4D-CT Parathyroid for the General Radiologist: A Pictorial Essay of Illustrative Cases

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Educational Objectives

- Describe the purpose of 4D-CT for evaluation of primary hyperparathyroidism and pre-surgical planning
- Review 4D-CT imaging protocol and illustrate the typical appearance of parathyroid adenomas
- Review the embryology of the parathyroid glands, and use it to facilitate a systematic search for embryologic and ectopic adenomas
- Illustrate a variety of cases of parathyroid adenomas in embryologic and ectopic locations and cases of multi-glandular disease
- Illustrate important mimics of parathyroid disease and important "incidental" findings in the neck and upper chest





Hyperparathyroidism

- Increased parathyroid hormone causes increased osteoclastic activity
- Subtypes
 - Primary
 - Parathyroid adenoma
 - Parathyroid hyperplasia
 - Parathyroid carcinoma
 - Secondary
 - Caused by chronic hypocalcemia
 - Most often due to renal osteodystrophy, vitamin D deficiency or malnutrition
 - Leads to increased parathyroid hormone levels
 - Tertiary
 - Autonomous adenoma caused by chronic overstimulation of hyperplastic glands, usually in the setting of renal insufficiency





Hyperparathyroidism

- 3 to 4 times more common in women than men
- 50-70 most common age range
- Majority of cases of <u>primary hyperparathyroidism</u> are caused by adenomas, (~90%)
- Less commonly there are multiple adenomas (~5%), multigland hyperplasia (~5%), or parathyroid carcinoma (<1%)
- Approximately 20% of adenomas are in ectopic locations (numbers vary in literature)
- Surgical excision of adenoma is treatment of choice in primary hyperparathyroidism





Hyperparathyroidism

- Historically, multigland exploration/excision was performed without pre-operative imaging
- In the early 2000s, minimally invasive resection emerged, and preop planning importance rose
 - Single adenoma resected through small incision; remaining glands left unexplored if PTH level drops
- Radiologist role is to:
 - Describe/localize high suspicion nodules
 - Identify ectopic adenomas
 - Recognize multigland hyperplasia or multiple adenomas
 - Highlight any relevant anatomical considerations





Imaging Technique

- Ultrasound
 - Inexpensive, fast, can be done by surgeons in office to localize typical adenomas greater than 1 cm.
- Dual phase Tc-99m Sestamibi scan
 - Early images 10-20 min post injection (activity in both thyroid & parathyroid)
 - Delayed (90 mins), activity in thyroid washes out, persists in adenoma
 - Can do SPECT/CT for better localization
- 4D CT parathyroid
 - Improved sensitivity of localization, particularly in ectopic/mediastinal locations
 - Helpful for pre-operative planning





CT parathyroid technique

- "4D" axial, sagittal, coronal, and dynamic enhancement characteristics
- Precontrast
- Arterial phase (25-30 seconds after injection of contrast commences)
- Venous phase (60-80 seconds after injection of contrast commences)
- At UCLA, our arterial phase extends from skull base to carina to help identify ectopic locations
- All three phases and planes should be evaluated; however the arterial phase is usually the most helpful



Anatomy

- Embryology is relevant to parathyroid adenomas
- Inferior glands (paired) arise from the 3rd branchial pouch
- Superior glands (paired) arise from the 4th branchial pouch
- These pouches are approximately at level of carotid bifurcation
- Parathyroid glands descend during development
- Inferior glands migrate with the thymus, and can be found from carotid bifurcation to carina, including within the thymus or thyroid gland
- Superior glands migrate with thyroid gland (less variable location)





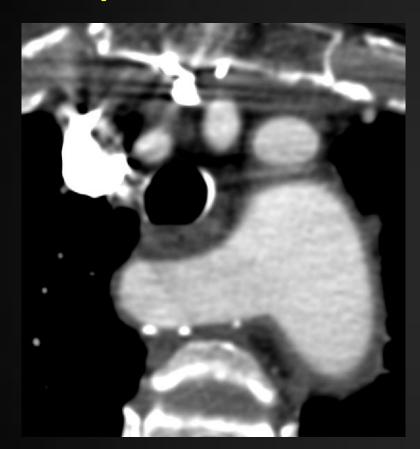
Anatomy

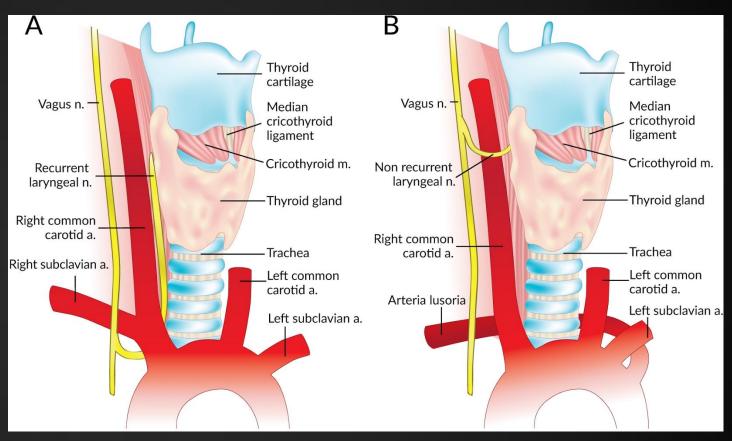
- Embryologic superior glands are located posterior to the upper third of the thyroid gland, often paraesophageal or within the tracheoesophageal groove
- Embryologic superior glands can fall caudally (descended) and can be found as low as within the mediastinum (typically posterior in location)
- Embryologic inferior glands are usually closely associated with the inferior thyroid (immediately inferior, posterior, lateral or anterior to the tip of the gland).
- Embryologic inferior glands are located more anteriorly (paratracheal, anterior mediastinum, closely associated with strap musculature)





Important to mention: Aberrant SCA





Aberrant SCA associated w/ nonrecurrent laryngeal nerve (not in TE groove)!





Enhancement Characteristics

- Typical parathyroid adenomas are:
 - Hypoattenuating to thyroid on precontrast
 - Arterially hyperenhancing
 - Rapid washout on venous phase
- Precontrast sequence is very important to differentiate thyroid tissue from parathyroid
- Start with arterial phase to search for possible adenoma, and then confirm on precontrast and delayed phase
- DDx of possible adenomas include thyroid (use precontrast to distinguish), lymph nodes (typically progressively enhance on venous phase, do not wash out)





Enhancement Characteristics

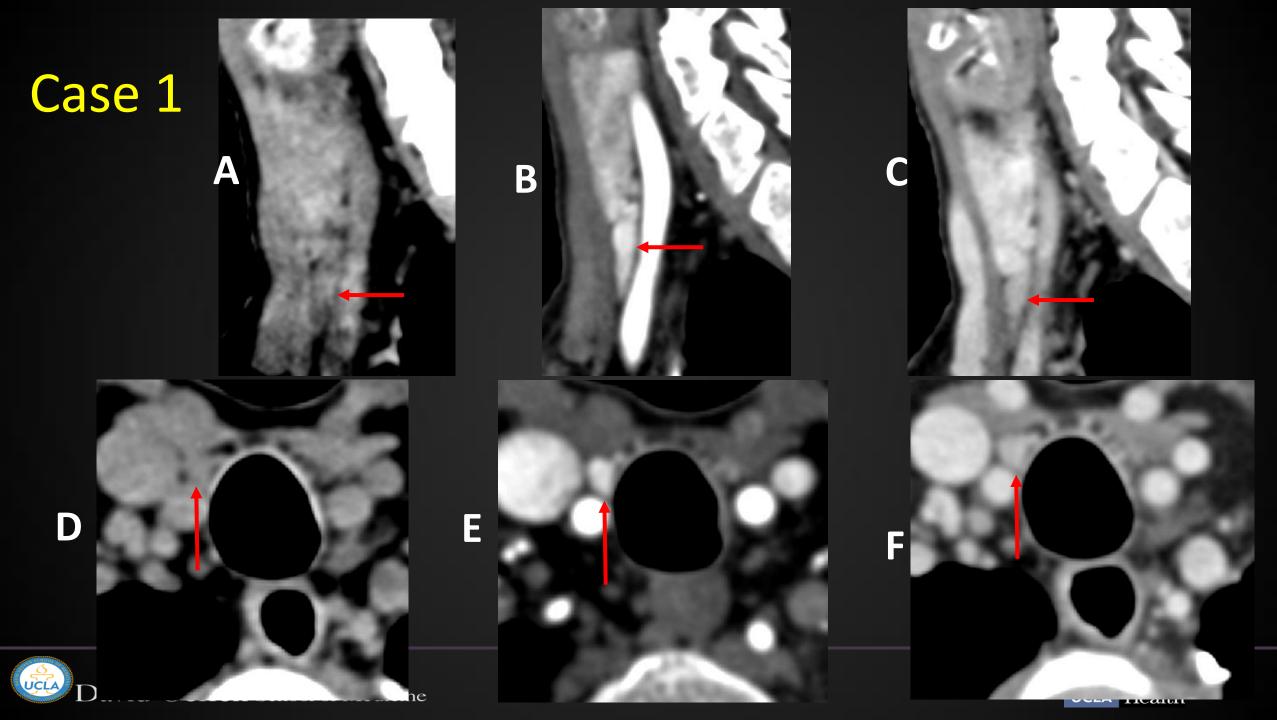
Caveats:

- In approximately one third of cases an adenoma will be isodense to thyroid on arterial and venous phases
- Patients with hypothyroidism demonstrate a lower intrinsic attenuation of the thyroid gland limiting the usefulness of the pre-contrast scan (these patients do not organify iodine as those with normally functioning thyroid glands)
- Cystic degeneration of an adenoma, streak artifact, suboptimal bolus timing, thyroiditis can alter enhancement characteristics



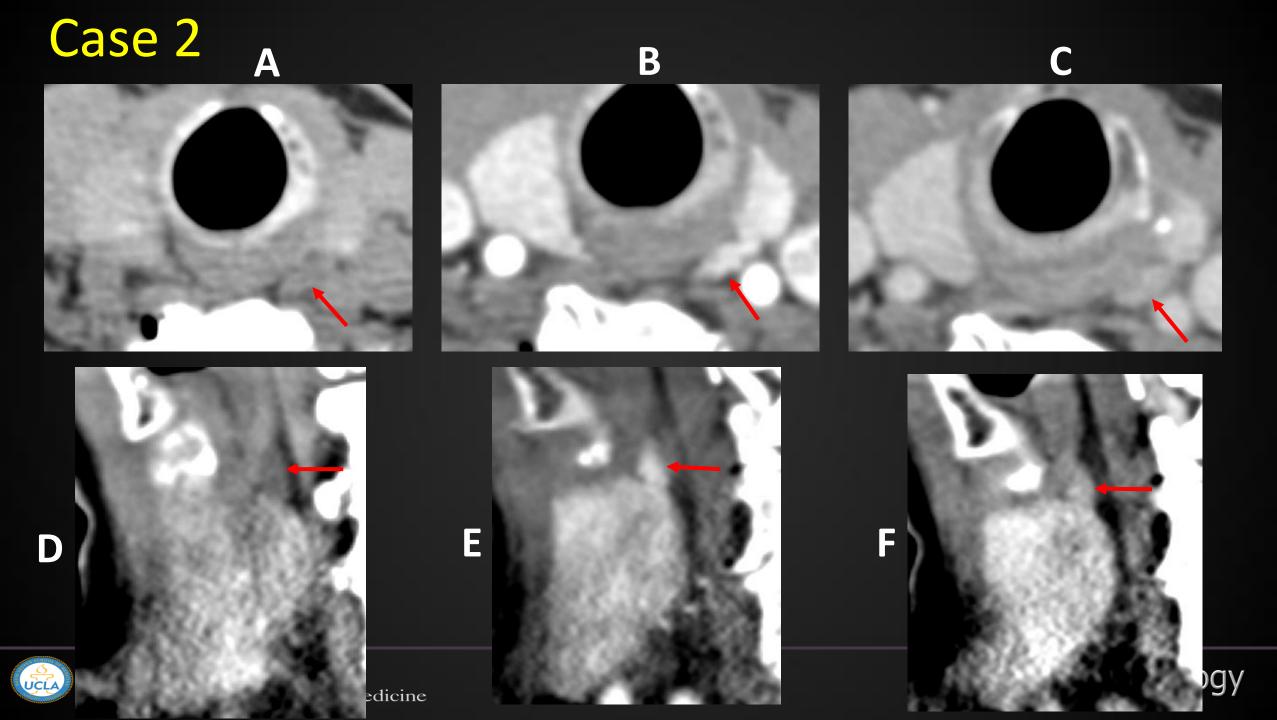
CASES





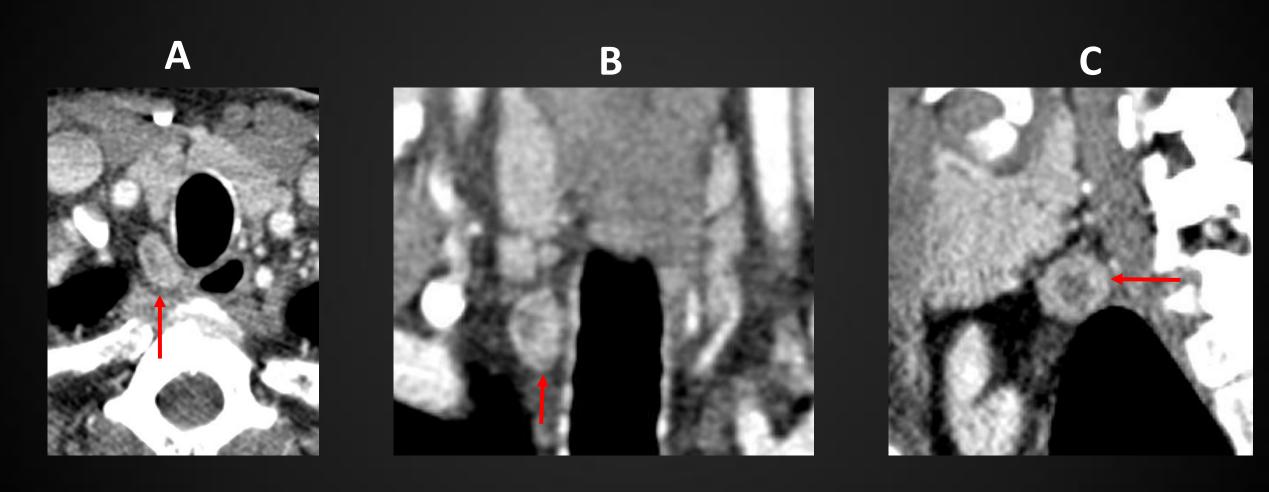
- Right sided embryologic inferior parathyroid adenoma
- Along inferior margin of right lobe of thyroid
- Paratracheal, closely associated with the overlying strap musculature
- Hypodense to thyroid on precontrast (A, D), arterially enhancing (B, E), and demonstrates washout on venous phase (C, F)





- Embryologic superior left parathyroid adenoma
- Paraesophageal, posterior to the superior aspect of the thyroid lobe
- Hypodense to thyroid on precontrast (A, D), arterially enhancing (B, E), demonstrates washout on venous phase (C, F)







- Right sided descended embryologic superior gland adenoma
- Parathyroid adenoma in right tracheoesophageal groove, along inferior aspect of the right thyroid lobe
- Arterial enhancement on axial (A), coronal (B), and sagittal (C) images with internal cystic change



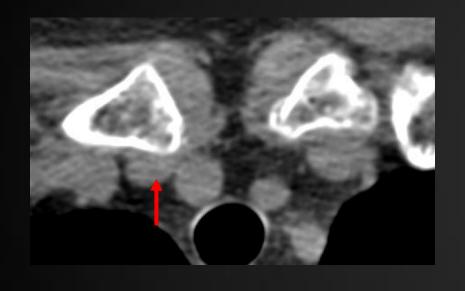
Case 4 B A D



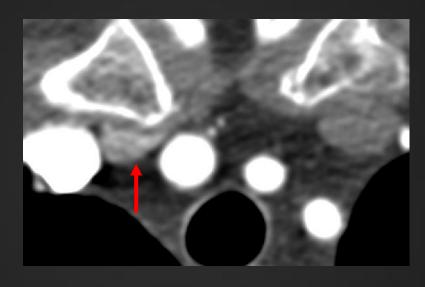


- Bilateral embryologic superior gland parathyroid adenomas
- Arterial enhancement on axial (A), coronal (B), and off-midline sagittal (C, D) images

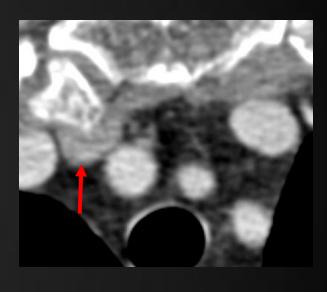








B



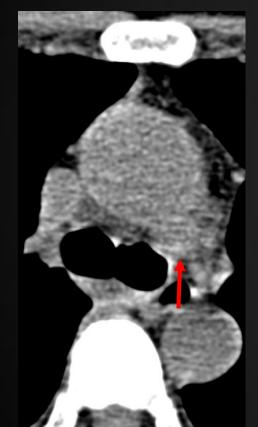
C



- Right sided mediastinal / intrathymic parathyroid adenoma
 - Along posterior margin of the right clavicular head
- Hypodense to thyroid on precontrast (A, D), arterial enhancement with cystic change (B, E), and venous washout (C, F)
- Can be easy to overlook on axial alone (A-C)
- Best identified on coronal reformats (D-F)



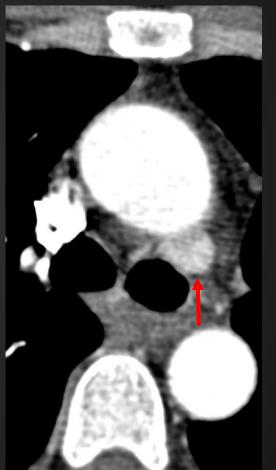
A



B



C

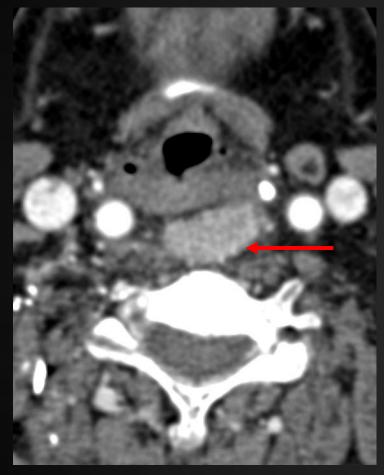


D



- Mediastinal (AP window) parathyroid adenoma
- Nodule in the AP window that is hypodense to thyroid on precontrast (A), arterially enhancing (B, C), and demonstrates venous washout (D)
- Remember to look all the way down to the AP window for mediastinal adenomas
- The inferior parathyroid gland migrates with the thymus and can be seen as low as the AP window / carinal level





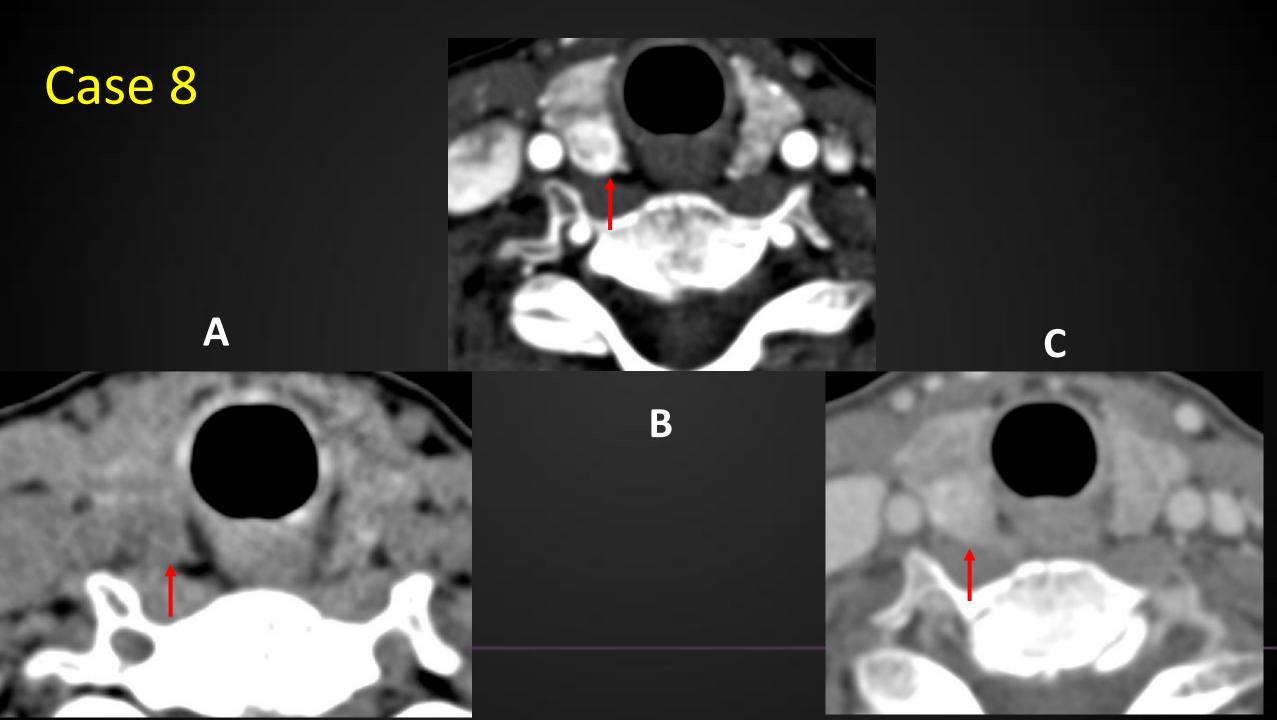




A

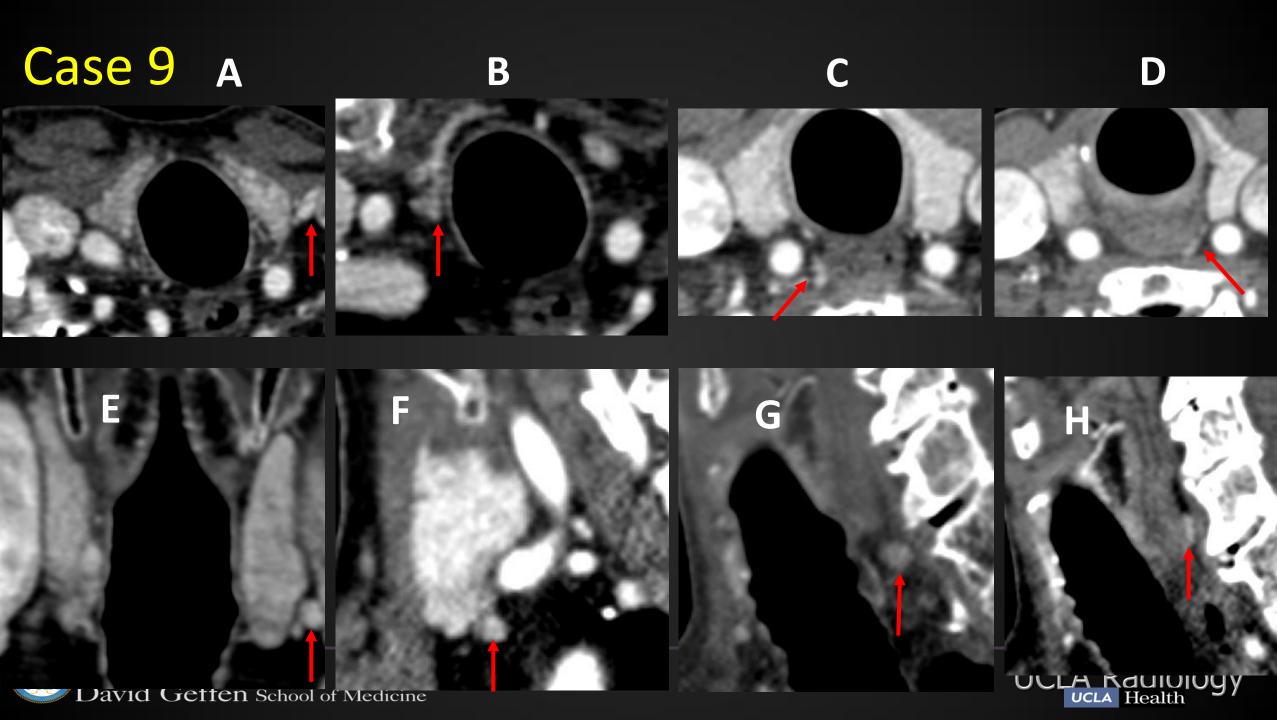
- Ectopic retropharyngeal parathyroid adenoma at the level of the supraglottic larynx
- Large hyperenhancing mass is seen on axial (A), sagittal (B) and coronal (C) aterial phase images





- Right sided intrathyroidal subcapsular parathyroid adenoma
- Nodule along the posterior subcapsular aspect of the right thyroid lobe is hypodense on precontrast (A), hyperenhancing on arterial phase(B), and washes out on delayed phase (C)
- Note background heterogeneity of the thyroid gland





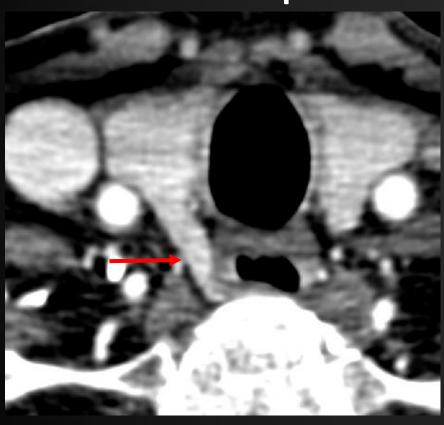
- Multigland hyperplasia
- All four embryologic glands are mildly enlarged:
 - Left inferior gland hyperplasia on axial (A) and coronal (E) images
 - Deep to the strap muscles, along inferior aspect of the left thyroid lobe
 - Right inferior gland hyperplasia on axial (B) and sagittal (F) images
 - Paratracheal, along inferior aspect of right thyroid lobe
 - Right superior gland hyperplasia on axial (C) and sagittal (G) images
 - Paraesophageal, along superior aspect of right thyroid lobe
 - Left superior gland hyperplasia on axial (D) and sagittal (H) images
 - Paraesophageal, along superior aspect of the left thyroid lobe



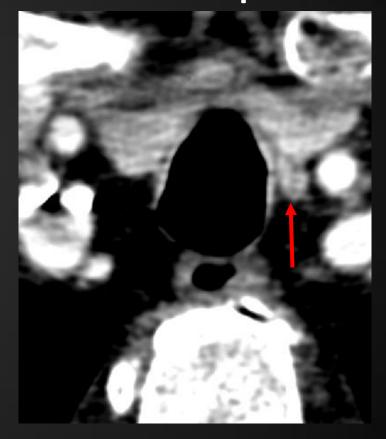


Case 10 – Which image shows the adenoma?

Axial arterial phase

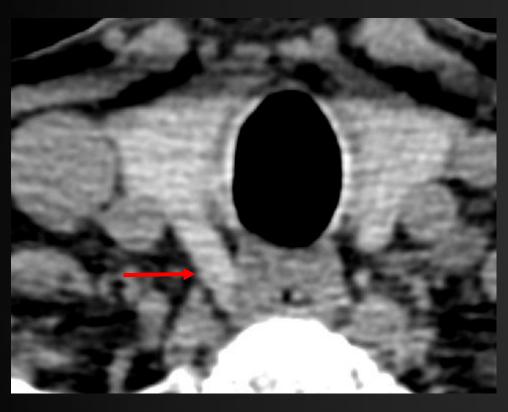


Axial arterial phase





Case 10 – Importance of Precontrast

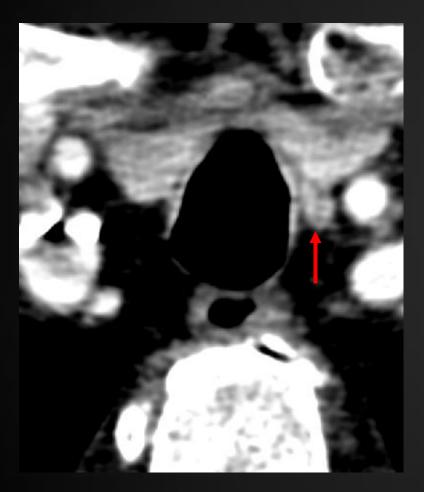


Axial precontrast: Right TE groove nodule isodense to thyroid, thus exophytic thyroid tissue (Zuckercandl tubercle of thyroid)



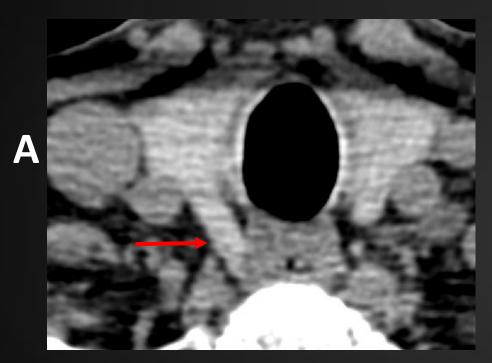
Axial precontrast: Left paratracheal nodule hypodense to thyroid on precontrast, therefore parathyroid adenoma

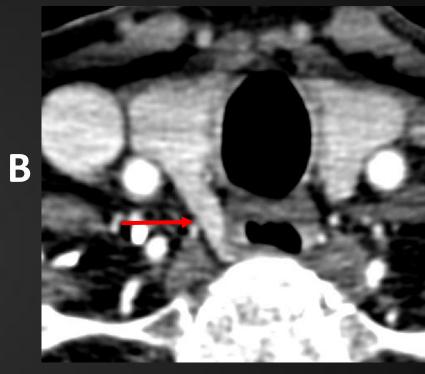




- -Left sided embryologic inferior parathyroid adenoma
- -Seen along the inferior margin of the thyroid lobe
- -Often paratracheal or closely associated with the strap musculature

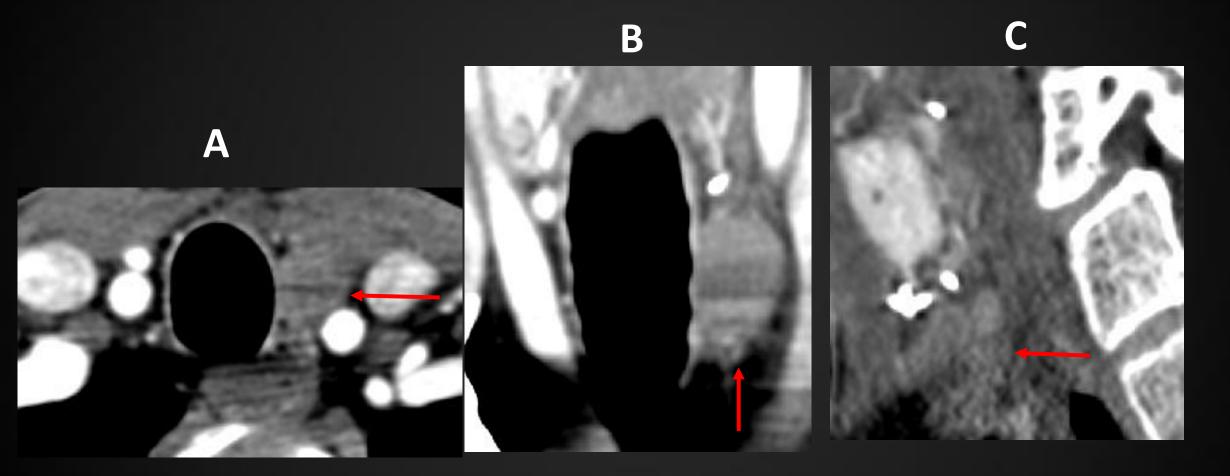






Zuckercandl tubercle. Right sided exophytic thyroid tissue in the tracheoesophageal groove, an anatomic variant characterized by tissue that is isodense to thyroid on precontrast phase (A) and iso-enhancing to thyroid on arterial phase (B)

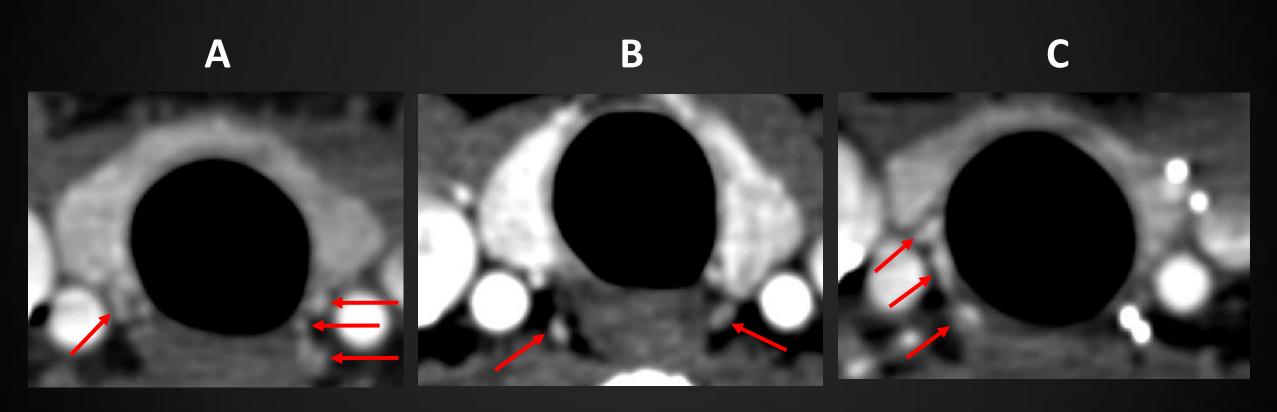




- Parathyroid carcinoma
 - Recurrent primary hyperparathyroidism after parathyroidectomy
- Poorly defined hypoenhancing mass with irregular borders in the parathyroidectomy bed on arterial phase axial (A), coronal (B) and sagittal (C) images
- Surgical clips from prior parathyroidectomy







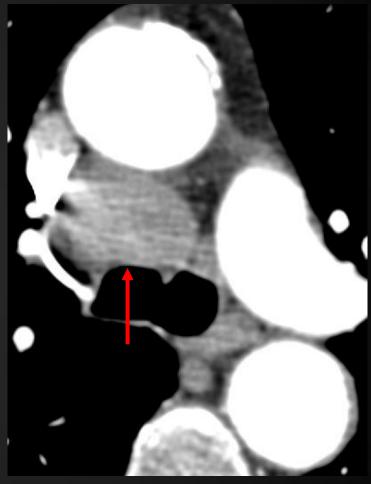


- Parathyromatosis after parathyroidectomy
- Multiple tiny nodules bilaterally in the tracheo-esophageal grooves seen on axial arterial phase images (A-C).
- Surgical clips from prior parathyroidectomy









В

- Biopsy confirmed poorly differentiated carcinoma of unknown primary
- Precarinal mass with only mild enhancement on arterial phase coronal (A), sagittal (B), and axial (C) images



Sestamibi immediate



Sestamibi delay



Case 14 A B



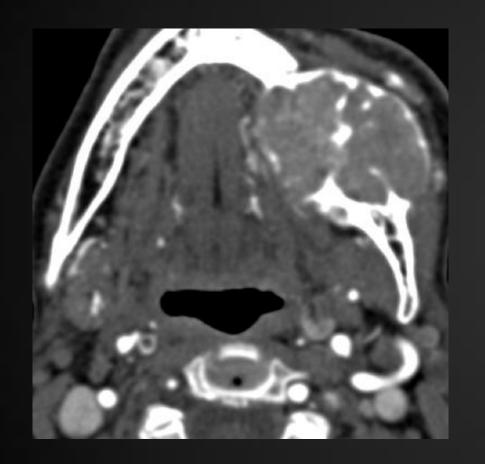


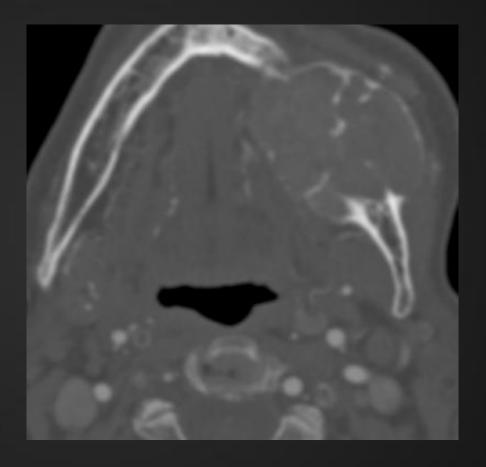
Thymoma

- Large lobulated anterior mediastinal mass
- Can mimic parathyroid adenoma on Sestamibi parathyroid or myocardial perfusion scans
 - Demonstrates increased uptake on the immediate phase Sestamibi with persistent uptake on delayed phase (same as parathyroid adenoma)
- Hypodense to thyroid on precontrast coronal and sagittal images (A, B) and hypoenhancing on arterial phase sagittal (C) images
- Can have areas of calcification and cystic/necrotic change









- Brown tumor of the mandible
- Expansile, lytic mass with internal calcification and internal hypderdense components (representing hemorrhage) in the left mandibular body related to primary hyperparathyroidism



A B

UCLA Radiology

- Commonly encountered incidental findings
- Spiculated left upper lobe mass compatible with pulmonary neoplasm on axial lung window CT (A)
- Superiorly directed left supraclinoid internal carotid artery aneurysm on axial arterial phase (B)
- Proximal internal carotid artery occlusion on sagittal arterial phase (C)



Take Home Points

- Primary Hyperparathyroidism is a relatively common disease associated with significant morbidity.
- Parathyroidectomy is a highly effective and definitive treatment for primary hyperparathyroidisim.
- Accurate preoperative localization is essential in planning for minimally invasive parathyroidectomies.
- 4D CT Parathyroid is emerging as the first line tool for preoperative localization, both for re-operation and initial presentation.
- High quality interpretation of 4D CT examinations should be part of the general radiologist's skillset.





References

- Ruda JM, Hollenbeak CS, Stack BC Jr. A systematic review of the diagnosis and treatment of primary hyperparathyroidism from 1995 to 2003. Otolaryngol Head Neck Surg 2005;132(3):359–72.
- Hoang JK, Sung WK, Bahl M, et al. How to perform parathyroid 4D CT: tips and traps for technique and interpretation. Radiology 2014;270(1):15–24.
- Sackett WR, Barraclough B, Reeve TS, et al. Worldwide trends in the surgical treatment of primary hyperparathyroidism in the era of minimally invasive parathyroidectomy. Arch Surg 2002;137(9):1055–59.
- Rodgers SE, Hunter GJ, Hamberg LM, et al. Improved preoperative planning for directed parathyroidectomy with 4-dimensional computed tomography. Surgery 2006;140(6):932–41.
- Bahl M, Sepahdari AR, Sosa JA, et al. Parathyroid adenomas and hyperplasia on four-dimensional CT scans: three patterns of enhancement relative to the thyroid gland justify a three-phase protocol.
 Radiology 2015;277(2):454-62.
- Sepahdari AR, Bahl M, Harari A, et al. Predictors of multigland disease in primary hyperparathyroidism: a scoring system with 4D-CT imaging and biochemical markers. Am J Neuroradiol 2015;36: 987–92.



