Artificial Intelligence in Medical Imaging

Artif Intell Med Imaging 2021 April 28; 2(2): 13-55





Published by Baishideng Publishing Group Inc

/ M /

Artificial Intelligence in **Medical Imaging**

Contents

Bimonthly Volume 2 Number 2 April 28, 2021

MINIREVIEWS

- 13 Artificial intelligence in radiation oncology *Yakar M, Etiz D*
- **32** Intrathyroidal ectopic thymus: Ultrasonographic features and differential diagnosis *Karavas E, Tokur O, Aydın S, Gokharman D, Uner C*
- **37** Current landscape and potential future applications of artificial intelligence in medical physics and radiotherapy

Ip WY, Yeung FK, Yung SPF, Yu HCJ, So TH, Vardhanabhuti V



Contents

Artificial Intelligence in Medical Imaging

Bimonthly Volume 2 Number 2 April 28, 2021

ABOUT COVER

Editorial Board Member of Artificial Intelligence in Medical Imaging, Guo-Lin Ma, MD, PhD, Professor, Department of Radiology, China-Japan Friend Hospital, Beijing 100029, China. maguolin1007@qq.com

AIMS AND SCOPE

The primary aim of Artificial Intelligence in Medical Imaging (AIMI, Artif Intell Med Imaging) is to provide scholars and readers from various fields of artificial intelligence in medical imaging with a platform to publish high-quality basic and clinical research articles and communicate their research findings online.

AIMI mainly publishes articles reporting research results obtained in the field of artificial intelligence in medical imaging and covering a wide range of topics, including artificial intelligence in radiology, pathology image analysis, endoscopy, molecular imaging, and ultrasonography.

INDEXING/ABSTRACTING

There is currently no indexing.

RESPONSIBLE EDITORS FOR THIS ISSUE

Production Editor: Yan-Xia Xing, Production Department Director: Yun-Xiaojian Wu; Editorial Office Director: Yun-Xiaojiao Wu.

NAME OF JOURNAL Artificial Intelligence in Medical Imaging ISSN ISSN 2644-3260 (online) LAUNCH DATE

June 28, 2020

FREQUENCY Bimonthly

EDITORS-IN-CHIEF Xue-Li Chen, Caroline Chung, Jun Shen

EDITORIAL BOARD MEMBERS https://www.wjgnet.com/2644-3260/editorialboard.htm

PUBLICATION DATE April 28, 2021

COPYRIGHT © 2021 Baishideng Publishing Group Inc

INSTRUCTIONS TO AUTHORS https://www.wjgnet.com/bpg/gerinfo/204

GUIDELINES FOR ETHICS DOCUMENTS https://www.wjgnet.com/bpg/GerInfo/287 **GUIDELINES FOR NON-NATIVE SPEAKERS OF ENGLISH**

https://www.wjgnet.com/bpg/gerinfo/240

PUBLICATION ETHICS https://www.wjgnet.com/bpg/GerInfo/288

PUBLICATION MISCONDUCT https://www.wjgnet.com/bpg/gerinfo/208

ARTICLE PROCESSING CHARGE https://www.wjgnet.com/bpg/gerinfo/242

STEPS FOR SUBMITTING MANUSCRIPTS https://www.wjgnet.com/bpg/GerInfo/239

ONLINE SUBMISSION https://www.f6publishing.com

© 2021 Baishideng Publishing Group Inc. All rights reserved. 7041 Koll Center Parkway, Suite 160, Pleasanton, CA 94566, USA E-mail: bpgoffice@wjgnet.com https://www.wjgnet.com



Artificial Intelligence in Medical Imaging

Submit a Manuscript: https://www.f6publishing.com

Artif Intell Med Imaging 2021 April 28; 2(2): 32-36

DOI: 10.35711/aimi.v2.i2.32

ISSN 2644-3260 (online)

MINIREVIEWS

Intrathyroidal ectopic thymus: Ultrasonographic features and differential diagnosis

Erdal Karavas, Oguzhan Tokur, Sonay Aydın, Dilek Gokharman, Cigdem Uner

ORCID number: Erdal Karavas 0000-0001-6649-3256; Oguzhan Tokur 0000-0003-3319-6663; Sonay Aydın 0000-0002-3812-6333; Dilek Gokharman 0000-0003-1166-0576; Cigdem Uner 0000-0002-4846-7764.

Author contributions: Karavas E, Tokur O, Aydin S, Gokharman D, Uner C contribute to the conceptualization, methodology, supervision, and the manuscript writing, review and editing; Karavas E, Tokur O, Avdin S contribute to the data curation, software; Karavas E, Tokur O, Gokharman D, Uner C contribute to the validation; Karavas E, Tokur O. Uner C contribute to the visualization; Karavas E, Tokur O, Aydin S, Gokharman D contribute to the writing, original draft of the manuscript.

Conflict-of-interest statement: The authors declare no conflict of interest.

Open-Access: This article is an open-access article that was selected by an in-house editor and fully peer-reviewed by external reviewers. It is distributed in accordance with the Creative Commons Attribution NonCommercial (CC BY-NC 4.0) license, which permits others to distribute, remix, adapt, build upon this work non-commercially, and license their derivative works

Erdal Karavas, Sonay Aydın, Department of Radiology, Erzincan Binali Yildirim University, Erzincan 24100, Turkey

Oguzhan Tokur, Dilek Gokharman, Department of Radiology, Ankara Training and Research Hospital, Ankara 06230, Turkey

Cigdem Uner, Department of Child Radiology, Ankara Sami Ulus Training and Research Hospital, Ankara 06560, Turkey

Corresponding author: Erdal Karavas, MD, MSc, Associate Professor, Department of Radiology, Erzincan Binali Yildirim University, Yalnızbağ Yerleşkesi, Erzincan 24100, Turkey. erdalkaravas@hotmail.com

Abstract

Intrathyroidal ectopic thymus (IET) is defined as an ectopic thymus tissue that is generally found incidentally and rarely in the thyroid gland in the pediatric group. It occurs as a result of disruption of the embryological migration path and the settling of the thymus tissue into the thyroid gland. In the differential diagnosis, it is mostly confused with thyroid nodules. Although thyroid nodules are less common in children than adults, the rate of malignancy is much higher. Therefore, knowing the general ultrasound findings of IET better may prevent unnecessary invasive attempts and surgical procedures. In this article, we tried to compile the key imaging findings of IET.

Key Words: Thymus; Intrathyroidal thymus; Ultrasonography; Diagnosis

©The Author(s) 2021. Published by Baishideng Publishing Group Inc. All rights reserved.

Core Tip: The unique ultrasonographic features of intrathyroidal ectopic thymus (IET) can be remembered as; well-circumscribed, hypoechoic eco pattern with linear or punctate echogenic foci resembling thymus, fusiform or oval shape, diameters smaller than 1 cm, location of middle and/or lower 1/3 part of thyroid gland, hypovascularity or avascularity and the same strain ratio values with the surrounding thyroid gland on elastography. Although some studies suggested cytopathological examination for the accurate diagnosis of a suspected IET case, majority of the previous studies stated that IET can be followed without the presence of any atypical findings. So that,



on different terms, provided the original work is properly cited and the use is non-commercial. See: htt p://creativecommons.org/License s/by-nc/4.0/

Manuscript source: Invited manuscript

Specialty type: Radiology, nuclear medicine and medical imaging

Country/Territory of origin: Turkey

Peer-review report's scientific quality classification

Grade A (Excellent): 0 Grade B (Very good): B, B, B Grade C (Good): C Grade D (Fair): D Grade E (Poor): 0

Received: March 17, 2021 Peer-review started: March 17, 2021 First decision: March 26, 2021 Revised: March 29, 2021 Accepted: April 20, 2021 Article in press: April 20, 2021 Published online: April 28, 2021

P-Reviewer: Jesus-Silva SG, Madian A, Pekiner FN, Stan F S-Editor: Wang JL L-Editor: A P-Editor: Xing YX



unnecessary surgical or interventional procedures can be avoided.

Citation: Karavas E, Tokur O, Aydın S, Gokharman D, Uner C. Intrathyroidal ectopic thymus: Ultrasonographic features and differential diagnosis. *Artif Intell Med Imaging* 2021; 2(2): 32-36 **URL:** https://www.wjgnet.com/2644-3260/full/v2/i2/32.htm

DOI: https://dx.doi.org/10.35711/aimi.v2.i2.32

INTRODUCTION

Thyroid lesions in children are mostly in the benign category, but malignant lesions can also be encountered, even rarely. Intrathyroidal ectopic thymus (IET) is a benign lesion that can be encountered in children and does not require treatment. Although it has typical sonographic aspects, it may be misdiagnosed as a thyroid nodule by radiologists who do not have sufficient experience. As a result, a process leading to interventional procedures or even surgery may occur. However, some of its features conflict with thyroid nodules. In these cases, a cytopathological evaluation must be made for the distinction. If IET common ultrasound (US) features are familiarized it can be distinguished from malignant pathologies of the thyroid.

DEVELOPMENT

Thymus tissue develops in the intrauterine 6th week from the 3rd and 4th branchial saccule. Bilaterally developing primordial thymus tissues descend to the anterior mediastinum in the 8th week and combine to form the bilobuled thymus tissue at the 9th week. This descent is from the mandibular angle level to the anterior mediastinum caudally and medially. During migration, the thymic remnant can replace ectopically anywhere along the descent line in the cervical region. Although some articles report that the most ectopic cervical location is intrathyroidal[1], some articles also report it is less common than other ectopic locations in the cervical region[2]. The reason for intrathyroidal localization is though to be due to the thyroid diverticulum being close to the 3rd branchial sac, although the thyroid tissue basically develops from the 1st and 2nd branchial sac[3].

IET was first described as pathologically by Gilmour in 1937[4]. It does not present with any clinical or physical examination findings in children and is often detected incidentally. Although it is a benign condition, cases showing malignant transformation have been reported in the literature[5]. Its prevalence has been reported between 0.99% and 5.9% in studies[1,4-6]. The mean age of onset varies between 1-10 years in studies[3,5,7,8]. Although not statistically significant, it has been reported slightly more frequently in men[1,3,8].

IET consists of 2 types as abutting and enclosed types, and abutting type is more common than the other. When abutting type is seen, it should be taken into consideration that ectopic thymus tissue may be extending from the anterior mediastinum to the thyroid gland and should be examined carefully[9].

The first diagnostic modality should be US to evaluate the IET and other thyroid lesions[5]. In US, it is observed as a well-circumscribed, hypoechoic lesion containing linear or punctate echogenic foci resembling thymus[1,3,5,7,8,10]. Its contours may be irregular in some cases[8].

Internal punctate echogenicities indicate the Hassall's Corpuscle and their typical histopathological appearance confirms the diagnosis of thymus tissue[10]. Also, because of the similarity of the echo structure and presence of internal echogenic foci, IET can resemble the thymus tissue in the anterior mediastinum[10,11] (Figure 1). When the shape characteristics are examined, IET mostly presents as fusiform or oval[1,3,5,9]. In studies, its dimensions were found to be smaller than 1 cm[3,8,11]. When the location in the thyroid gland is evaluated, it is mostly observed in the middle and lower 1/3 part and more commonly in the posterior parts[1-3,11]. The reason is thought to be that the thymus develops under the pharyngeal sac, where the thyroid develops during embryological development[1]. Only 2 IETs were observed in the upper pole in the study of Erol *et al*[2].



Karavas E et al. Intrathyroidal ectopic thymus: Ultrasonographic diagnosis

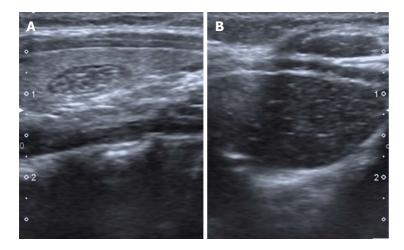


Figure 1 Eight-year-old asymptomatic girl. Longitudinal sonographic images obtained with 7 MHz linear transducer. A: Intrathyroidal ectopic thymus with typical ultrasonographic findings; hypoechoic, fusiform appearance with linear and punctate echogenic focil; B: The resemblance with mediastinal thymic tissue can be seen easily.

In color Doppler examination, IET is hypovascular or avascular compared to surrounding thyroid tissue[1,3,10,11] (Figure 2). Only a few cases have been reported to be isovascular[1,3,11]. Erol *et al*[2] stated that the observation of a vascular structure that passes through the nodule without any compression is a useful hint to differentiate IET and the other nodules.

Elastography was used in two studies, showing that IET had the same stiffness as the surrounding thyroid tissue and the average strain ratio (SR) was defined as 0.99[5,8].

In US follow-up, the dimensions of IET either remain stable or show regression[2,3,12]. Additionally, hormones and enzymes associated with the thyroid gland were also evaluated, and no significant relationship was found with IET[3,5].

In the presence of typical findings of IET described above, absence of palpable thyroid nodule and cervical lymphadenopathy, no prior risk to increased risk of malignancy (radiation exposure to the neck, family history), follow-up will be enough for the management of IET cases[1,2,5,9,10,12]. In the study of Januś *et al*[5], it was underlined that US features of IET can be similar to papillary thyroid cancer (especially diffuse sclerosing variant]. In these cases it can be challenging to differentiate the two entities, and elastography can be helpful[5,8]. Exceptionally, Stasiak *et al*[8] emphasized that cytopathological examination should be performed in cases with suspected IET and the diagnosis could only be reliable in this way[8].

IET is generally confused with thyroid nodules (Figure 3). In pediatric patients, the incidence of thyroid nodules is between 0.2%-1.5%, much less common than adults, but the rate of malignancy is higher[9]. In addition, bilateral lesions in children do not reduce the suspicion of malignancy because thyroid cancer is more common in children when it is multifocal and bilateral[8]. Therefore, further examinations should be made and the distinction between benign and malignant nodules should be revealed. If the ultrasound findings are insufficient for differentiation and the suspicion continues, fine needle aspiration biopsy should be performed first. There are some cases diagnosed by hemi lobectomy in the literature. Although IET is a benign process, it should be known that it may undergo malignant transformation such as thymoma, thymic carcinoma and lymphoblastic lymphoma[5,8]. There are some findings that support the malignant nature of a hypoechoic nodule echo pattern, solid component, ill-defined contour, irregular or round shape, microcalcification, increased vascularity, and pathological lymph node presence[8,9].

Focal thyroiditis is also included in the differential diagnosis. Focal thyroiditis is more hypoechoic and mostly does not contain diffuse echogenic focuses. The contour of focal thyroiditis is ill-defined. Vascularity is also increased (Figure 4)[2].

Intrathyroidal parathyroid gland is similar to IET, but it is differentiated by the absence of echogenic punctate foci. Clinical symptoms and laboratory findings are also useful in differential diagnosis[5].

Intrathyroidal esophageal diverticulum should be considered in differential diagnosis. Its echo pattern is isoechoic or hypoechoic compared to the surrounding thyroid tissue. Internal and peripheral echogenic focuses can be seen. In the differentiation, showing the relationship with the esophagus, changing shape with



AIMI https://www.wjgnet.com

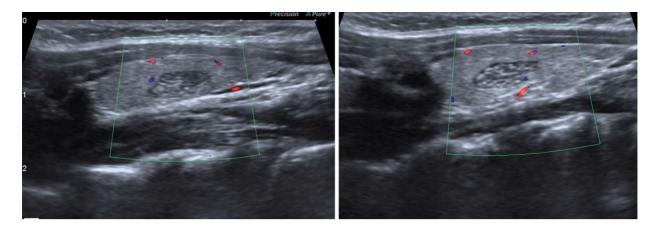


Figure 2 Eight-year-old asymptomatic girl. Longitudinal sonographic images obtained with 7 MHz linear transducer. Both intrathyroidal ectopic thymus cases are hypo vascular in comparison within the surrounding thyroid parenchyma.

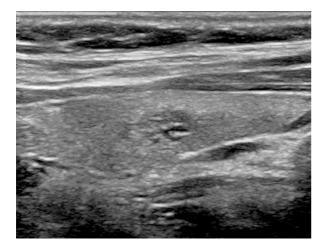


Figure 3 Nine-year-old male with hypothyroidism symptoms. Longitudinal sonographic images obtained with 7 MHz linear transducer. A thyroid nodule presenting with a similar appearance with intrathyroidal ectopic thymus tissue.

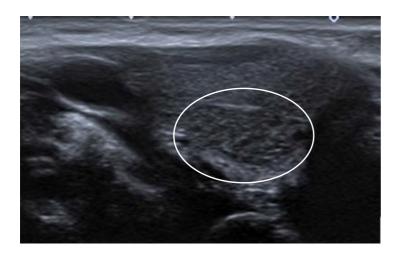


Figure 4 Ten-year-old male with recurrent cough. Longitudinal sonographic images obtained with 7 MHz linear transducer. Abutting type intrathyroidal ectopic thymus. The lesion is located at the lower half of the gland, it has unclear margins. This appearance can be confused with focal thyroiditis.

swallowing and a comet tail artifact due to the inside air can be helpful[1].

Hashimoto thyroiditis nodular form has a solid, hypoechoic echo structure and echogenic punctate focus that can also be observed. It can be well-circumscribed or illdefined. Laboratory findings may be normal in this type of Hashimoto thyroiditis. Therefore, cytopathological correlation is required[1].



Zaishidena® AIMI | https://www.wjgnet.com

Artificial intelligence (AI) has been playing an increasing role in the diagnosis of thyroidal disease[13], however as far as we know there is not a study concentrating on the use of AI in detecting IET in the English literature.

CONCLUSION

The unique ultrasonographic features of IET can be remembered as wellcircumscribed, hypoechoic eco pattern with linear or punctate echogenic foci resembling thymus, fusiform or oval shape, diameters smaller than 1 cm, location of middle and/or lower 1/3 part of thyroid gland, hypovascularity or avascularity and the same SR values with the surrounding thyroid gland on elastography. Although some studies suggested cytopathological examination for the accurate diagnosis of a suspected IET case, most stated that IET can be followed since there is absence of atypical findings. Therefore, unnecessary surgical or interventional procedures can be avoided.

REFERENCES

- Yildiz AE, Elhan AH, Fitoz S. Prevalence and sonographic features of ectopic thyroidal thymus in children: A retrospective analysis. *J Clin Ultrasound* 2018; 46: 375-379 [PMID: 29575022 DOI: 10.1002/jcu.22590]
- 2 Erol OB, Şahin D, Bayramoğlu Z, Yılmaz R, Akpınar YE, Ünal ÖF, Yekeler E. Ectopic intrathyroidal thymus in children: Prevalence, imaging findings and evolution. *Turk J Pediatr* 2017; 59: 387-394 [PMID: 29624218 DOI: 10.24953/turkjped.2017.04.004]
- 3 Aydin S, Fatihoglu E, Kacar M. Intrathyroidal ectopic thymus tissue: a diagnostic challenge. *Radiol Med* 2019; 124: 505-509 [PMID: 30710204 DOI: 10.1007/s11547-019-00987-0]
- 4 for the Report: "Molecular remission of infant B-ALL after infusion of universal TALEN gene-edited CAR T cells" by W. Qasim, H. Zhan, S. Samarasinghe, S. Adams, P. Amrolia, S. Stafford, K. Butler, C. Rivat, G. Wright, K. Somana, S. Ghorashian, D. Pinner, G. Ahsan, K. Gilmour, G. Lucchini, S. Inglott, W. Mifsud, R. Chiesa, K. S. Peggs, L. Chan, F. Farzeneh, A. J. Thrasher, A. Vora, M. Pule, P. Veys. *Sci Transl Med* 2017; **9** [PMID: 28202780 DOI: 10.1126/scitranslmed.aam9292]
- 5 Januś D, Kalicka-Kasperczyk A, Wójcik M, Drabik G, Starzyk JB. Long-term ultrasound follow-up of intrathyroidal ectopic thymus in children. *J Endocrinol Invest* 2020; 43: 841-852 [PMID: 31902058 DOI: 10.1007/s40618-019-01172-w]
- 6 Fukushima T, Suzuki S, Ohira T, Shimura H, Midorikawa S, Ohtsuru A, Sakai A, Abe M, Yamashita S, Suzuki S; Thyroid Examination Unit of the Radiation Medical Center for the Fukushima Health Management Survey. Prevalence of ectopic intrathyroidal thymus in Japan: the Fukushima health management survey. *Thyroid* 2015; 25: 534-537 [PMID: 25778711 DOI: 10.1089/thy.2014.0367]
- 7 Bang MH, Shin J, Lee KS, Kang MJ. Intrathyroidal ectopic thymus in children: A benign lesion. Medicine (Baltimore) 2018; 97: e0282 [PMID: 29620644 DOI: 10.1097/MD.000000000010282]
- 8 Stasiak M, Adamczewski Z, Stawerska R, Krawczyk T, Tomaszewska M, Lewiński A. Sonographic and Elastographic Features of Extra- and Intrathyroidal Ectopic Thymus Mimicking Malignancy: Differential Diagnosis in Children. *Front Endocrinol (Lausanne)* 2019; 10: 223 [PMID: 31110490 DOI: 10.3389/fendo.2019.00223]
- 9 Yildiz AE, Ceyhan K, Sıklar Z, Bilir P, Yağmurlu EA, Berberoğlu M, Fitoz S. Intrathyroidal Ectopic Thymus in Children: Retrospective Analysis of Grayscale and Doppler Sonographic Features. J Ultrasound Med 2015; 34: 1651-1656 [PMID: 26269296 DOI: 10.7863/ultra.15.14.10041]
- 10 Vlachopapadopoulou EA, Vakaki M, Karachaliou FE, Kaloumenou I, Kalogerakou K, Gali C, Michalacos S. Ectopic Intrathyroidal Thymus in Childhood: A Sonographic Finding Leading to Misdiagnosis. *Horm Res Paediatr* 2016; 86: 325-329 [PMID: 27756075 DOI: 10.1159/000450724]
- Tritou I, Raissaki M. Intrathyroidal ectopic thymus tissue: emphasis on details. *Radiol Med* 2019; 124: 1064-1065 [PMID: 31286340 DOI: 10.1007/s11547-019-01061-5]
- 12 Frates MC, Benson CB, Dorfman DM, Cibas ES, Huang SA. Ectopic Intrathyroidal Thymic Tissue Mimicking Thyroid Nodules in Children. J Ultrasound Med 2018; 37: 783-791 [PMID: 28850707 DOI: 10.1002/jum.14360]
- 13 Thomas J, Haertling T. AIBx, Artificial Intelligence Model to Risk Stratify Thyroid Nodules. *Thyroid* 2020; 30: 878-884 [PMID: 32013775 DOI: 10.1089/thy.2019.0752]

AIMI https://www.wjgnet.com



Published by Baishideng Publishing Group Inc 7041 Koll Center Parkway, Suite 160, Pleasanton, CA 94566, USA Telephone: +1-925-3991568 E-mail: bpgoffice@wjgnet.com Help Desk: https://www.f6publishing.com/helpdesk https://www.wjgnet.com

