

# **Precision Medicine with 3D Ultrasound**

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## Abstract

*Introduction:* Currently, B-mode ultrasound (US) is the primary imaging modality in diagnosing thyroid nodules (TNs). B-mode is a two-dimensional US (2D US) imaging display. Recent studies suggest a role for strain and shear wave elastography for evaluating TN as well. Three-dimensional US (3D-US) has the potential to enhance the diagnostic accuracy and precision for thyroid cancer (TC) detection.

*Materials and Methods:* An experienced ultrasonographer (G.A.) evaluated the patient using the following techniques: B-mode, strain and shear wave elastography, and 3D-US followed by fine needle aspiration biopsy (FNAB). Laboratory measurements were performed at LabCorp. Informed consent was obtained.

**Case:** A 28-year-old woman referred for hypothyroidism. Her primary doctor initiated levothyroxine 50 mcg daily 6 months prior. At the time of her visit, her thyroid stimulating hormone (TSH) was 2.8 (0.45–4.5 ulU/mL) and both thyroid peroxidase and thyroglobulin antibodies were elevated, suggestive of Hashimoto's thyroiditis. Her thyroid US showed a heterogeneous gland with an isoechoic TN in the right lobe measuring  $7.7 \times 6.3 \times 7$  mm. Strain elastography showed diffuse and patchy tissue stiffness throughout the gland, suggestive of tissue fibrosis caused by Hashimoto's thyroiditis. This study did

not distinguish target TN from the surrounding tissue. Shear wave elastography of the TN revealed moderately increased stiffness compared with surrounding tissue. The shear wave velocity (SWV) measurement for the TN was 3.1 m/s. 3D-US examination demonstrated an isoechoic TN with irregular margins, and the volume was 0.119 cm<sup>3</sup>. FNAB of the TN was performed. Cytopathology was diagnostic for papillary thyroid cancer (PTC), Bethesda Category VI. Subsequent total thyroidectomy confirmed a 7 mm PTC with positive surgical margins caused by thyroid capsule invasion and no clear-cut evidence of extra-thyroid extension.

*Discussion:* This case showcases the recent technological advances in TN imaging. Our objective is to provide an improved approach to TN management. The American College of Radiology Thyroid Imaging Reporting and Data System stratifies the malignancy risk of TN primarily based on the size and B-mode US features. This model does not recommend FNAB for any TN <10 mm regardless of malignancy risk.<sup>1</sup> This is our observation that with 3D-US the size cutoff of TN might not be an issue as with B-mode or elastography. Irregularities of the TN can be seen with 3D-US with small and large nodules equally. The finding of irregular margins on 3D-US and consulting with the patient lead us to perform FNAB. Recent publications in the journal of VideoEndocrinology showed utilizations of 3D-US in diagnosing parathyroid adenomas and TNs.

3D-US technology improves view of the target lesion by adding a third dimension, coronal view, to the transverse and longitudinal views of B-mode US.<sup>2,3</sup> B-mode imaging provides excellent view of TNs. However, it has a low sensitivity for predicting TC.<sup>4</sup> Prospective TN studies have demonstrated that adding elastography to B-mode imaging improves sensitivity of US technology for detecting TC.<sup>5–10</sup> In a prospective study with 707 TN, we showed that a single cutoff analysis for predicting malignancy in TNs, a maximum SWV of 3.54 m/s had the best sensitivity. The mean SWV for benign nodules was 2.71 m/s. The mean SWV for malignant nodules was 3.96 m/s.<sup>6</sup> In this particular case strain and shear wave were not as helpful. The discrepancy between the two systems has been described in cases with severe Hashimoto's thyroiditis associated with tissue fibrosis.<sup>6</sup> In our experience, the presence of autoimmune thyroid disease increases the risk for malignancy. Recent publications reported an association between differentiated TC and autoimmune thyroid disease and/or TSH when all Bethesda classifications were included.<sup>11–13</sup>

**Conclusion:** 3D-US technology in conjunction with B-mode may improve diagnostic accuracy in detecting TC.

No competing financial interests exist. Runtime of video: 2 mins 30 secs

Keywords: 3D ultrasound, thyroid nodule, thyroid cancer

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### Cite this video

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#### References

- 1. Tessler FN, Middleton WD, Grant EG, et al. ACR thyroid imaging, reporting and data system (TI-RADS): White paper of the ACR TI-RADS committee. J Am Coll Radiol 2017;587–595
- 2. Azizi G, Mayo ML, Keller J, Farrell J, Malchoff C. Diagnosis of parathyroid adenomas with new ultrasound imaging modalities. VideoEndocrinology. 2019;6. DOI: 10.1089/ve.2019.0163.
- 3. Azizi G, Faust K, Mayo M, Farrell J, Malchoff C. Diagnosis of thyroid nodule with new ultrasound imaging modalities. VideoEndocrinology 2020;7. DOI: 10.1089/ve.2020.0173.

- 4. Gharib H, Papini E, Paschke R, Duick DS, Valcavi R, Heged€us L, Vitti P. American Association of Clinical Endocrinologists, Associazione Medici Endocrinologi, and European Thyroid Association Medical guidelines for clinical practice for the diagnosis and management of thyroid nodules: Executive summary of recommendations. AACE/AME/ETA Task Force on Thyroid Nodules. Endocr Pract 2010;16:468–475.
- 5. Xu JM, Xu XH, Xu HX, et al. Conventional US, US elasticity imaging, and acoustic radiation force impulse imaging for prediction of malignancy in thyroid nodules. Radiology 2014;272:577–586.
- 6. Azizi G, Keller JM, Mayo ML, Piper K, Puett D, Earp KM, Malchoff CD. Thyroid nodules and shear wave elastography: A new tool in thyroid cancer detection. Ultrasound Med Biol 2015;41:2855–2865.
- 7. Liao LJ, Chen HW, Hsu WL, Chen YS. Comparison of strain elastography, shear wave elastography, and conventional ultrasound in diagnosing thyroid nodules. J Med Ultrasound 2019;27:26–32.
- 8. Azizi G, Keller J, Lewis M, Puett D, Rivenbark K, Malchoff C. Performance of elastography for the evaluation of thyroid nodules: A prospective study. Thyroid 2013;23:734–740.
- Bhatia KS, Lam AC, Pang SW, Wang D, Ahuja AT. Feasibility study of texture analysis using ultrasound shear wave elastography to predict malignancy in thyroid nodules. Ultrasound Med Biol 2016;42:1671–1680.
- Friedrich-Rust M, Vorlaender C, Dietrich CF, et al. Evaluation of strain elastography for differentiation of thyroid nodules: Results of a prospective DEGUM multicenter study. Ultraschall Med 2016;37:262–270.
- 11. Azizi G, Keller JM, Lewis M, et al. Association of Hashimoto's thyroiditis with thyroid cancer. Endocr Relat Cancer 2014;21:845–852.
- Haymart MR, Repplinger DJ, Leverson GE, et al. Higher serum thyroid stimulating hormone level in thyroid nodule patients is associated with greater risks of differentiated thyroid cancer and advanced tumor stage. J Clin Endocrinol Metab 2008;93:809–814.
- 13. Kim ES, Lim DJ, Baek KH, et al. Thyroglobulin antibody is associated with increased cancer risk in thyroid nodules. Thyroid 2010;20:885–891.

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