

Columnar Cell Variant of Papillary Carcinoma of the Thyroid Gland in a 22-Year-Old African Female

Yemisi Oluseyi Kila Uvie-Emegbo, John Olufemi Ogunbiyi, Nasir Akanmu Ibrahim¹, Sa'eid Funsho Ahmad¹

Department of Pathology and Forensic Medicine, Lagos State University Teaching Hospital, ¹Department of Surgery, Division of General Surgery, Lagos State University Teaching Hospital, Lagos, Nigeria

Abstract

Columnar cell variant of papillary carcinoma of the thyroid gland is a rare and aggressive thyroid malignancy with the clinical course predicated on the clinical stage, with the presence or absence of extrathyroid invasion. The tumors tend to occur in the elderly. We herein report a case of columnar cell variant of papillary carcinoma of the thyroid gland with lymph node metastasis in a 22-year-old African female who presented with 2 years' history of anterior neck swelling. The clinical examination, imaging, and cytological examination were suggestive of a benign neoplasm. An initial subtotal thyroidectomy was histologically diagnosed to be a columnar variant of papillary carcinoma. She subsequently had a completion thyroidectomy with resection of residual malignant thyroid tissue and lymph node, which showed metastasis. The patient is on follow-up.

Keywords: Papillary thyroid carcinoma, thyroid cancer, thyroidectomy

Received on: 23-09-20 **Review completed on:** 20-01-21 **Accepted on:** 24-01-21 **Published on:** 09-08-21

INTRODUCTION

Among endocrine cancers, thyroid cancer remains the most common, and its incidence has continuously increased worldwide,^[1] except in Africa where the trend has not been substantiated as detection has been insufficient.^[2] Papillary thyroid carcinoma (PTC) is the most common histological type among the thyroid cancers globally and accounts for 80%–85% of all thyroid cancers.^[1,3] Figures for Nigeria are variable and mostly lower than these but, in many cases, at least Papillary carcinoma is predominant.^[4-6] PTC is commonly seen in countries having iodine sufficient or iodine excess diets, but the most important etiological factor is said to be its association with radiation, especially occurring early in young age.^[7] PTC can occur at any age, but most tumors are diagnosed in patients in the third to fifth decades of life. Women are more frequently affected than men in a ratio ranging between 2:1 and 4:1.^[8,9]

Several variants of PTC have been documented, and each histopathologic variant of PTC has a varied clinical course different from the classical PTC.^[10] The columnar cell variant of PTC (CCPTC) is a rare variant that was first described by Evans in 1986, and it accounts for 0.15-0.2% of all PTCs.^[11,12]

Recently, the revised American Thyroid Association guidelines categorized variants of PTC according to their biological behavior and classified the CCPTC as an aggressive type with widespread dissemination and a fatal outcome.^[12]

CASE REPORT

A 22-year-old woman presented with an anterior neck swelling of 2 years' duration. There was neither associated pain nor difficulty with breathing, but the swelling progressively increased more to the left side of the neck. There was no complaint of obstructive symptoms, no weight loss, and no hypothyroid or hyperthyroid symptoms. There was also no known history of exposure to radiation. Examination revealed a palpable mobile mass more to the left with the left lateral margin about 7.0 cm from the midline. The mass was about 6.0

Address for correspondence: Dr. Yemisi Oluseyi Kila Uvie-Emegbo, Department of Pathology and Forensic Medicine, Lagos State College of Medicine, 1-5 Oba Akinjobi Way, Ikeja, Lagos, Nigeria.
E-mail: yemisi.kila@lasucom.edu.ng

This is an open access journal, and articles are distributed under the terms of the Creative Commons Attribution-NonCommercial-ShareAlike 4.0 License, which allows others to remix, tweak, and build upon the work non-commercially, as long as appropriate credit is given and the new creations are licensed under the identical terms.

For reprints contact: WKHLRPMedknow_reprints@wolterskluwer.com

How to cite this article: Uvie-Emegbo YO, Ogunbiyi JO, Ibrahim NA, Ahmad SF. Columnar cell variant of papillary carcinoma of the thyroid gland in a 22-year-old African female. *Ann Trop Pathol* 2021;12:35-8.

Access this article online

Quick Response Code:



Website:
www.atpjjournal.org

DOI:
10.4103/atp.atp_51_20

cm long and moved with swallowing but not with protrusion of the tongue. The lower margin could not be palpated, and the overlying skin appeared normal. There was no palpable cervical lymph node.

Thyroid function test was normal, while an ultrasound scan of the neck revealed a left thyroid mass with cystic degeneration. Fine-needle aspiration cytology was suggestive of a benign mass. Thoracic inlet and neck radiographs of the neck showed a goiter with retrosternal extension. At surgery, the left lobe of the gland was mostly involved, and a subtotal thyroidectomy was done. Gross examination of the tumor showed a bilobular thyroid gland with an isthmus. The first lobe measured 4.0 cm × 2.5 cm × 1.0 cm, while the second lobe measured 5.5 cm × 3.5 cm × 2.5 cm. The whole specimen weighed 42 g. Cut sections from the smaller lobe showed a brownish appearance while the cut surface of the larger lobe revealed a gray-tan-colored appearance with areas of hemorrhage. Microscopically, the isthmus and the smaller lobe showed proliferating epithelial follicles with variable amounts of colloid and nothing to suggest malignancy. However, sections from the larger, second, lobe showed proliferating epithelial cells forming papillae with sometimes prominent vascular core. The cells lining the papillary core are predominantly tall columnar with vesicular nuclei arranged in a pseudostratified pattern [Figure 1]. Focally, there was invasion of the capsule [Figure 2].

Based on the initial hematoxylin and eosin sections, the diagnosis of invasive PTC (columnar cell variant) was made with a comment to exclude metastasis from the lower gastrointestinal tract.

She subsequently had a completion thyroidectomy surgery. The thyroid tissue showed residual tumor with lymph node tumor metastasis [Figure 3]. The patient is alive and clinically stable with no recurrences or metastasis as at the time of reporting, which is only 12 months since diagnosis.

DISCUSSION

PTC was considered an indolent tumor with excellent prognosis.^[13] It however represents a heterogeneous group of tumors that exhibit marked variability in macroscopic and histologic appearance. The CCPTC is a rare subset of PTC, which was initially considered to be aggressive by its having tall columnar cells, but this opinion has since been reinforced more by the nature of its biological behavior, almost always extending beyond the thyroid capsule into extrathyroid tissues by the time of its diagnosis, a rapid growth rate, and a high rate of recurrence.^[14,15] It often presents as an asymptomatic or enlarging neck mass, which can be encapsulated. It may also present as an infiltrating mass.

The macroscopic appearance of the PTC is quite variable, with lesional tumor appearing anywhere within the gland. Typically, PTC is often small in size with averages of 2–3 cm but may occasionally be large. The lesions are typically firm

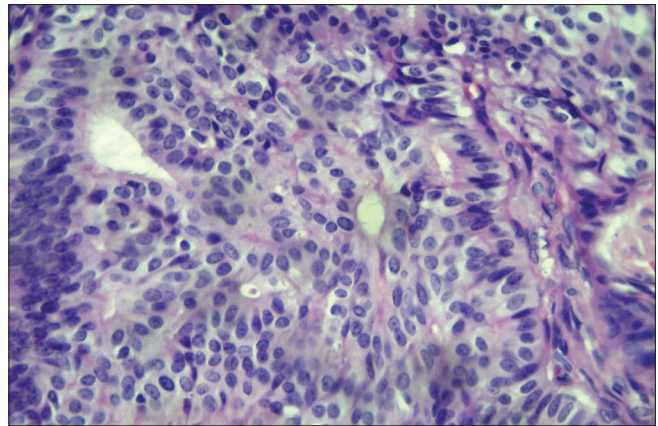


Figure 1: Columnar cell variant of papillary carcinoma (H and E, ×400) Histologic sections show cells lining are predominantly tall columnar with vesicular nuclei arranged in a pseudostratified pattern

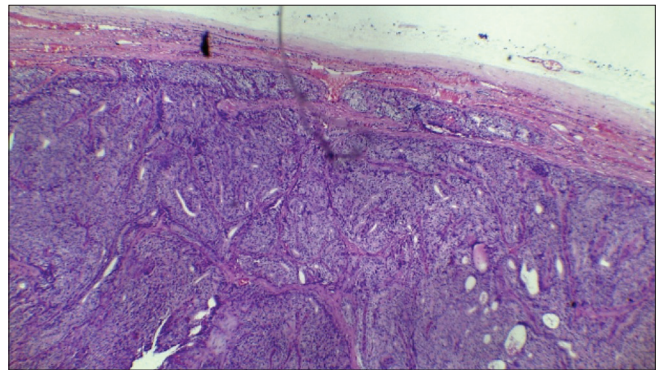


Figure 2: Capsular invasion (H and E, ×100)

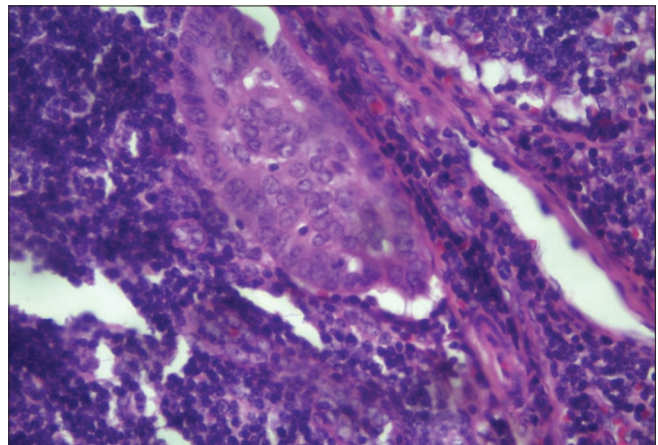


Figure 3: Lymph node tumor metastasis (H and E, ×100)

with a tan or white appearance, with infiltrating borders and often gritty cut surface. Cystic changes are relatively common especially within lymph node metastasis and calcification is a common feature.^[14,16]

The microscopic appearance of PTC is also based on constellation of architectural and cytologic features. The architectural patterns include papillary, trabecular, micro and macrofollicular, solid, diffuse sclerosing, oncotic, and

tall or columnar cell variant.^[14,16] Cystic spaces lined by one or occasionally several layers of cells with crowded oval nuclei (the neoplastic epithelial cells with characteristic enlarged optically clear, empty “Orphan Annie” nuclei, nuclear grooves, cytoplasmic pseudoinclusions, and overlapping nuclei) are commonly present. Psammoma bodies can be seen in up to 50% of cases. Lymphatic invasion is common and is often associated with a cystic growth pattern in the lymph node. The stroma is often abundant and fibrous.^[14,16]

CCPTC tumors however, often extend beyond the thyroid capsule and are usually larger (>6.0 cm) than the conventional PTC. These tumors tend to occur more in older persons, with mean age of 44 years.^[8] The majority of cases present with lymph node metastasis and even distant metastasis.

Microscopically, the histopathologic features used to define CCPTC include the presence of columnar appearing cells with nuclear stratification, scant cytoplasm, and absent minimal features of PTC.^[16,17] In the current, we found a morphologic spectrum including papillary and micropapillary patterns.

In this case report, the distinguishing features of the aggressive nature of the CCPTC is yet to be established. With follow-up and more extensive ancillary examination, the exact biological behavior in the case can be concluded.

Despite this emphasis that CCPTC is highly aggressive and fatal, some studies have reported more favorable outcomes in certain patients who have the encapsulated form of CCPTC.^[18,19] In addition, it has also been suggested that tumors confined to the thyroid gland are associated with an excellent prognosis.^[20,21] In our own case, the columnar cell carcinoma shows capsular invasion and lymph node metastasis, which suggest guarded prognosis as no distant metastasis observed was demonstrated. It is necessary to emphasize that further ancillary investigations such as nuclear and PET scans were not carried out to exclude distant spread at the time of surgery and histological diagnosis. The patient however is still alive 12 months after the initial diagnosis was made.

In well-differentiated PTC, genetic alterations such as V600E missense mutation have been documented. In this mutation, valine is substituted by glutamic acid in the gene encoding the serine threonine kinase, b-raf (BRAF^{V600E}) resulting in activation of the receptor tyrosine kinase signaling cascade. The BRAF driver mutation has been described as an indicator of the progression and aggressiveness of PTC.^[22,23]

Some studies show a 33% detection of BRAF in CCPTC, which is comparable to findings in its overall prevalence in well-differentiated PTC.^[21,24] The presence of BRAF mutation may therefore suggest progression and aggressiveness of the tumor.^[24] BRAF has not been studied in this case because we do not have the facility to do so. However, some other studies do not agree that BRAF does show a correlation with the known prognostic variable.^[25]

Surgical resection is the main form of treatment. Encapsulated tumors are managed conservatively while, metastases, recurrences, and unencapsulated tumors that invade beyond the capsule are managed aggressively.^[26,27]

CONCLUSION

The biological behavior of this tumor is predicated on the clinical stage, with the presence or absence of extrathyroidal invasion being the singular most important parameter that treatment is based apart from the morphologic appearance and the clinical stage as well. Our patient has a tumor with capsular invasion and lymph node metastasis. With clinical follow-up, the disease-free period, recurrence or distant metastasis will be observed to further emphasize the nature of the tumor.

Declaration of patient consent

The authors certify that they have obtained all appropriate patient consent forms. In the form the patient(s) has/have given his/her/their consent for his/her/their images and other clinical information to be reported in the journal. The patients understand that their names and initials will not be published and due efforts will be made to conceal their identity, but anonymity cannot be guaranteed.

Financial support and sponsorship

Nil.

Conflicts of interest

There are no conflicts of interest.

REFERENCES

- Pellegriti G, Frasca F, Regalbuto C, Squatrito S, Vigneri R. Worldwide increasing incidence of thyroid cancer: Update on epidemiology and risk factors. *J Cancer Epidemiol* 2013;2013:965212.
- Kilfoy BA, Zheng T, Holford TR, Han X, Ward MH, Andreas Sjodin A, *et al.* International patterns and trends in thyroid cancer incidence, 1973-2002. *Cancer Causes Control* 2009;20:525-31.
- Papp S, Asa SL. When thyroid carcinoma goes bad: A morphological and molecular analysis. *Head Neck Pathol* 2015;9:16-23.
- Thomas JO, Ogunbiyi JO. Thyroid cancers in Ibadan, Nigeria. *East Afr Med J* 1995;72:231-3.
- Emmanuel I, Ramalan M, Ochigbo A, Akpa P, Yakubu D, *et al.* Malignant thyroid lesions: A histopathological perspective. *J Adv Med Res* 2019;29:1-10. [doi.: 10.9734/2019/v29i1230149].
- Solomom R, Iliyasu Y, Mohammed AZ. Histopathological pattern of thyroid lesions in Kano, Nigeria: A 10-year retrospective review (2002-2011). *Niger J Basic Clin Sci* 2015;12:55-60.
- Hunt JL. Radiation induced thyroid diseases. *Pathol Case Rev* 2009;14:224-230.
- Jemal A, Siegel R, Ward E, Hao Y, Xu J, Thun MJ. Cancer statistics, 2009. *CA Cancer J Clin* 2009;59:225-49.
- Enewold L, Zhu K, Ron E, Marrogi AJ, Stojadinovic A, Peoples GE, *et al.* Rising thyroid cancer incidence in the United States by demographic and tumor characteristics, 1980-2005. *Cancer Epidemiol Biomarkers Prev* 2009;18:784-91.
- Lee JH, Shin JH, Lee HW, Oh YK, Hahn SK, Ko EY, *et al.* Sonographic and cytopathologic correlation of papillary thyroid cancer variant. *J Ultrasound Med* 2015;34:1-15
- Evans HL. Encapsulated columnar cell neoplasm of the thyroid. A report of four cases suggesting a favourable prognosis. *Am J Surg Pathol* 1996;20:1205-11.
- Haugen BR, Alexander EK, Bible KC, Doherty GM, Mandel SJ,

- Nikiforov YE, *et al.* 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. *Thyroid* 2016;26:1-33.
13. Clark OH. Thyroid cancer and lymph node metastases. *J Surg Oncol* 2011;103:615-8.
 14. Lewis SL. Thyroid. In: Humphrey PA, Dehner LP, Pfeifer JD, editors. *The Washington Manual of Surgical Pathology*. 3rd ed., Ch. 24. Philadelphia: Wolters Kluwer Publisher; 2012.
 15. Coca-Pelaz A, Shah JP, Hernandez-Prera JC, Ghossein RA, Rodrigo JP, Hartl DM. Papillary thyroid cancer-aggressive variant and impact on management: A narrative review. *Adv Ther* 2020;37:3112-28.
 16. Williams MD, El-Naggar A. Thyroid gland. In: Gattuso NP, Reddy VR, David O, Spitz D, Haber MH, editors. *Differential Diagnosis in Surgical Pathology*. 2nd ed., Saunders Elsevier Publisher 2010. 978-1-4160-4580-9.
 17. Bongiovanni M, Mermod M, Canberk S, Saglietti C, Sykiotis GP, Pusztaszeri M, *et al.* Columnar cell variant of papillary thyroid carcinoma: Cytomorphological characteristics of 11 cases with histological correlation and literature review. *Cancer Cytopathol* 2017;125:389-97.
 18. Kebebew E, Greenspan FS, Clark OH, Woeber KA, Grunwell J. Extent of disease and practice patterns for medullary thyroid cancer. *J Am Coll Surg* 2005;200:890-6.
 19. Huang WT, Yang ST, Wang SH, Chan HM, Chai CY. Encapsulated columnar cell carcinoma of thyroid: A case report. *Kaoshiung J Med Sci* 2005;21:241-4.
 20. Cho J, Shin JH, Hahn SY, Oh YL. Columnar cell variant of papillary thyroid carcinoma: Ultrasonographic and clinical differentiation between the indolent and aggressive types. *Korean J Radiol* 2018;19:1000-5.
 21. Gaertner EM, Davidson M, Wenig BM. The columnar cell variant of thyroid papillary carcinoma. Case report and discussion of an unusually aggressive thyroid papillary carcinoma. *Am J Surg Pathol* 1995;19:940-7.
 22. Xing M. BRAF mutation in papillary thyroid cancer: Pathogenic role, molecular bases, and clinical implications. *Endocr Rev* 2007;28:742-62.
 23. Sujoy V, Pinto A, Nosé V. Columnar cell variant of papillary thyroid carcinoma: A study of 10 cases with emphasis on CDX2 expression. *Thyroid* 2013;23:714-9.
 24. Chen JH, Faquin WC, Lloyd RV, Nosé V. Clinicopathological and molecular characterization of nine cases of columnar cell variant of papillary thyroid carcinoma. *Mod Pathol* 2011;24:739-49.
 25. Fonseca D, Murthy SS, Tagore R, Rao V, Rao CS, Raju KV, *et al.* BRAF status in variants of papillary thyroid carcinoma. *Int J Head Neck Pathol* 2018;1:41-47.
 26. Yunta PJ, Ponce JL, Prieto M, Merino F, Sancho-Fornos S. The important of a tumour capsule in columnar cell thyroid carcinoma: A report of two cases and review of literature. *Thyroid* 1999;9:815-819.
 27. Akslen LA, LiVolsi VA. Prognostic significance of histologic grading compared with subclassification of papillary thyroid carcinoma. *Cancer* 2000;88:1902-8.