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Review article

COEXISTENCE OF HASHIMOTO'S THYROIDITIS WITH PAPILLARY CARCINOMA THYROID: A RARE CASE REPORT WITH REVIEW OF LITERATURE

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ABSTRACT

Hashimoto's thyroiditis is an inflammatory disease of the thyroid gland. It has an autoimmune etiology. A higher incidence of papillary thyroid carcinoma with Hashimoto's thyroiditis was reported in several studies. 51 year old female patient presented with a swelling in front of the neck region since 5 years. Clinical examination revealed a swelling about 4x4x3 cm, smooth, tender, non-pulsatile and moved with deglutition. Ultrasonography revealed multinodular goiter without evidence of lymphadenopathy. Thyroid profile was done. Patient was euthyroid. FNAC reported as benign lesion. Hemithyroidectomy was done. Grossly thyroidectomy specimen i.e. hemithyroid 6x3x3 cm was received which was externally capsulated and nodular. Cut section showed a greyish white area and cystic areas each of size 1x1 cm filled with haemorrhagic and mucoid material respectively. Microscopy showed thyroid follicles with lymphoid infiltrate in the stroma forming follicles with germinal centres. Hurthle cell change was also noted. Section from both cystic areas showed plenty of complex branching papillae with fibrovascular core lined by cuboidal cells showing ground glass nuclei. The case was diagnosed as papillary carcinoma in Hashimoto's thyroiditis. The frequency of the association of Hashimoto's thyroiditis and differentiated thyroid carcinoma is approximately 30%. However, the presence of Hashimoto's thyroiditis has no effect on the diagnostic evaluation and management of papillary carcinoma of thyroid. Yet, one has to keep an eye for the features of papillary carcinoma in case of Hashimoto's thyroiditis. So a thorough grossing of thyroid specimen is recommended especially in patients who have Hashimoto's thyroiditis.

Key words: Hashimoto's thyroiditis, papillary carcinoma thyroid, coexistence.

INTRODUCTION

Hashimoto's thyroiditis, characterized by the presence of diffuse lymphocytic and plasma cell infiltration of the thyroid parenchyma and reactive germinal centres, is most typically seen in the adult population with a female predominance.¹ Papillary carcinoma is defined as a malignant epithelial tumour showing evidence of follicular cell differentiation and characterized by nuclear distinctive feature.²

Several studies report a higher rate of papillary thyroid carcinoma in patients with Hashimoto's

thyroiditis indicating possible correlation between the two diseases.³⁻⁵

There is approximately 30% frequency of the coexistence of Hashimoto's thyroiditis and differentiated thyroid carcinoma. The presence of coexistent Hashimoto's thyroiditis does not affect the diagnostic evaluation and management of papillary thyroid cancer.⁶

CASE REPORT

51 years old female patient presented with swelling in front of the neck region since 5 years. Patient had difficulty in swallowing and change in voice since 2 months. Clinical examination revealed a swelling about 4x4x3 cm, smooth, tender, non-pulsatile and moved with deglutition. On ultrasonography thyroid gland appeared diffusely bulky with well defined nodules. It was reported as features suggestive of multinodular goiter without evidence of lymphadenopathy. Thyroid profile was done. Patient was Euthyroid. FT₄ – 1.06 [N.R.- 0.8-1.9 ng/dl] FT₃ – 3.05 [N.R.- 1.5-4.1 pg/dl] TSH – 0.973 [N.R.- 0.4-4 Uiu/ml] FNAC reported as benign lesion. hemithyroidectomy was done. Grossly thyroidectomy specimen i.e. hemithyroid of size 6x3x3 cm was received which was externally capsulated and nodular. Cut section showed a greyish white area and cystic areas each of size 1x1 cm filled with haemorrhagic and mucoid material respectively. (Fig-1) Microscopy showed thyroid follicles with lymphoid infiltrate in the stroma forming follicles with germinal centers. (Fig-2,3)



Fig 1: Cut section of a thyroid showing nodule with cystic and haemorrhagic areas

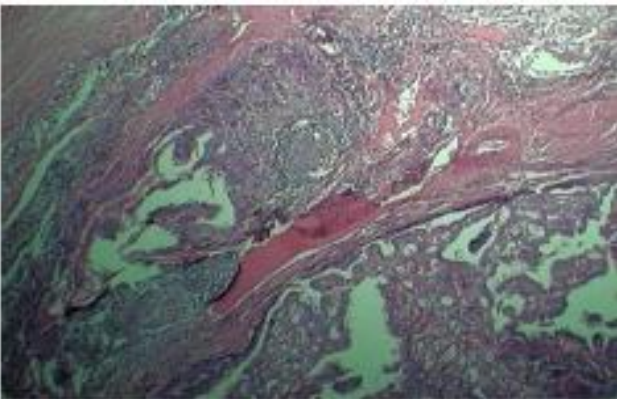


Fig 2: Thyroid having lymphoid follicles & papillary carcinoma (H& E 10x)

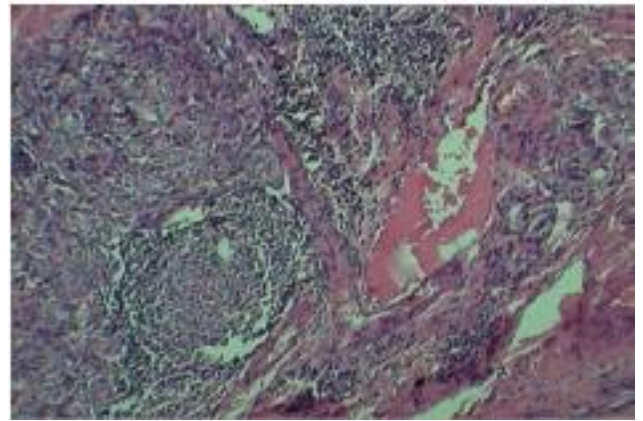


Fig 3: Lymphoid follicles with germinal centres (H& E 10x)

Hurthle cell change was also noted. Section from both cystic areas showed plenty of complex branching papillae with fibrovacuolar core lined by cuboidal cells showing ground glass nuclei. (Fig-4,5) The case was diagnosed as papillary carcinoma in Hashimoto's thyroiditis.

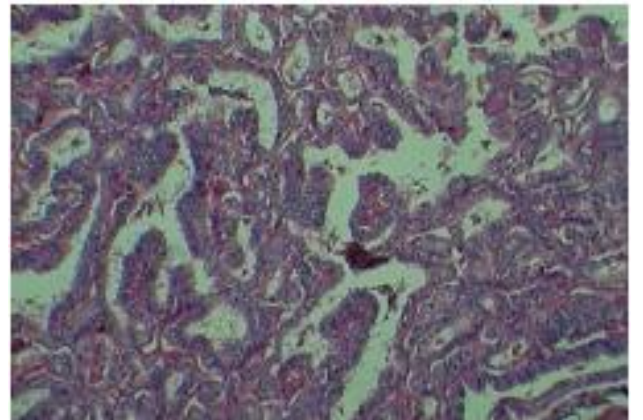


Fig 4: Papillary carcinoma (H& E 10x)

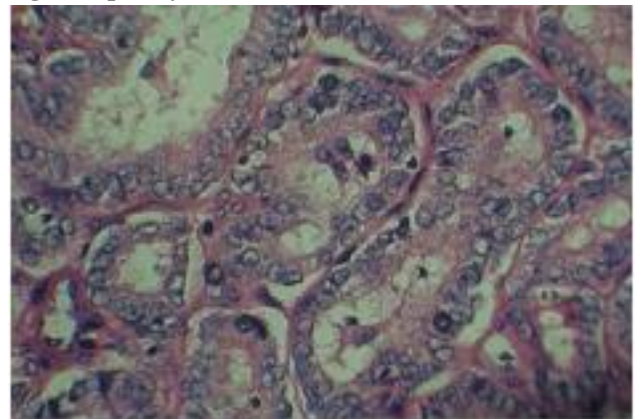


Fig 5: Papillae and ground glass like appearance of nuclei. (H& E 40x)

DISCUSSION

Hakaru Hashimoto, a Japanese surgeon, working in Berlin, Germany, first described Hashimoto's thyroiditis as a histological diagnosis. It is a part of

the spectrum of autoimmune thyroid diseases. It is known that women express thyroid autoimmunity more frequently than men and this tendency is even more obvious in the postmenopausal period.⁷ Papillary thyroid cancer is the most common form of cancer in the thyroid. It is 2.5 times more likely to develop in women than in men.⁸ In our case, patient is of 51 years female.

The relationship between Hashimoto's thyroiditis and papillary thyroid carcinoma was first proposed by Daily, et al. in 1955. A clear association between the two diseases among patients of different ethnic origin was determined by Okayasu et al. The causative relationship between Hashimoto's thyroiditis and Papillary Carcinoma thyroid is not yet clear, careful observation of Hashimoto's thyroiditis patient is recommended. The literature quotes a number of proposed mechanisms of both of these diseases and some attempts are made to explain the association. For example, Wirtschafter et al. described expression of the RET/PTC1 and RET/PTC3 oncogenes in Hashimoto's thyroiditis patient.⁸

Arif, et al. concluded papillary thyroid carcinoma and Hashimoto's thyroiditis overlap in morphological features, immunohistochemical pattern and most importantly, molecular profile. Although considered a 'benign' condition, Hashimoto's thyroiditis can harbour the RET/PTC rearrangement which is an early specific marker that is strongly associated with papillary thyroid carcinoma.⁹

In addition, expression of p63 in Hashimoto's patients with papillary thyroid cancer was found by Unger, et al. This was further examined by Burstein, et al. who proposed the two diseases are both initiated by pluripotent p63 positive stem cell remnants.⁸

Larson, et al. investigated this relationship based on the link between chronic inflammation and cancer, resulting from chronic immune response activation leading to repeated cellular damage and alteration of stromal elements. Their work revealed that patients with HT were 3 times more likely to present with associated well-differentiated thyroid carcinoma in comparison to patients without HT, supporting the existence of a link between chronic inflammation and cancer development.¹

According to Pino et al an immunological and autoimmune mechanism can be possible in etiopathogenesis of papillary carcinoma stimulating lymphocytic infiltration.¹⁰

Segal K et al¹¹ states that Hashimoto's thyroiditis does not appear to be a premalignant lesion. Thyroid carcinoma originated in the proliferating epithelium of Hashimoto's thyroiditis does not have any evidence. It would appear that thyroid carcinoma stimulate the development of HT in some patients. Autoimmune inflammatory reaction and the circulating antibodies hamper growth and metastasis of carcinoma of thyroid gland.¹¹

Neoplastic transformation is a multistep process that results in a continuous spectrum from the normal (physiological) state to a fully established neoplasm.⁹ The crux of papillary thyroid carcinoma diagnosis relies on nuclear changes: overlapping elongated ground glass nuclei with grooves and pseudoinclusions are characteristic and are most reliable features. In fact, nuclear features are the essential diagnostic component and although frequently associated with papillae, the diagnosis of papillary thyroid carcinoma can be made in their absence. The gold standard nuclear features for the diagnosis of papillary thyroid carcinoma are related to RET/PTC rearrangement.⁹

Total thyroidectomy is the surgical procedure of choice for treatment of Hashimoto's thyroiditis with papillary thyroid carcinoma.¹² The survival of the patients who have papillary thyroid cancer may be superior in coexistent Hashimoto's thyroiditis.¹³

There is a need to be cautious while screening FNAC smears if any focus of papillary thyroid carcinoma is seen. A thorough grossing of thyroid specimen is recommended. If sample sections are not taken properly and careful grossing is not done then foci of microcarcinoma may be missed in a patient who has Hashimoto's thyroiditis.¹⁴

CONCLUSION

There is approximately 30% frequency of the coexistence of Hashimoto's thyroiditis and differentiated thyroid carcinoma. Relationship between Hashimoto's thyroiditis and papillary thyroid carcinoma was first proposed by Daily, et al. in 1955. A clear association between the two diseases among patients of different ethnic origin was determined by Okayasu et al.

The literature quotes a number of proposed mechanisms of both of these diseases and some attempts are made to explain the association. For example, Wirtschafter et al. described expression of

the RET/PTC1 and RET/PTC3 oncogenes in Hashimoto's thyroiditis patient.⁸ Arif et al. concluded that Neoplastic transformation is a multistep process that results in a continuous spectrum from the normal (physiological) state to a fully established neoplasm.⁹ Expression of p63 in Hashimoto's patients with papillary thyroid cancer was found by Unger et al. According to Pino et al an immunological and autoimmune mechanism can be possible in etiopathogenia of papillary carcinoma stimulating lymphocytic infiltration.¹⁰ Segal K, et al. States that Hashimoto's thyroiditis does not appear to be a premalignant lesion.

The presence of coexistent Hashimoto's thyroiditis has no effect on the diagnostic evaluation and management of papillary carcinoma of thyroid. Yet, one has to keep an eye for the features of papillary carcinoma in case of Hashimoto's thyroiditis. So a thorough grossing of thyroid specimen is recommended especially in patients who have Hashimoto's thyroiditis.

Conflict of interest: None

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