

EUROPEAN JOURNAL OF PHARMACEUTICAL AND MEDICAL RESEARCH

www.ejpmr.com

Review Article
ISSN 2394-3211
EJPMR

18F-FDG PET/CT THYROID INCIDENTALOMA REVEALING PAPILLARY THYROID CARCINOMA: A CASE REPORT AND LITERATURE REVIEW

S. El Mselmi*, F. Foukoué, N. Abaouz and N. Ismaili Alaoui

*Department of Nuclear Medicine, Hassan II University Hospital Center, Fez, Morocco. Faculty of Medicine and Pharmacy, Sidi Mohamed BenAbdellah University, Fez, Morocco.

*Corresponding Author: Dr. S. El Mselmi

Department of Nuclear Medicine, Hassan II University Hospital Center, Fez, Morocco.

Article Received on 08/09/2020

Article Revised on 28/09/2020

Article Accepted on 18/10/2020

ABSTRACT

Thyroid incidentalomas are rarely discovered during 18F-FDG PET/CT exams for extrathyroid conditions. We report a case of papillary thyroid cancer discovered during 18F-FDG PET/CT examination carried out as part of the management of Hodgkin lymphoma. Through this observation, we underline the interest of exploring any hypermetabolic thyroid focus visualized on 18F-FDG PET/CT, by fine needle aspiration biopsy to determine its nature which may be neoplastic.

KEYWORDS: Thyroid incidentaloma; 18F-FDG; PET/CT; thyroid; cancer; Papillary carcinoma.

INTRODUCTION

Thyroid incidentalomas (TI) are described as thyroid nodule discovered accidentally during imaging studies performed for unrelated pathologies of the thyroid. [1,2] The progress in technology, especially improvements the quality of images in ultrasonography (USG), computed tomography (CT), magnetic resonance imaging (MRI), positron emission tomography/computed tomography (PET/CT) with different tracers such as 18 fluoro-2-deoxy-d-glucose (FDG), radiolabeled prostatespecific membrane antigen (PSMA), Fluorine-18 (F18), or Carbon-11 (C11) have significantly increased the rate detection of thyroid incidentalomas. [3,4] However, the risk of malignant etiology differs depending on the imaging modality used. [5,6]

We reported the case of papillary thyroid carcinoma discovered during a 18F-FDG PET/CT performed in the management of non-Hodgkin's lymphoma. Through this observation, we underlined the value of exploring any hypermetabolic thyroid focus visualized onPET/CT to determine its nature that may be neoplastic.

OBSERVATION

A 71-year-old patient treated for non-Hodgkin's lymphoma who has undergon ileal resection with anastomosis followed by adjuvant chemotherapy. The 18F-FDG PET/CT examination conducted as part of the therapeutic evaluation, showed a fairly intense hypermetabolic lesion in the left laterotracheal region (SUVmax at 10.12) with a thyroid appearance (Fig 1). Complement by cervical ultrasound was performed showing a left thyroid nodule of 30x20 mm in diameter classified score 5 of the TIRADS classification, with a second nodule of 25 mm in diameter classified as TIRADS 4. A total thyroidectomy was performed; the anatomopathological examination was in favor of a 1.5 cm papillary thyroid carcinoma invading the surrounding thyroid parenchyma massively with thyroid capsule impairment and presence of poorly differentiated foci without the presence of vascular emboli. The patient received six weeks after surgery a therapeutic supplement with iodine 131 dosed at 3.7 GBq (100 mCi). Serum thyroglobulin level after thyroid hormone withdrawal was 176 ng/ml, the anti-thyroglobulin antibodies were negative. Post-therapeutic I-131 whole body scan showed an uptake in thyroid bed compatible with thyroid remnants without foci of pathological iodine uptake at a distance.

www.ejpmr.com Vol 7, Issue 11, 2020. ISO 9001:2015 Certified Journal 124

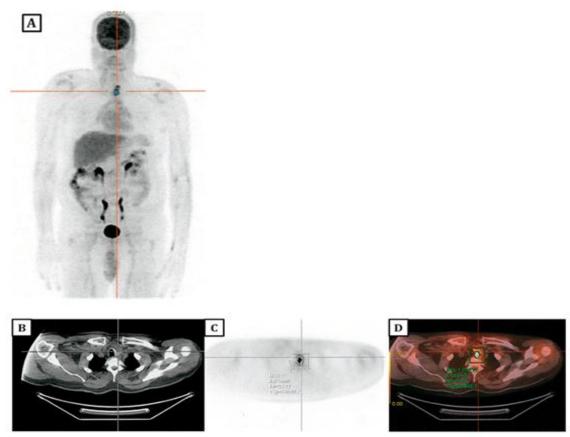


Figure 1: A: whole body 18F-FDG positron emission tomography (PET). B: Axial section computed tomography (CT), C: axial section 18F-FDG PET, D: fusion 18F -FDG PET/CT showing a focal FDG uptake on the left thyroid lobe of a patient followed-up with FDG PET/CT for non-Hodgkin's lymphoma; the histology report after thyroidectomy revealed papillary thyroid carcinoma incidentally identified in PET/CT.

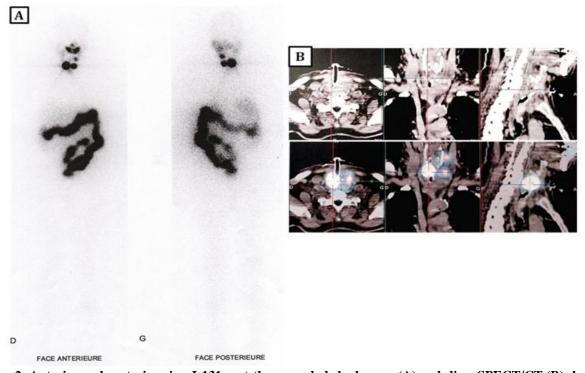


Figure 2: Anterior and posterior view I-131 post-therapy whole body scan (A) and slices SPECT/CT (B) showing an uptake in thyroid bed compatible with thyroid remnants without foci of pathological iodine uptake at a distance.

www.ejpmr.com Vol 7, Issue 11, 2020. ISO 9001:2015 Certified Journal 125

DISCUSSION

Thyroid incidentaloma, by definition, is any thyroid lesion discovered incidentally during a medical imaging examination performed for various extra-thyroid pathologies. The incidence of detecting a thyroid nodule on palpation ranges between 2% and 7%. [7,8] However, the incidence increases to 14%–46% using USG [9,10,11] and 16-56% using CT or MRI. [9,12,13]

The incidence of a TI demonstrated by 18F-FDG PET/CT compared to other imaging techniques is rare. Several authors reported a TI incidence to range between 0.2 and 8.9%. [14,15,16,17] This can be explained by the absence of the glucose transporter 1 (GLUT-1) in normal thyroid cells which is necessary for the uptake of glucose into the cell across the plasma membranes; and also by the fact that the thyroid gland uses mostly the free fatty acids for its energy metabolism. [18]

However, although rare, TI discovered on 18F-FDG PET/CT deserve special attention in exploration since the malignancy rate reported in the literature is about 8 to 64%. [4,14,17,19-20]

FDG can be captured by thyroid parenchyma in a diffuse or focused manner. The diffuse nature of hypermetabolism suggests a benign situation such as an inflammatory thyroid, [21,22] but does not exclude that a malignant lesion may coexist. However, these circumstances are very rare. [22,23]

The interpretation of the FDG uptake is facilitated by the possibility of coupling the PET images to the CT and by calculation of the SUV (Standardized Uptake Value). This latter remains a semi-quantitative parameter that reflects metabolic activity, but does not constitute a specific factor of malignancy. In this regard, some studies reported a statistically significant difference between SUVmax of benign lesions and the value of malignant ones, [24,25,26,27] but most do not find a cut-off value of SUVmax for differentiating a malignant from a benign nodule. Although it is known that malignant lesions have SUVmax value higher than benign ones. [23,28,29] In a retrospective study, Wang et al. analyzed different SUV thresholds, and considered that a value of 3.3 has a high sensitivity, and, taking into consideration the risk of malignancy, it is better to overinvestigate these cases than to under-investigate them. [30] Also, Larg et al considered that SUVmax greater than 4 was significantly associated with the risk of malignancy.[31] In our patient, the SUVmax value of the hypermetabolic thyroid focus was high, in the order of 10.12. However, other controversial studies have not shown a significant difference between SUVmax from benign and malignant lesions. [20, 23, 29, 32, 33]

Another parameter that has been evaluated by some authors is about tumor size. Indeed for some of them, there is a positive correlation between the SUV value max and tumor size, so the diameter of the lesion

influences the FDG uptake.^[31] Others found no significant correlation between SUVmax values and the size of malignant nodules.^[34] Measuring the SUV value in a small lesion can be challenging, because of the partial volume effect, which can cause an underestimation of the radiopharmaceutical uptake, so lower values of SUV are found.^[41,42] This issue is still being debated and no definitive conclusions could be drawn regarding the impact of SUV value on diagnostic driving for these patients.

The current guidelines for thyroid nodules have not established specific management plans for thyroid incidentalomas detected on 18F-FDG PET/CT. Only the American Thyroid Association (ATA) guidelines recommend performing Fine Needle Aspiration Biopsy (FNAB) for all thyroid nodules incidentally discovered on 18F-FDG PET/CT and confirmed through USG, having a dimension of >1 cm and a suspicious USG criteria. [35] In this regard, there are studies that stated that the FNAB procedure performed for thyroid nodules with suspicious USG features had benign results in 36–75% of cases, and therefore a large number of thyroid surgeries were avoided. [26,36] These results demonstrate the importance of this technique in the evaluation process of thyroid incidentalomas.

The most common primary malignancies where TIs were identified were breast cancer, Hodgkin lymphoma, colon cancer, and malignant melanoma. [15,17,31,38] In previous studies, the most common histological type of malignant thyroid incidentalomas most frequently diagnosed is represented by papillary thyroid carcinoma. [4,17,26,37] with an excellent prognosis and high survival rate. However, the undifferentiated type, anaplastic and medullary cacinoma or metastases of solid cancers are not exceptional. [17,31]

It should be noted that the capture of 18F-FDG is more pronounced in the histological sub types that are not very differentiated or anaplastics. Indeed Grabellus et al.^[39] concluded that the differentiation of thyroid carcinoma was generally accompanied by an over expression of GLUT-1 responsible for the capture of FDG. However, some thyroid carcinoma, although well differentiated, are greedy for FDG due to the presence of a pejorative histology, such as peri-thyroid extension and lympho vascular invasion that make them greedy at the FDG, as demonstrated by Choi et al.^[40]

CONCLUSION

At the end of this work, thyroid incidentalomas discovered on 18F-FDG PET/CT although rare deserve a supplement by Cervical USG to clarify the nodular nature and possible FNAB if needed, regardless of the SUVmax value, given the high risk of malignancy in this imaging modality.

www.ejpmr.com | Vol 7, Issue 11, 2020. | ISO 9001:2015 Certified Journal | 126

REFERENCES

- 1. Delivanis, D.A.; Castro, M.R. Thyroid Incidentalomas. In Thyroid Nodules; Humana Press: Cham, Switzerland, 2017; 153-167.
- Vassiliadi, D.A.; Tsagarakis, S. Endocrine Incidentalomas—Challenges Imposed by Incidentally Discovered Lesions. Nat. Publ. Gr, 2011; 7: 668-680.
- 3. Luster, M.; Duntas, L.; Wartofsky, L. The Thyroid and Its Diseases for the Clinician, 1st ed.; Springer International Publishing: Cham, Switzerland, 2019.
- Tamburello, A.; Treglia, G.; Albano, D.; Bertagna, F.; Giovanella, L. Prevalence and Clinical Significance of Focal Incidental 18F-FDG Uptake in Di_erent Organs: An Evidence-Based Summary. In Clinical and Translational Imaging; Springer: Milano, Italy, 2017; 525-532.
- Uppal A, White MG, Nagar S, Aschebrook-Kilfoy B, Chang PJ, Angelos P, et al. Benign and malignant thyroid incidentalomas are rare in routine clinical practice: A review of 97,908 imaging studies. Cancer Epidemiol Biomarkers Prev, 2015; 24: 1327–31.
- Asmar A, Simonsen L, Bolow J, Asmar M. Work-up of thyroid incidentalomas identified by 18Ffluorodeoxyglucose PET/CT. Ugeskr Laeger, 2017; 179: 2160893.
- 7. Dean DS, Gharib H. Epidemiology of thyroid nodules. Best Pract Res Clin Endocrinol Metab, 2008; 22: 901-911.
- 8. Hegedus L. Clinical practice. The thyroid nodule. N Eng J Med, 2004; 351: 1764-1771.
- 9. Jin J, Wilhelm SM, McHenry CR. Incidental thyroid nodule: patterns of diagnosis and rate of malignancy. Bitter J Surg, 2009; 197: 320-324.
- Bruel AV, Maes A, Potter TD, Mortelmans L, Drijkoningen M, Bo Van Damme BV, Delaere P, Bouillon R. Clinical relevance of thyroid fluorodeoxyglucose-whole body positron emission tomography incidentaloma. J Clin Endocrinol Metab, 2002; 87: 1517-1520.
- 11. Taheri MS, Hemadi H, Haghighatkhah HR, Kamyar K, Shakiba M, Jalali AH. Prevalence of incidental thyroid nodules diagnosed by ultrasound in an Iranian population. Iran J Radiol, 2008; 5: 19-23.
- 12. Youserm DM, Huang T, Loevner LA, Langlotz CP. Clinical and economic impact of incidental thyroid lesions found with CT and MR. Am J Neuroradiol, 1997; 18: 1423-1428.
- Shetty SK, Maher MM, Hahn PF, Halpern EF, Aquino SL. Significance of incidental thyroid lesions detected on CT: correlation among CT, sonography and pathology. Am J Roentgenol, 2006; 187: 1349-1356.
- Suh, C.H.; Kim, H.J.; Lee, J.J.; Kim, W.G. Thyroid Incidentalomas Detected on 18 F-Fluorodeoxyglucose Malignant Risk Stratification and Management Plan. Thyroid, 2018; 28: 762-768.
- 15. Soelberg KK, Bonnema SJ, Brix TH, Hegeds L. Risk of malignancy in thyroid incidentalomas

- detected by 18F-fluorodeoxyglucose positron emission tomography: a systematic review. Thyroid, 2012; 22: 918-925.
- Hagenimana, N.; Dallaire, J.; Valley, E.; Belzile, M. Thyroid Incidentalomas on 18FDG-PET/CT: A Metabolico-Pathological Correlation. J. Otolaryngol. Head Neck Surg, 2017: 46.
- 17. Bertagna, F.; Treglia, G.; Piccardo, A.; Giubbini, R. Diagnostic and Clinical Significance of F18-FDG-PET/CT Thyroid Incidentalomas. J. Clin. Endocrinol. Metab, 2012; 97: 3866–3875.
- 18. Macheda ML, Rogers S, Best JD. Molecular and cellular regulation of glucose transporter (GLUT) proteins in cancer. J Cell Physiol, 2005; 202: 654-662.
- Barrio, M.; Czernin, J.; Yeh, M.W.; Palma Diaz, M.F.; Gupta, P.; Allen-Auerbach, M.; Schiepers, C.; Herrmann, K. The Incidence of Thyroid Cancer in Focal Hypermetabolic Thyroid Lesions: An18F-FDG PET/CT Study in More than 6000 Patients. Nucl Med. Common, 2016; 37.
- 20. Chen, W.; Parsons, M.; Torigian, D.A.; Zhuang, H.; Alavi, A. Evaluation of Thyroid FDG Uptake Incidentally Identified on FDG-PET/CT Imaging. Nucl Med. Common, 2009; 30: 240–244.
- 21. Rothman IN, Middleton L, Stack BC Jr, Riggs AT, Bartel TB, Bodenner DL. The incidence of diffuse fluorodeoxyglucose positron emission tomography (PET) uptake in the thyroid of patients with autoimmune thyroiditis. Eur Arch Otolaryngol, 2011; 268: 1501-1504.
- 22. Kurata S, Ishibashi M, Hiromatsu Y, Kaida H, Miyake I, Uchida M, Hayabuchi N. Diffuse and diffuse-plus-focal uptake in the thyroid gland identified by using FDG-PET: prevalence of thyroid cancer and Hashimoto's thyroiditis. Ann Nucl Med, 2007; 21: 325-330.
- 23. Kim TY, Kim WB, Ryu JS, Gong G, Hong SJ, Shong YK. 18F-fluorodeoxyglu-cose uptake in thyroid from positron emission tomogram (PET) for evaluation in cancer patients: high prevalence of malignancy in thyroid PET incidentaloma. Laryngoscope, 2005; 115(6): 1074-8.
- 24. Ho TY, Liou MJ, Lin KJ, Yen TC Prevalence and significance of thyroid uptake detected by 18F-FDG PET. Endocrine, 2011; 40: 297-302.
- 25. Kim BH, Na MA, Kim IJ, Kim SJ, Kim YK Risk stratification and prediction of cancer of focal thyroid fluorodeoxyglucose uptake during cancer evaluation. Ann Nucl Med, 2010; 24: 721-728.
- 26. Pagano L, Sama' MT, Morani F, Prodam F, Rudoni M, Boldorini R, Valente G, Marzullo P, Baldelli R, Appetecchia M, Isidoro C, Aimaretti G Thyroid incidentaloma identified by 18F-fluorodeoxyglucose positron emission tomography with CT (FDG-PET/CT): clinical and pathological relevance. Clin Endocrinol (Oxf), 2011; 75: 528-534.
- 27. Kang BJ, O JH, Baik JH, Jung SL, Park YH, Chung SK Incidental thyroid uptake on F-18 FDG PET/CT:

- correlation with ultrasonography and pathology. Ann Nucl Med, 2009; 23: 729737.
- 28. Al-Nahhas A, Khan S, Gogbashian A, Banti E, Rampin L, Rubello D. Review. 18F-FDG PET in the diagnosis and follow-up of thyroid malignancy. In Vivo, 2008; 22(1): 109-14.
- 29. Bogsrud TV, Karantanis D, Nathan MA et al. The value of quantifying 18F-FDG uptake in thyroid nodules found incidentally on whole-body PET-CT. Nucl Med Commun, 2007; 28(5): 373-81.
- Wong, J.; Liu, K.; Siu, C.; Jones, S.; Sovka, M.; Wilson, D.; Wiseman, S.M. Management of PET Diagnosed Thyroid Incidentalomas in British Columbia Canada: Critical Importance of the PET Report. Am. J. Surg, 2017, 213; 950-957.
- Larg, M.-I.; Apostu, D.; Pes, tean, C.; Gabora, K.; Bedulescu, I.C.; Olariu, E.; Piciu, D. Evaluation of Malignancy Risk in 18F-FDG PET/CT Thyroid Incidentalomas. Diagnostics, 2019; 9: 92.
- 32. Bonabi S, Schmidt F, Broglie MA, Haile SR, Stoeckli SJ 2 February 2012 Thyroid incidentalomas in FDG-PET/CT: prevalence and clinical impact. Eur Arch Otorhinolaryngol doi:10.1007/s00405-012-1941-7
- 33. Pampaloni MH, Win AZ Prevalence and characteristics of incidentalomas discovered by whole body FDG PET CT. Int J Mol Imaging, 2012; 476763.
- 34. Sencan Eren M, Ozdogan O, Gedik A, Ceylan M, Guray Durak M, et al. The incidence of 18 F-FDG PET/CT thyroid incidentalomas and the prevalence of malignancy: a prospective study. Turk J Med Sci, 2016; 46: 840–847. doi: 10.3906/sag-1503-26.
- Haugen, B.R.; Alexander, E.K.; Bible, K.C.;
 Doherty, G.M.; Mandel, S.J.; Nikiforov, Y.E.;
 Pacini, F.; Randolph, G.W.; Sawka, A.M.;
 Schlumberger, M.; et al. 2015 American Thyroid
 Association Management. Thyroid, 2016; 26: 1-133.
- Karo, J.J.; Miller, M.M.; McLeary, R.D.; uz Zafar,
 S. Ultrasonic Evaluation of Thyroid Nodules. In Ultrasound in Medicine; Springer: Boston, MA, USA, 2012; 295-304.
- Demir, 19. Kose, N.; Ozkan, E.; U.S.an, U.S.; Arace, G.; Erdo 'gan' M.F.Clinical Significance of Thyroid Incidentalomas Identified by 18F-FDG PET/CT. It's Nucl.Mr. Med. Common, 2016; 37: 715-720.
- 38. Chung SR, Choi YJ, Suh CH, et al. Thyroid incidentalomas detected on 18F-fluorodeoxyglucose positron emission tomography with computed tomography: malignant risk stratification and management plan. It's Thyroid, 2018; 28(6): 762-768.
- 39. Grabellus F, Nagarajah J, Bockisch A, Schmid KW, Sheu SY Glucose carry 1 expression, tumor proliferation, and iodine/ glucose uptake in thyroid cancer with emphasis on poorly differentiated thyroid carcinoma. Clin Nucl Med, 2012; 37: 121-127.

- 40. Choi JW, Yoon YH, Yoon YH, Kim SM, Koo BS Characteristics of primary papillary thyroid carcinoma with false-negative findings on initial (18)F-FDG PET/CT. Ann Surg Oncol, 2011; 6: 1306-1311.
- 41. Soret, M.; Bacharach, S.L.; Buvat, I. Partial-Volume Efect in PET Tumor Imaging. J. Nucl. Med, 2007; 48: 932–945.
- 42. Teo, B.-K.; Seo, Y.; Bacharach, S.L.; Carrasquillo, J.A.; Libutti, S.K.; Shukla, H.; Hasegawa, B.H.; Hawkins, R.A.; Franc, B.L. Partial-Volume Correction in PET: Validation of an Iterative Postreconstruction Method with Phantom and Patient Data. J. Nucl. Med, 2007; 48: 802–810.

www.ejpmr.com | Vol 7, Issue 11, 2020. | ISO 9001:2015 Certified Journal | 128