



LITERATURE REVIEW

Thyroid hemiagenesis: description of two clinical cases, anatomical review, and critical synthesis of the literature

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Abstract

Introduction: Thyroid hemiagenesis (THA) is a rare congenital thyroid condition in which one of the lobes of the gland fails to develop. The etiology remains uncertain, with the main hypotheses suggesting genetic alterations in thyroid embryogenesis associated with failure of the gland to descend to its final position. Additionally, the condition may be present in both asymptomatic and euthyroid patients, as well as in those with thyroid diseases such as hyperthyroidism, nodular goiter, and neoplasia. **Objective:** To critically review the literature on the subject, detailing known aspects of anatomy in THA, and the demographic findings of patients with this condition, while also adding two case reports to the literature. **Methods:** An extensive, systematized review of the MEDLINE database was carried out using the descriptor “thyroid hemiagenesis”. A total of 214 related articles up to October 2022 were retrieved. Out of these, 154 described new cases in the literature and contained demographic and epidemiological data of the reported patients. Studies without new case reports, systematic reviews, and those lacking described or available demographic data—rendered inadmissible for this review—were excluded from the analysis. **Results:** Our review identified the following prevalence and characteristics: 520 cases of THA were reported, and in 50 of them, absence of the isthmus also coexisted. Several thyroid conditions and statuses have been associated with hemiagenesis of the gland. **Conclusion:** THA is more prevalent in female, young, and euthyroid patients. Although rare, it is vital for head and neck surgeons to be aware of this possibility, thereby avoiding unnecessary manipulation of the absent thyroid bed and, consequently, preventing complications.

Keywords: thyroid dysgenesis; thyroidectomy; embryology; genetics.

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Introduction

Thyroid hemiagenesis (THA) is a rare congenital thyroid condition in which one of the lobes of the gland develop. The etiology remains uncertain, with the main hypotheses suggesting genetic alterations in thyroid embryogenesis associated with failure of the gland to descent to its final position. Additionally, the condition may be present in both asymptomatic and euthyroid patients, as well as in those with thyroid diseases such as hyperthyroidism, nodular goiter, and neoplasia.



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THA has a prevalence of 0.05% to 0.5% in the population, and its diagnosis is generally incidental¹. This condition is more frequent in women, with a ratio of 7:1²; however, this increased prevalence in females may be related to the fact that this group has higher rates of thyroid-related diseases and, therefore, more investigations related to the gland. Lastly, the left side is more commonly affected, accounting for about 86.7% of cases in some studies³, and absence of the isthmus may also coexist in some cases⁴.

THA can be diagnosed through careful physical examination, as well as by ultrasonography, computed tomography, or scintigraphy of the thyroid. On physical examination, on the affected side, it is possible to easily palpate the edge of the trachea, and the sternocleidomastoid muscle is often much closer to the midline, sometimes covering the trachea, and not distant from it, as in the usual symptomatology. Among the imaging examinations, ultrasonography is the one that offers the best cost-benefit ratio for this diagnosis⁵.

The objective of this study was to report two cases of THA, their intraoperative findings, as well as detail known aspects of its anatomy. Additionally, a systematic review was conducted, grouping the epidemiological and anatomical data of this condition described in the literature.

Methods

Systematic review

An extensive and systematized review of the MEDLINE database was conducted using the descriptor “thyroid hemiagenesis”. A total of 214 related articles were retrieved up to October 2022. Out of these, 154 described new cases in the literature and contained demographic and epidemiological data of the reported patients, and were included in this study, with individual data shown in Table 1. Studies without new case reports, systematic reviews, and those lacking described or available demographic data—rendered inadmissible for this review — were excluded from the analysis.

Table 1. Demographic, pathological, and anatomical characteristics of cases of thyroid hemiagenesis (THA) found in the systematic review.

Study	Number of patients (age or age range)	Sex	Thyroid disease	Anatomical description, surgical or imaging findings
Gursoy et al. ⁶	12 (20-81)	9F/3M	Hashimoto's thyroiditis (n = 4)	Left hemiagenesis (n = 12)
			Multinodular goiter (n = 4)	
			Toxic adenoma (n = 1)	Absence of isthmus (n = 6)
			Euthyroid (n = 3)	
Velayutham et al. ⁷	1 (18)	1M	Subclinical hypothyroidism	Right hemiagenesis and absence of isthmus Ectopic suprahyoid and infrahyoid thyroid gland

Captions: F = female; M = male; NB = newborn; n = number; -: information not given.

Table 1. Continued...

Study	Number of patients (age or age range)	Sex	Thyroid disease	Anatomical description, surgical or imaging findings
Campenni et al. ⁸	1 (36)	1M	Graves' Disease Papillary thyroid carcinoma	Left hemiagenesis
Bhartiya et al. ⁹	1 (40)	1F	Euthyroid	Left hemiagenesis Multinodular goiter on the right side
Sasaki et al. ¹⁰	1 (28)	1F	Graves' Disease	Right hemiagenesis
Peña et al. ¹¹	1 (55)	1F	Euthyroid	Left hemiagenesis
Shaha and Gujarati ¹²	1 (30)	1F	Papillary thyroid carcinoma	Right hemiagenesis
Wang et al. ¹³	1 (31)	1F	Hashimoto's Thyroiditis	Left hemiagenesis Absence of left parathyroids
Matsumura et al. ¹⁴	1 (35)	1F	Thyrotoxicosis	Left hemiagenesis
Sharma et al. ¹⁵	1 (33)	1F	Lymphocytic thyroiditis	Left hemiagenesis
Isreb et al. ¹⁶	1 (75)	1F	Primary hyperparathyroidism	Left hemiagenesis
Kocakusak et al. ¹⁷	1 (38)	1F	Hyperthyroidism	Left hemiagenesis
Cansu et al. ¹⁸	1 (45)	1F	Graves's Disease	Left hemiagenesis and absence of isthmus
De Sanctis et al. ¹⁹	3 (6-13)	3M	Euthyroid (n=3)	Left hemiagenesis (n = 1)
				Right hemiagenesis (n = 1)
				Left hemiagenesis + absence of isthmus (n = 1)
Pizzini et al. ²⁰	1 (54)	1M	Euthyroid	Left hemiagenesis
Maiorana et al. ²¹	12 (11-14)	5F/7M	Euthyroid	Left hemiagenesis (n = 12)
Rajmil et al. ²²	2 (29-50)	2F	Euthyroid	Left hemiagenesis and absence of isthmus (n = 2)
Shabana et al. ⁵	6 (6-12)	4F/2M	Euthyroid	Left hemiagenesis (n = 3)
				Left hemiagenesis + absence of isthmus (n = 3)
Hamburger et al. ²³	1 (14)	1F	Mixed tumor: papillary and follicular carcinoma	Left hemiagenesis
Çitgez et al. ²⁴	1 (32)	1F	Follicular neoplasm	Left hemiagenesis
Park et al. ²⁵	1 (32)	1F	Euthyroid	Right hemiagenesis and absence of isthmus

Captions: F = female; M = male; NB = newborn; n = number; -: information not given.

Table 1. Continued...

Study	Number of patients (age or age range)	Sex	Thyroid disease	Anatomical description, surgical or imaging findings
Buyukdereli et al. ²⁶	3 (41-59)	1F/2M	Hyperthyroidism (n = 2)	Left hemiagenesis (n = 1)
			Euthyroid (n = 1)	Right hemiagenesis (n = 2)
Chuang et al. ²⁷	1 (30)	1F	Euthyroid	Right hemiagenesis Ectopic sublingual thyroid
Sereke et al. ²⁸	1 (32)	1F	Euthyroid	Left hemiagenesis
Kartini et al. ²⁹	1 (59)	1F	Hurthle cell carcinoma	Right hemiagenesis
Schneider et al. ³⁰	13	11F/2M	Graves' disease	-
			Hashimoto's Thyroiditis	
			Hypofunctional nodules	
			Toxic goiter	
Mikosch et al. ³¹	1 (21)	1M	Euthyroid	Left hemiagenesis and absence of isthmus
			Medullary thyroid carcinoma	Right hemiagenesis and absence of isthmus (n = 1)
Wang et al. ³²	2 (49-60)	2F	Papillary thyroid cancer	Left hemiagenesis (n = 1)
				Right hemiagenesis (n = 1)
Hsieh et al. ³³	2 (25-48)	1F/1M	Euthyroid	Left hemiagenesis (n = 1)
				Right hemiagenesis (n = 1)
Gandla et al. ³⁴	1 (20)	1F	Papillary thyroid cancer	Right hemiagenesis Absence of upper thyroid vessels ipsilateral to the hemiagenesis
Peteiro-Gonzalez et al. ³⁵	5 (16-74)	5F	Euthyroid (n = 2)	Right hemiagenesis and absence of isthmus (n = 1)
			Subclinical hypothyroidism (n = 2)	Left hemiagenesis (n = 3)
			Hyperthyroidism (n = 1)	Right hemiagenesis (n = 1)
Sari et al. ³⁶	4 (9-73)	2F/2M	Subclinical hyperthyroidism (n = 1)	Left hemiagenesis (n = 3)
			Hypothyroidism (n = 1)	Right hemiagenesis (n = 1)
			Euthyroid (n = 2)	
McHenry et al. ³⁷	7	-	Adenoma (n = 1)	Left hemiagenesis (n = 5)
			Follicular thyroid carcinoma (n = 1)	Right hemiagenesis (n = 2)
			Colloid nodule (n = 2)	
			Graves' disease (n = 2)	
			Simple goiter (n = 1)	

Captions: F = female; M = male; NB = newborn; n = number; -: information not given.

Table 1. Continued...

Study	Number of patients (age or age range)	Sex	Thyroid disease	Anatomical description, surgical or imaging findings
Karabay et al. ³⁸	1 (45)	1F	Multinodular goiter	Right hemiagenesis and absence of isthmus
Ferrari et al. ³⁹	1 (15)	1F	Primary hyperparathyroidism	Left hemiagenesis
Jha et al. ⁴⁰	1 (9)	1F	Subclinical hypothyroidism	Right hemiagenesis and absence of isthmus
Suzuki et al. ⁴¹	67 (0-23)	41F/26M	Euthyroid (n = 67)	Right hemiagenesis (n = 12) Left hemiagenesis (n = 55)
Kada et al. ⁴²	1 (48)	1F	Parathyroid carcinoma, secondary hyperparathyroidism	Right hemiagenesis
Szczepanek-Parulska et al. ¹	2 (39-45)	2F	Goiter (n = 2)	Left hemiagenesis (n = 2)
Oruci et al. ⁴³	1 (66)	1F	Parathyroid adenoma	Right hemiagenesis
Wu et al. ⁴⁴	5 (29-89)	4F/1M	Subclinical hyperthyroidism (n = 1), hypothyroidism (n = 1), euthyroid (n = 3)	Left hemiagenesis (n = 5)
Gurleyik et al. ⁴⁵	2 (25-49)	1F/1M	Hyperthyroidism (n = 1), Graves' disease (n = 1)	Left hemiagenesis (n = 2), multinodular goiter (F)
Alqahtani et al. ⁴⁶	2 (36-40)	1F/1M	Primary hyperparathyroidism to the left side with thyroid papillary microcarcinoma (n = 1), papillary carcinoma (n = 1)	Right hemiagenesis associated with suspicious multinodular left lobe, presence of isthmus, and parathyroid adenoma (n = 1) Left hemiagenesis with presence of isthmus and nodule in the right lobe (n = 1)
Yildiz et al. ⁴⁷	1 (13)	1M	Goiter	Left hemiagenesis
Chun et al. ⁴⁸	1 (1)	1F	Congenital hypothyroidism (n = 1)	Right hemiagenesis
Kirdak et al. ⁴⁹	1 (59)	1M	Subclinical hypothyroidism and plunging goiter	Left hemiagenesis
Bosco et al. ⁵⁰	1 (50)	1F	Hashimoto's Thyroiditis	Left hemiagenesis and absence of isthmus; parathyroids and inferior laryngeal nerve in the usual location
Ayaz et al. ⁵¹	1 (8)	1M	Euthyroid	Left hemiagenesis, minimal hyperplasia on the right side

Captions: F = female; M = male; NB = newborn; n = number; -: information not given.

Table 1. Continued...

Study	Number of patients (age or age range)	Sex	Thyroid disease	Anatomical description, surgical or imaging findings
Nsame et al. ⁵²	1 (23)	1F	Graves' disease with progression to Hashimoto's thyroiditis after use of anti-thyroid drugs	Left hemiagenesis, absence of isthmus
Gurleyik et al. ⁵³	1 (51)	1F	Hyperthyroidism	Left hemiagenesis, absence of isthmus; toxic adenoma in the right lobe
Sato et al. ⁵⁴	1 (64)	1F	Papillary carcinoma	Left hemiagenesis; 1.4 cm tumor on the side of the missing lobe
Reya et al. ⁵⁵	1 (21)	1F	Subclinical hyperthyroidism	Left hemiagenesis
Aydogan et al. ⁵⁶	1 (38)	1F	Thyroiditis with intermittent thyrotoxicosis	Left hemiagenesis; submandibular ectopic thyroid
Verma et al. ⁵⁷	3 (33-41)	3F	Euthyroid	Left hemiagenesis and absence of isthmus (n = 3)
Mikosch et al. ⁵⁸	16 (29-71)	13F/3M	Graves' disease (n = 1)	Left hemiagenesis (n = 10)
			Euthyroid (n = 5)	Left hemiagenesis + absence of isthmus (n = 5)
			Nodular goiter (n = 7)	Right hemiagenesis (n = 1)
			Diffuse goiter (n = 2)	
			Hashimoto's thyroiditis (n = 1)	
Korpál-Szczyrska et al. ⁵⁹	2 (7-15)	2F	Euthyroid	Left hemiagenesis and absence of isthmus (n = 2)
Ammar et al. ⁶⁰	1 (16)	1F	Euthyroid	Right hemiagenesis. Mobile, fibroelastic, 1 cm cyst in the left lobe
Baldini et al. ⁶¹	1 (41)	1F	Graves' disease	Left hemiagenesis
Cansu et al. ¹⁸	1 (45)	1F	Graves' disease	Left hemiagenesis and absence of isthmus
Kim et al. ⁶²	1 (2)	1M	Congenital hypothyroidism	Right hemiagenesis
			Microduplication of 22q11.2	
Szczepanek-Parulska et al. ⁶³	65 (40.9 ±19.7)	56F/9M	Hyperthyroidism (n = 7)	-
			Hypothyroidism (n = 20)	
			Goiter (n = 18)	
			Euthyroid (n = 28)	

Captions: F = female; M = male; NB = newborn; n = number; -: information not given.

Table 1. Continued...

Study	Number of patients (age or age range)	Sex	Thyroid disease	Anatomical description, surgical or imaging findings
Wu et al. ⁶⁴	1 (48)	1F	Euthyroid	Left hemiagenesis Multinodular nodule in the right lobe
Shah et al. ⁶⁵	1 (72)	1F	Hypothyroidism	Left hemiagenesis and absence of isthmus
Berker et al. ⁶⁶	10 (21-63)	8F/2M	Graves' disease (n = 1)	Left hemiagenesis (n = 7)
			Hashimoto's Thyroiditis (n = 1)	Right hemiagenesis (n = 3)
			Nodular goiter (n = 4)	
			Multinodular goiter (n = 2)	
			Euthyroid (n = 8)	
Cakir et al. ⁶⁷	1 (55)	1F	Graves' disease and nodular goiter	Left hemiagenesis and absence of isthmus
Szczepanek-Parulska et al. ⁶⁸	2 (10-13)	1F/1M	Hyperthyrotropinemia (n = 1) Goiter (n = 2)	Left hemiagenesis (n = 2)
Yang et al. ⁶⁹	1 (26)	1F	Goiter Lingual thyroid	Left hemiagenesis
Chang et al. ⁷⁰	2 (50-61)	2F	Euthyroid (n = 2)	Left hemiagenesis (n = 2)
Kroeker et al. ⁷¹	1 (41)	1M	Parathyroid adenoma (hypercalcemia and nephrolithiasis)	Left hemiagenesis
Ruchala et al. ²	40 (12-79)	35F/5M	Hyperthyroidism (n = 4)	Left hemiagenesis (n = 35)
			Hypothyroidism (n = 10)	Right hemiagenesis (n = 5)
			Euthyroid (n = 26)	
Castanet et al. ⁷²	22 (2-89)	14F/8M	Hypothyroidism (n = 9)	Right hemiagenesis (n = 16)
			Hyperthyroidism (n = 1)	Left hemiagenesis (n = 6)
			Euthyroid (n = 12)	
Faulkner et al. ⁷³	1 (31)	1F	Graves' disease	Left hemiagenesis
Burman et al. ⁷⁴	3 (41-58)	2F/1M	Graves' disease associated with thyrotoxicosis (n = 1)	Left hemiagenesis (n = 3)
			Euthyroid (n = 1)	
			Primary myxedema (n = 1)	
Russotto et al. ⁷⁵	1 (23)	1F	Euthyroid	Right hemiagenesis
Bough et al. ⁷⁶	1 (51)	1F	Metastatic thyroid adenocarcinoma	Left hemiagenesis

Captions: F = female; M = male; NB = newborn; n = number; -: information not given.

Table 1. Continued...

Study	Number of patients (age or age range)	Sex	Thyroid disease	Anatomical description, surgical or imaging findings
Nakamura et al. ⁷⁷	1 (28)	1F	Graves' disease with progression to hypothyroidism	Right hemiagenesis
Prakash et al. ⁷⁸	7 (8-65)	6F/1M	Toxic adenoma (n = 2)	Left hemiagenesis (n = 5)
			Colloid goiter (n = 3)	Right hemiagenesis (n = 2)
			Multinodular toxic goiter (n = 1)	
			Hashimoto's thyroiditis (n = 1)	
Greening et al. ⁷⁹	3 (47-52)	1F/2M	Adenoma (n = 2) Papillary adenocarcinoma (n = 1)	Left hemiagenesis, absence of isthmus (n = 3)
Sakurai et al. ⁸⁰	1 (42)	1M	Primary hyperparathyroidism	Right hemiagenesis
Tiwari et al. ⁸¹	1 (37)	1F	Euthyroid	Right hemiagenesis, absence of isthmus
Lee et al. ⁸²	1 (69)	1F	Papillary thyroid cancer and adenomatous hyperplasia of the right lobe	Left hemiagenesis
Philip et al. ⁸³	1 (50)	1F	Graves' disease with thyrotoxicosis	Right hemiagenesis
Melnick et al. ⁴	4 (29-58)	2F/2M	Euthyroid (n = 4)	Left hemiagenesis (n = 4)
				Absence of isthmus (n = 2)
Huang et al. ⁸⁴	1 (47)	1F	Papillary thyroid cancer and ectopic thyroid tissue	Right hemiagenesis
Ozgen et al. ⁸⁵	1 (29)	1F	Graves' disease with ophthalmopathy	Left hemiagenesis
Nayak et al. ⁸⁶	1 (45 days)	1F	Lingual cyst	Left hemiagenesis, absence of isthmus
Shechner et al. ⁸⁷	2 (33-35)	2F	Graves' disease (n = 2)	Left hemiagenesis (n = 1)
			Thyrotoxicosis (n = 1)	Right hemiagenesis (n = 1)
Meringolo et al. ⁸⁸	1 (64)	1F	Subclinical hypothyroidism with progression to Graves' disease and orbitopathy	Left hemiagenesis
Liu et al. ⁸⁹	1 (43)	1F	Graves' disease Thyrotoxicosis	Left hemiagenesis, absence of isthmus
Aslaner et al. ⁹⁰	1 (44)	1F	Multinodular goiter	Left hemiagenesis

Captions: F = female; M = male; NB = newborn; n = number; -: information not given.

Table 1. Continued...

Study	Number of patients (age or age range)	Sex	Thyroid disease	Anatomical description, surgical or imaging findings
Mydlarz et al. ⁹¹	1 (55)	1F	Primary hyperparathyroidism (adenoma)	Left hemiagenesis
Ng TT et al. ⁹²	1 (37)	1M	Euthyroid	Parapharyngeal cyst on the right side Right hemiagenesis
Hervás et al. ⁹³	1 (45)	1M	Graves' disease	Right hemiagenesis
Papi et al. ⁹⁴	1 (42)	1F	Euthyroid	Left hemiagenesis, absence of isthmus Pseudo-nodules
Stagi et al. ⁹⁵	1 (12)	1F	Williams syndrome Down syndrome	Left hemiagenesis
Nebesio et al. ⁹⁶	1 (10)	1F	Congenital hypothyroidism Graves' disease	Left hemiagenesis
Borges et al. ⁹⁷	1 (-)	1F	Congenital hypothyroidism	Left hemiagenesis
Lee et al. ⁹⁸	1 (44)	1F	Graves' disease	Left hemiagenesis
Eroglu et al. ⁹⁹	1 (27)	1F	Hyperparathyroidism	Right hemiagenesis Right parathyroid adenoma
Karatag et al. ¹⁰⁰	1 (59)	1F	Papillary carcinoma	Left hemiagenesis
Léger et al. ¹⁰¹	3 (-)	1F/2M	Euthyroid (n = 3)	Left hemiagenesis (n = 2) Right hemiagenesis (n = 1) Absence of isthmus (n = 2)
Abbassi et al. ¹⁰²	2 (4-11)	1F/1M	Euthyroid	Left hemiagenesis (n = 2)
Woods et al. ¹⁰³	1 (31)	1F	Hyperparathyroidism	Left hemiagenesis
Khatri et al. ¹⁰⁴	1 (41)	1F	Papillary adenocarcinoma	Right hemiagenesis
Hsu et al. ¹⁰⁵	1 (NB)	1F	Congenital hypothyroidism	Right hemiagenesis Ectopic sublingual thyroid
Friedman et al. ¹⁰⁶	6 (18-67)	4F/2M	Toxic goiter (n = 4) Hyperthyroidism (n = 2)	Left hemiagenesis (n = 5) Right hemiagenesis (n = 1)
Konno et al. ¹⁰⁷	1 (38)	1F	Goiter Right aortic arch	Left hemiagenesis (absence of left thyroid arteries on angiography)
Zangeneh et al. ¹⁰⁸	1 (31)	1F	Graves' disease	Left hemiagenesis

Captions: F = female; M = male; NB = newborn; n = number; -: information not given.

Table 1. Continued...

Study	Number of patients (age or age range)	Sex	Thyroid disease	Anatomical description, surgical or imaging findings
Kirmizibekmez et al. ¹⁰⁹	12 (2-3 months)	-	Congenital hypothyroidism (n=12)	Left hemiagenesis (n = 5) Right hemiagenesis (n = 7)
Jain et al. ¹¹⁰	1 (17)	1F	Goiter	Left hemiagenesis
Golinger et al. ¹¹¹	1 (52)	1F	Euthyroid	Left hemiagenesis (cystic, enlarged, right lobe and thin isthmus)
Cerqueira et al. ¹¹²	13 (children)	6F/7M	Congenital hypothyroidism(n=13)	-
Kebapçılar et al. ¹¹³	1 (43)	1F	Graves' disease Hypercalcemia	Left hemiagenesis
Rosenberg et al. ¹¹⁴	1 (3)	1M	Euthyroid	Left hemiagenesis
Hsu et al. ¹¹⁵	2 (NB)	2F	Congenital hypothyroidism (n=2)	-
Duarte et al. ¹¹⁶	5 (6-14)	2F/3M	Euthyroid	-
Shibutani et al. ¹¹⁷	1 (58)	1F	Subacute thyroiditis	Right hemiagenesis
Saydam et al. ¹¹⁸	1 (25)	1F	Euthyroid	Left hemiagenesis
Ruchala et al. ¹¹⁹	1 (49)	-	Hashimoto's thyroiditis with progression to Graves' disease and progression to hypothyroidism after combined radiotherapy and iodine therapy	Left hemiagenesis
Folsom et al. ¹²⁰	1 (18)	M	Nodular hyperplasia	Left hemiagenesis (isthmectomy, black isthmus and right lobe intraoperatively)
Véliz et al. ¹²¹	1 (35)	1F	Graves' disease	Left hemiagenesis
Duh et al. ¹²²	1 (-)	-	Hyperparathyroidism (hyperplastic parathyroidectomy; glands and nerves in usual positions)	Left hemiagenesis
Letonturier et al. ¹²³	8 (-)	8F	Simple goiter (n = 4); thyrotoxicosis (n = 2); cold nodule (n = 2)	Left hemiagenesis (n = 6) and Right hemiagenesis (n = 2)
Golden et al. ¹²⁴	1 (37 weeks)	-	Malformations (pectus excavatum, hypoplastic right clavicle, dextroscoliosis)	Right hemiagenesis
Ramos et al. ¹²⁵	3 (NB)	2F/1M	Congenital hypothyroidism	-
Bando et al. ¹²⁶	1 (40)	1F	Hashimoto's thyroiditis with progression to Graves' disease	Right hemiagenesis and absence of isthmus

Captions: F = female; M = male; NB = newborn; n = number; -: information not given.

Table 1. Continued...

Study	Number of patients (age or age range)	Sex	Thyroid disease	Anatomical description, surgical or imaging findings
Sheridan et al. ¹²⁷	1 (29)	1F	Hashimoto's Thyroiditis	Left hemiagenesis and absence of isthmus
Berni et al. ¹²⁸	1 (35)	1F	Carcinoma of the thyroglossal duct cyst	Right hemiagenesis
Tonacchera et al. ¹²⁹	6 (mean 20.5 years)	-	Euthyroid	Left hemiagenesis (n = 6)
Serdengeçti et al. ¹³⁰	1 (8)	1F	Graves' disease with ophthalmopathy	Left hemiagenesis
Alam et al. ¹³¹	1 (45)	1F	Goiter	Right hemiagenesis
Cammareri et al. ¹³²	1 (-)	1F	Williams syndrome; subclinical hypothyroidism	Left hemiagenesis
Woo et al. ¹³³	1 (5)	1F	Cervical chondrocutaneous branchial remnant	Right hemiagenesis
Gursoy et al. ¹³⁴	1 (19)	1M	Dilated cardiomyopathy and hypergonadotropic hypogonadism (DCM-HH syndrome)	Left hemiagenesis
Piera et al. ¹³⁵	3 (40-54)	3F	Adenomatous goiter (n = 3)	Left hemiagenesis (n = 2) and Right hemiagenesis (n = 1)
Supakul et al. ¹³⁶	2 (NB)	1F/1M	Congenital hypothyroidism (n = 2); ectopic thyroid (n = 1)	Right hemiagenesis (n = 2)
Hopwood et al. ¹³⁷	2 ^{5,34}	2F	Lymphocytic thyroiditis (n = 1); ectopic thyroid (n = 1)	Left hemiagenesis (n = 2)
McLean et al. ¹³⁸	1 (NB)	1M	Congenital hypothyroidism	Left hemiagenesis and absence of isthmus
Braga-Basaria et al. ¹³⁹	1 (44)	1F	Complex nodule	Left hemiagenesis
De Remigis et al. ¹⁴⁰	2 (18-66)	2F	Euthyroid	Left hemiagenesis (n = 2)
Hashemipour et al. ¹⁴¹	16 (RN)	5F/11M	Congenital hypothyroidism	-
Passeri et al. ¹⁴²	1 (RN)	-	Congenital hypothyroidism	Right hemiagenesis
Beltrão et al. ¹⁴³	1 (>3 years)	-	Ectopic thyroid; Congenital hypothyroidism	Left hemiagenesis
Devos et al. ¹⁴⁴	1 (-)	1F	Congenital hypothyroidism	Left hemiagenesis
Retnam et al. ¹⁴⁵	1 (26)	1F	Euthyroid	Left hemiagenesis
Marwaha et al. ¹⁴⁶	1 (-)	1F	Nodular goiter with lymphocytic thyroiditis	Left hemiagenesis
Leckie et al. ¹⁴⁷	2 (-)	2F	Nodular goiter (n = 2)	Left hemiagenesis (n = 2)

Captions: F = female; M = male; NB = newborn; n = number; -: information not given.

Table 1. Continued...

Study	Number of patients (age or age range)	Sex	Thyroid disease	Anatomical description, surgical or imaging findings
Tonacchera et al. ¹⁴⁸	4	4F	Euthyroid (n = 2); Subclinical hypothyroidism (n = 2)	Left hemiagenesis (n = 4)
Perry et al. ¹⁴⁹	3 (NB)	3F	Congenital hypothyroidism	Left hemiagenesis (n = 1) and Right hemiagenesis (n = 2)
Vaswani et al. ¹⁵⁰	1 (70)	1F	Adenomatous nodules	Left hemiagenesis
Srichomkwun et al. ¹⁵¹	1 (8)	1F	Congenital hypothyroidism	-
Lazzarin et al. ¹⁵²	1 (40)	1F	Hashimoto's Thyroiditis	-
Vakili et al. ¹⁵³	1 (14 months)	1F	Congenital hypothyroidism	Left hemiagenesis
Nair et al. ¹⁵⁴	2 (18 and 45 years)	2F	Euthyroid (n = 2)	Left hemiagenesis (n = 1) Right hemiagenesis (n = 1)

Captions: F = female; M = male; NB = newborn; n = number; -: information not given.

Results

Case reports

These are two patients: the first, a 58-year-old woman with hypothyroidism undergoing hormonal replacement, and the second, a 48-year-old euthyroid man, both diagnosed with thyroid nodules through routine ultrasonographic examinations. The ultrasonography of both patients, evaluated retrospectively, was suspicious for hemiagenesis of the left thyroid lobe, as shown in Table 2. The investigation with fine-needle aspiration (FNA) biopsy revealed a nodule suspected of malignancy in the woman (Bethesda IV) and consistent with papillary carcinoma (Bethesda VI) in the man. For both cases, total thyroidectomy was indicated as the treatment for this condition.

The thyroidectomy was performed under general anesthesia, with intraoperative monitoring of the vagal and laryngeal nerves and high-frequency bipolar forceps. The resection of the right lobe occurred in the usual manner without complications, including the detachment of the isthmus and the pyramidal lobe, respectively, from the pre-tracheal and pre-laryngeal fasciae. In both procedures, it was decided to keep the entire gland in a single piece. When approaching the left thyroid bed, only a projection of the lateral portion of the isthmus over the trachea was noticed, and the complete absence of the left lobe. There was no contact of the remaining thyroid tissue with the inferior laryngeal nerve or the parathyroids, which were found in their usual positions, as shown in Figure 1. In the female patient's case, there was no evidence of vascularization of the thyroid stump by the left cervical vessels, and in the male patient's case, only the middle thyroid vein crossing the inferior laryngeal nerve was observed.

Table 2. Ultrasound features of two cases of left-lobe THA.

Case	Characteristics of the Nodules	Right lobe	Left lobe	Isthmus	Estimated gland volume
Woman, 58 years old	<p>Nodule 1: predominantly solid, isoechoic, with well-defined contours, without intervening echogenic foci, in the upper/middle third of the right lobe, measuring 3.8 x 2.3 x 1.8 cm, with peripheral and central flow.</p> <p>Nodule 2: mixed, isoechoic, with well-defined contours, without intervening echogenic foci, in the lower third at the transition between the right lobe and the isthmus, measuring 3.7 x 3.8 x 1.7 cm, with peripheral and central flow.</p>	5.8 x 2.2 x 2.2 cm (estimated volume: 14.0 mL).	3.9 x 0.9 x 1.2 cm (estimated volume: 2.1 mL)	4.4 x 2.7 x 1.9 cm (estimated volume: 11.3 mL)	27.4 mL
Man, 48 years old	<p>Solid, hypoechoic, with well-defined contours, without microcalcifications, in the upper third of the right lobe, measuring 1.0 x 1.0 x 0.5 cm, with peripheral and central vascularization.</p>	5.8 x 2.7 x 2.3 cm (estimated volume: 18.4 mL)	2.1 x 1.0 x 0.8 cm (estimated volume: 0.8 mL)	3.0 x 3.0 x 0.4 cm (estimated volume: 2.0 mL)	21.2 mL

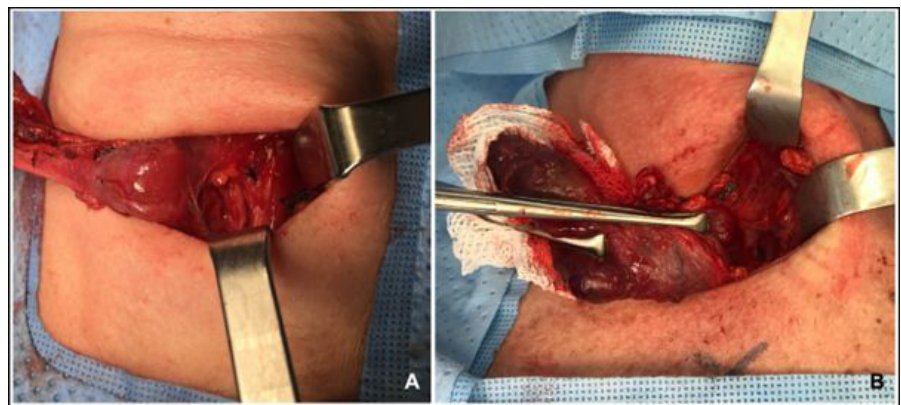


Figure 1. Intraoperative view showing the projection of the lateral portion of the isthmus over the trachea and the complete absence of the left lobe. **A** - 48-year-old man with papillary carcinoma of the right thyroid lobe. **B** - 58-year-old woman with lymphocytic thyroiditis.

A small exploration of the left bed was also conducted, with no signs of detached goiter evident. In the surgical specimen (Figure 2), one can note the vicarious right lobe, the small projection of the isthmus into the left thyroid bed, and the absence of the left lobe in both cases.

The postoperative course for both patients proceeded without complications. Inflammatory nodules due to lymphocytic thyroiditis were confirmed in the woman, and papillary carcinoma in the man, who did not require adjuvant treatment because of the low risk of the tumor and an excellent biochemical response. The patients are in outpatient follow-up, the first for 7 years and the second for 3 years, respectively on appropriate replacement and hormonal suppression.

Results from the systematic review

With the articles included in the review¹⁻¹⁵⁴, 520 patients with descriptions of THA were identified, of whom 50 (9.6%) also coexisted with the absence of the isthmus. Out of the total cases, 363 (74.1%) were women and 127 were men (in 30 cases, the patient's sex was not described). Most of these patients, or 313 cases (75.6%), presented with hemiagenesis of the left lobe, and 101 with hemiagenesis of the right lobe, with the absence of this information in 106 reports. The ages at diagnosis for the condition ranged from newborns to patients as old as 89 years, with a weighted average age of 30 years for the studies that described it.

Although many cases of THA were present in asymptomatic and euthyroid patients (234 cases - 73.1%), having been discovered in incidental findings of examinations or through screening proposed by the study, 34 patients (10.6%) had hyperthyroidism and 52 (16.3%) had hypothyroidism, in the 320 cases where thyroid function was described. Some conditions were more frequently found in patients with THA, such as Graves' disease (27 cases), Hashimoto's thyroiditis (14 cases), thyroid carcinoma (18 cases), ectopic thyroid tissue (9 cases), and congenital hypothyroidism (62 cases). Other rarer conditions, and consequently with a smaller number of associations, were also found in this review, such as Williams Syndrome.

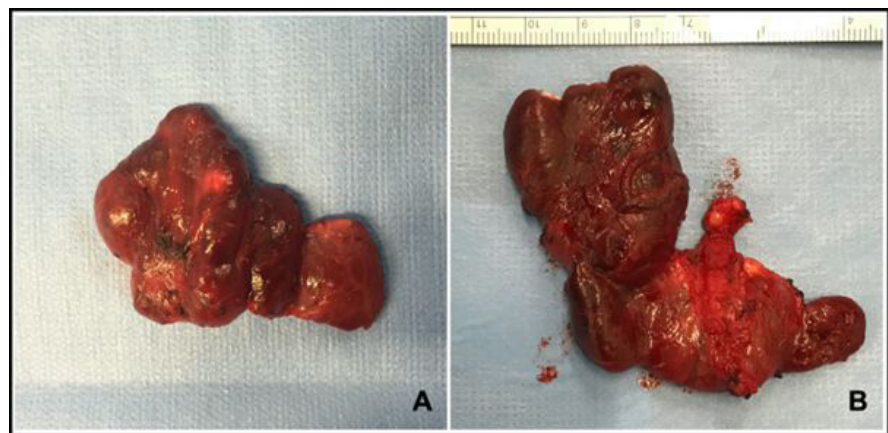


Figure 2. Specimens demonstrating the absence of the left thyroid lobe and compensatory right lobe. **A** - 48-year-old man with papillary carcinoma of the right thyroid lobe; **B** - 58-year-old woman with lymphocytic thyroiditis.

Discussion

This study demonstrates two cases of THA in patients with different glandular diseases. The condition should be suspected, especially in the evaluation of the ultrasound examination with the description of a vicarious thyroid lobe and the contralateral lobe with very reduced dimensions. Although rare, it is essential that head and neck surgeons be aware of this possibility, avoiding unnecessary manipulation of the absent thyroid bed and, consequently, preventing complications.

The current review showed a higher prevalence of left-sided THA and in females, with normal thyroid function being the condition most frequently described. The two cases reported in this study had distinct diseases, as well as completely different demographic and clinical data, showing that the condition should be assessed regardless of these characteristics.

According to the surveyed studies, the prevalence of THA in the general population was estimated between 0.05 and 0.2%⁶, but its real prevalence remains uncertain because of the number of asymptomatic and undiagnosed cases. Maiorana et al.²¹ conducted a study detecting THA in 24,032 children in Sicily, Italy, using ultrasound, and found a similar prevalence of 0.05% (12 children). Shabana et al.⁵ carried out the same type of study in 2,845 children in Belgium, where six cases of THA were detected (0.2%).

The causes of this phenomenon are also uncertain to date. One of the most accepted explanations is that sporadic cases of THA are caused by epigenetic factors, resulting in a failure in the final positioning and development of the thyroid gland⁸. The main genes related to THA are the following thyroid transcription factors: TTF1 (NKX2-1), TTF2 (FOXE1), and PAX8. Their co-expression is exclusively associated with the thyroid gland. They have an action in cellular differentiation, and mutations are linked to thyroid dysgenesis, mainly THA or ectopic thyroid⁷².

Some anatomical variations associated with thyroid hemiagenesis have been described. Gandla et al.³⁴ observed in one patient the absence of ipsilateral superior thyroid vessels to the hemiagenesis, a finding noted intraoperatively. Konno et al.¹⁰⁷ described the absence of left thyroid arteries, ipsilateral to the side of hemiagenesis, visualized in angiography. Folsom et al.¹²⁰ reported a case in which black staining of the isthmus was observed intraoperatively as a result of prior use of minocycline. The other studies did not describe other findings of anatomical changes.

Conclusion

Thyroid hemiagenesis (THA) was found to be more prevalent in female, young, and euthyroid patients. However, considering all the reports available in the literature, along with the two from the present study, this condition should be contemplated regardless of the patients' demographic scenarios and associated thyroid diseases. Although rare, it is vital for head and neck surgeons to be aware of this possibility, avoiding unnecessary manipulation of the absent thyroid bed and, consequently, preventing complications.

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