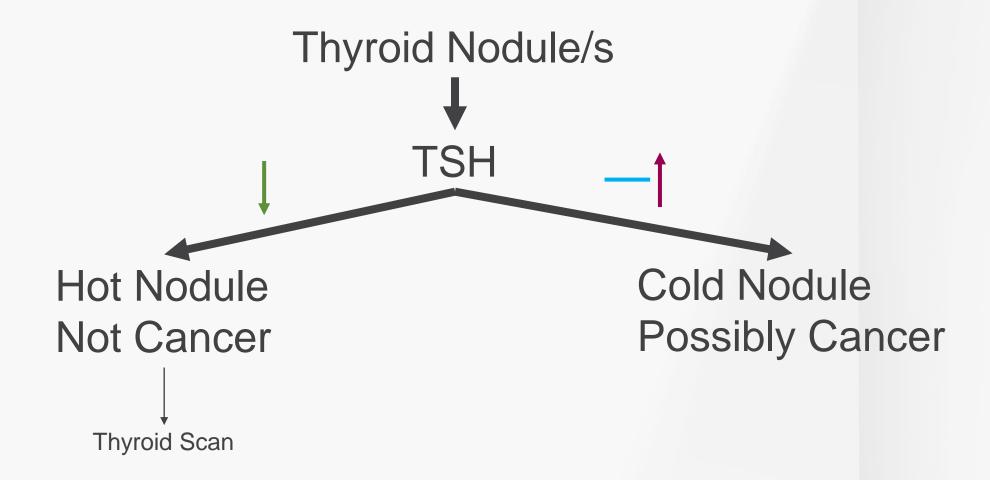
- Determine if Hot or Cold
 - TSH level















THYROID

Volume 26, Number 1, 2016

© American Thyroid Association

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DOI: 10.1089/thy.2015.0020

2015 American Thyroid Association Management Guidelines for Adult Patients with Thyroid Nodules and Differentiated Thyroid Cancer

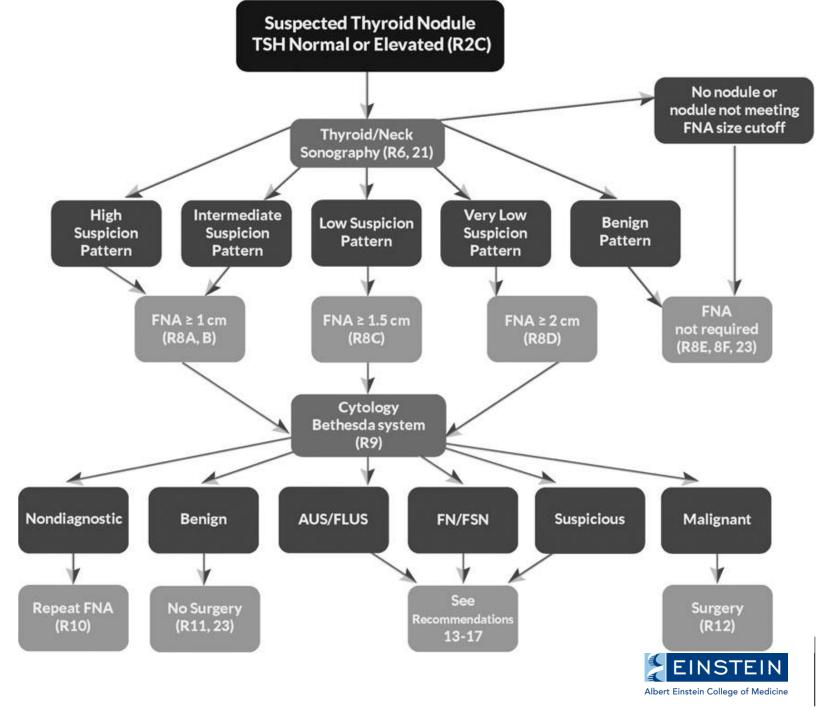
The American Thyroid Association Guidelines Task Force on Thyroid Nodules and Differentiated Thyroid Cancer

Bryan R. Haugen,^{1,*} Erik K. Alexander,² Keith C. Bible,³ Gerard M. Doherty,⁴ Susan J. Mandel,⁵ Yuri E. Nikiforov,⁶ Furio Pacini,⁷ Gregory W. Randolph,⁸ Anna M. Sawka,⁹ Martin Schlumberger,¹⁰ Kathryn G. Schuff,¹¹ Steven I. Sherman,¹² Julie Ann Sosa,¹³ David L. Steward,¹⁴ R. Michael Tuttle,¹⁵ and Leonard Wartofsky¹⁶



SPECIAL ARTICLE







TIRADS

TIRADS criteria	Nodule description
Composition	Cystic (0)
	Spongiform (0)
	Mixed cystic and solid (1)
	Solid (2)
Echogenicity	Anechoic (0)
	Hyperechoic or isoechoic (1)
	Hypoechoic (2)
	Very hypoechoic (3)
Shape	Wider-than-tall (0)
	Taller-than-wide (3)
Margin	Smooth (0)
	Ill-defined (0)
	Lobulated or irregular (2)
	Extra-thyroidal extension (3)
Echogenic foci	None or large comet-tail artifacts (0)
	Macrocalcifications (1)
	Peripheral (rim) calcification (2)
	Punctate echogenic foci (3)





TIRADS classification (points)	Description	Management
TIRADS 1 (0)	Benign	No FNA
TIRADS 2 (2)	Not suspicious	No FNA
TIRADS 3 (3)	Mildly suspicious	FNA if \geq 2.5 cm; active surveillance if \geq 1.5 cm
TIRADS 4 (4–6)	Moderately suspicious	FNA if \geq 1.5 cm; active surveillance if \geq 1 cm
TIRADS 5 (≥7)	Highly suspicious	FNA if \geq 1.0 cm; active surveillance if \geq 0.5 cm





High Suspicion >70-90% hypoechoic, Interrupted rim calcification with soft tissue extrusion nodule with irregular margins, hypoechoic, irregular margins, extrathyroidal extension hypoechoic irregular margins microcalcifications hypoechoic nodule irregular margin hypoechoic taller than wide suspicious left lateral lymph node Intermediate Suspicion 10-20% hypoechoic sol regular mergin hypoechoic solid regular margin malignancy Low Suspicion 5-10% isoechoic solid regular margin partially cystic with eccentric solid area partially cystic with eccentric solid areas hyperechoic solid regular margin of Very low Risk Suspicion <3% partially cystic no suspicious features partially cystic no suspicious features spongiform Benign <1%





Table 6. Sonographic Patterns, Estimated Risk of Malignancy, and Fine-Needle Aspiration Guidance for Thyroid Nodules

Sonographic pattern	US features	Estimated risk of malignancy, %	FNA size cutoff (largest dimension)		
High suspicion	Solid hypoechoic nodule or solid hypoechoic component of a partially cystic nodule with one or more of the following features: irregular margins (infiltrative, microlobulated), microcalcifications, taller than wide shape, rim calcifications with small extrusive soft tissue component, evidence of ETE	>70–90 ^a	Recommend FNA at ≥1 cm		
Intermediate suspicion	Hypoechoic solid nodule with smooth mar- gins <i>without</i> microcalcifications, ETE, or taller than wide shape	10–20	Recommend FNA at ≥1 cm		
Low suspicion	Isoechoic or hyperechoic solid nodule, or partially cystic nodule with eccentric solid areas, <i>without</i> microcalcification, irregular margin or ETE, or taller than wide shape.	5–10	Recommend FNA at ≥1.5 cm		
Very low suspicion	Spongiform or partially cystic nodules with- out any of the sonographic features de- scribed in low, intermediate, or high suspicion patterns	<3	Consider FNA at ≥2 cm Observation without FNA is also a reasonable option		
Benign	Purely cystic nodules (no solid component)	<1	No biopsy ^b		





Table 8. The Bethesda System for Reporting Thyroid Cytopathology: Diagnostic Categories and Risk of Malignancy^a

Diagnostic category	Estimated/predicted risk of malignancy by the Bethesda system, % ^a	Actual risk of malignancy in nodules surgically excised, % median (range) ^b		
Nondiagnostic or unsatisfactory	1–4	20 (9–32)		
Benign	0–3	2.5 (1–10)		
Atypia of undetermined significance or follicular lesion of undetermined significance	5–15	14 (6–48)		
Follicular neoplasm or suspicious for a follicular neoplasm	15–30	25 (14–34)		
Suspicious for malignancy Malignant	60–75 97–99	70 (53–97) 99 (94–100)		





FNA is main stay

Endocrine (2019) 65:365–370 https://doi.org/10.1007/s12020-019-01973-2

ORIGINAL ARTICLE

Thyroid core needle biopsy: patients' pain and satisfaction compared to fine needle aspiration

Hyo Jin Kim¹ · Yeo Koon Kim (o² · Jae Hoon Moon³ · June Young Choi⁴ · Sang Il Choi²

Received: 27 March 2019 / Accepted: 27 May 2019 / Published online: 15 June 2019 © Springer Science+Business Media, LLC, part of Springer Nature 2019





Medical therapy of benign thyroid nodules

None





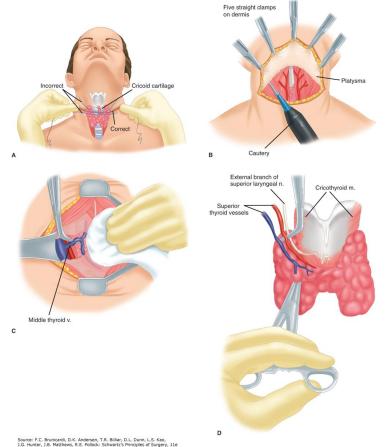
Malignant Management

- Improve OS and DSS. Reduce persistent/recurrent disease. Accurate staging. Minimize morbidity.
- Facilitate adjuvant therapy
- Malignancy > 4cm = total thyroidectomy +/- selective lymph node dissection
- Malignancy >1cm and <4cm without extrathyroidal extension or LN+ disease = either lobectomy or total thyroidectomy
- Malignancy >1cm and <4cm with contralateral benign nodules = variable
- Malignancy <1cm without extrathyroidal extension or LN+ = lobectomy or active surveillance





How I do it.





- Incision in a skin crease, SCM to SCM
- Rarely transect the strap muscles.
- Best with a second assist to retract muscles





Post Operative Complications

Complication	%
Vocal cord paralysis	
Transient	2.5
Permanent	0.3
Prolonged postoperative hypocalcemia (>7 days)	3.7, 0.6
Post operative bleeding	< 1

If parathyroid resected or in jeopardy, I autotransplant in left SCM marked with hemoclip I send all patients home same day if PACU PTH is > 9.

Pts sent home on Calcium Carbonate (500mg QID) and Calcitriol (.25ug BID)

Redo Surgery or Dysphonia = Fiberoptic Laryngoscopy





Drains and Thyroidectomy

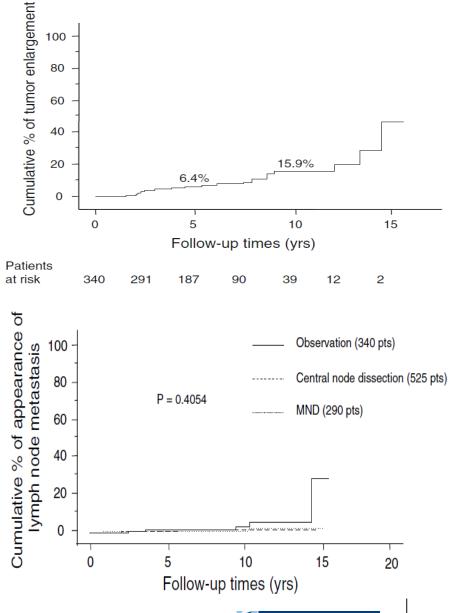
- Data doesn't support routine use
- No significant difference in
 - Re-op rates
 - Respiratory distress
 - Wound infections
 - Post op fluid collections
 - Longer LOS

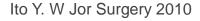




Malignant Management

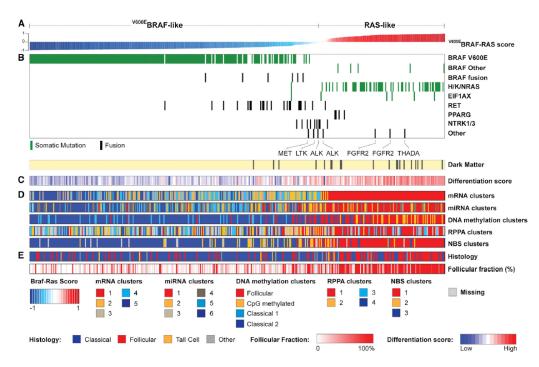
- Disease Specific
 Mortality <1%
- Papillary TC <1cm
- Dx by US FNA
- 340 observed, 1,055 surgery
- Mean f/u 74 months



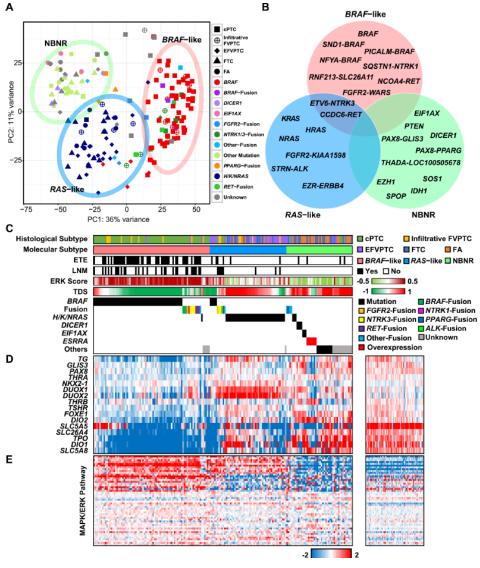




Molecular Testing



Cancer Genome Atlas Research Network, Cell. 2014



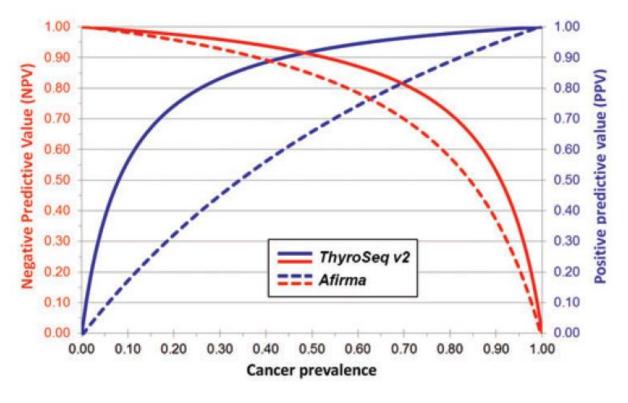
Yoo SK, et. al. PLOS 2016





ThyroSeq

- ThyroSeq 2.1 "rule in" and "rule out" test
 - PPV 91%, NPV 92%
 - 14 genes
 - 42 gene fusions
 - 8 genes assessed for cell composition of FNA sample
 - follicular cells vs parathyroid cells vs parafollicular cells (C-cells)



Nikiforov YE Thyroid 2015





AUS/FLUS: Bethesda III

Nikiforov et al., 2015 ThyroSeq v2

- PPV 77%
- NPV 97%

Alexander et al., 2012

- Afirma
 - PPV 38%
 - NPV 95%

FN/SFN: Bethesda IV

Nikiforov et al., 2014

ThyroSeq v2

- PPV 83%
- NPV 96%

Alexander et al., 2012

Afirma

- PPV 37%
- NPV 94%





Accuracy of FNA for Large Nodules

TABLE 3: Diagnostic indices of fine-needle aspiration cytology (FNAC) in the five thyroid nodule size categories.

Group	A (≤0.5 cm)	B (>0.5–1 cm)	C (>1-2 cm)	D (>2-4 cm)	E (>4 cm)	p value
Sensitivity	96.8	94.9	98.7	86.7	50	0.006
Specificity	100	93.9	100	100	100	0.575
PPV	100	98.8	100	100	100	0.745
NPV	76.9	77.5	97	92	85.7	0.076
Diagnostic accuracy	96.8	94.8	99	94.7	87.5	

PPV: positive predictive value; NPV: negative predictive value.





False Negatives and TIRADS for >3cm Benign Nodules

Table 3. Diagnostic performance according the TIRADS category of all cases and surgery cases.

TIRADS	All cases (n = 632)					Surgery cases (n = 164)						
category	Nodules recommended for surgery *	TP	FP	FN	SEN (%,95%CI)	PPV (%,95%CI)	Nodules underwent surgery	TP	FP	FN	SEN (%,95%CI)	PPV (%,95%CI)
3,4a,4b,4c	632 (100)	23	609	0 (0)	100 (85.2–100)	3.6 (2.3–5.4)	164 (100)	22	142	0 (0)	100 (84.6–100)	13.4 (8.6–19.6)
4a,4b,4c	313 (49.5)	20	293	3 (13.0)	87.0 (66.4– 97.2)	6.4 (5.4–7.5)	86 (52.4)	19	67	3 (13.6)	86.4 (65.1– 97.1)	22.1 (18.2– 26.5)
4b,4c	97 (15.3)	10	87	13 (56.5)	43.5 (23.2– 65.5)	10.3 (6.5–16.0)	33 (20.1)	9	24	13 (59.1)	40.9 (20.7– 63.7)	27.3 (16.8– 41.1)
4c	17 (2.7)	2	15	21 (91.3)	8.7 (1.1–28.0)	11.8 (3.2–35.5)	2 (1.2)	1	1	21 (95.4)	4.6 (0.1–22.8)	50 (6.1–93.9)

Numbers in parentheses are percentages and 95% confidence interval

TP: true-positive, FP: false-positive, FN: false-negative, SEN: sensitivity, PPV: positive predictive value





^{*}Number of surgeries that would have been performed had the TIRADS category been applied to the overall sample of nodules as cytology-image discordance

FDG Avid Nodules?

THYROID Volume 22, Number 9, 2012 © Mary Ann Liebert, Inc.

DOI: 10.1089/thy.2012.0005

THYROID RADIOLOGY AND NUCLEAR MEDICINE

Risk of Malignancy in Thyroid Incidentalomas Detected by ¹⁸F-Fluorodeoxyglucose Positron Emission Tomography: A Systematic Review

Kerstin Kathrine Soelberg, Steen Joop Bonnema, Thomas Heiberg Brix, and Laszlo Hegedüs

- ≅35% of ¹⁸FDG-PET positive nodules are cancerous.
- >1cm FNA indicated.





Conclusions

- DTC has great outcomes if treated well
- TSH and US are key 1st steps
- TIRADS and Bethesda Criteria frame management
- Genetics in atypical nodules should be considered
- Minimalist approach to surgery is standard with significant evidence for equivalent outcomes compared to more surgery
- FDG Avidity is a risk factor for cancer
- Surgical intervention for large nodules appropriate due to decreased accuracy of FNA





Montefiore DOING MORE