

STUDY ON HISTOPATHOLOGICAL SPECTRUM OF FOLLICULAR CELL DERIVED THYROID DISEASES

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ABSTRACT**BACKGROUND**

In clinical practice, disorders of thyroid are common worldwide and incidence of which varies with geographical as well as nutritional factors. About twelve percent of Indian adults have palpable goitre. Nowadays, thyroidectomy is a routine surgical procedure offering the chances of cure to many patients having goitre.

Aim of the study is to find out the common indications of thyroidectomy in our centre and to study the frequency and histopathological spectrum of follicular cell derived lesions in thyroidectomy specimens.

MATERIALS AND METHODS

It is a record-based retrospective study conducted in a tertiary care teaching centre in central Kerala. All thyroidectomy specimens received in the Department of Pathology in our centre from January 2011 to December 2015 for histopathological study were included in this study. Lesions involving ectopic thyroid tissue were excluded. Data including patient's age, sex, type of surgery done and histopathological diagnosis were collected from the records and analysed by SPSS package 16.0.

RESULTS

The commonest indication for thyroidectomy in all age group was follicular cell derived lesions of thyroid. Frequency of follicular cell derived lesions was 99.2% among thyroid specimens. Most common follicular cell derived lesion was multinodular colloid goitre. Papillary carcinoma was the commonest follicular cell derived malignant lesion of thyroid. Peak incidence of both neoplastic and non-neoplastic category of follicular cell derived lesions was noted in the age group of 40-49 years. Frequency of follicular cell derived lesions was significantly higher in females than males.

CONCLUSION

Current study showed comparatively higher prevalence rate of thyroid lesions and the peak incidence of follicular cell derived lesions in the age group of 40-49 years. This difference may be due to the influence of nutritional and environmental factors, so more observations and studies are warranted in this aspect.

KEYWORDS

Thyroid Disease, Thyroidectomy, Goitre, Follicular Cells.

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BACKGROUND

Thyroid gland is the largest of all endocrine glands. In clinical practice developmental, inflammatory, hyperplastic and neoplastic diseases of thyroid are common worldwide.^[1] Thyroid diseases are of great importance, because most of them are amenable to medical or surgical management. According to a projection from various studies on thyroid disease, it has been estimated that about 42 million people in India suffer from thyroid diseases.^[2] Hilly areas like Himalayan regions as well as coastal areas like Goa, Gujarat and Kerala are endemic for thyroid diseases. The government of India has

started universal iodisation of salt owing to the huge burden of thyroid diseases in the country.^[3] Goitre refers to enlarged thyroid gland, which is a common surgical problem in tertiary care hospitals. Biosynthetic defects, iodine deficiency, autoimmune diseases and nodular diseases can lead to goitre, though by different mechanisms. It is reported that twelve percent of Indian adults have palpable goitre.^[4] Nodular colloid goitre is estimated to affect at least 200 million people worldwide involving all races in all climates. Hashimoto's thyroiditis is reported about ten times more common in coastal areas of India.^[5] Based on National Cancer Registry Program reports of India, the highest relative frequency of thyroid cancer was in Thiruvananthapuram.^[2] Thyroid cancer accounts for about one percent of all malignancy in developed countries and thyroid carcinoma is the most common malignancy of the endocrine system. Carcinomas of follicular cell origin far outnumber those of para-follicular cell origin and non-epithelial malignancies. Incidence of papillary carcinoma of thyroid has increased worldwide.^[6] Prevalence of different thyroid disease varies with geographical and nutritional factors. Nowadays thyroidectomy is a routine surgical procedure because of the introduction of safe

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anaesthesia, antiseptics, fine surgical instruments and developments of new techniques, offering the chances of cure to many patients.^[7] Our study was aimed to find out the common indications of thyroidectomy in our centre and to study the frequency and histopathological spectrum of follicular cell derived lesions in thyroidectomy specimens.

MATERIALS AND METHODS

This is a retrospective study approved by Scientific Review Committee and Institutional Review Board and performed in a tertiary care teaching centre serving urban and rural population of central Kerala. All thyroidectomy specimens received for histopathological study and diagnosed in the Department of Pathology of our centre during a period of five years from January 2011 to December 2015 were included in this study. Lesions involving ectopic thyroid tissue were excluded. Data including patient’s age, sex, type of surgery done and morphological diagnosis were collected from the records. The collected data were analysed by SPSS package 16.0. Statistical significances were calculated by Chi square test and P values < 0.05 were considered statistically significant.

RESULTS

During the study period, a total of 60310 surgical specimens were received in the Department of Pathology of our centre and 3.6% of which was thyroid specimens. Table 1 shows the frequency of thyroid specimens in relation to total surgical specimens received each year.

Year		2011	2012	2013	2014	2015	Total
Specimen	Total	10270	11200	12494	13073	13273	60310
Number	Thyroid	361	437	455	412	495	2160
%		3.5	3.9	3.6	3.2	3.7	3.6

Table 1. Frequency of Thyroid Specimens

Patients who underwent thyroidectomy belong to age ranging from 8 to 90 years (Table 2). The gender distribution was predominantly females, constituting 86.6% of total cases and male-to-female ratio was 1:6.4.

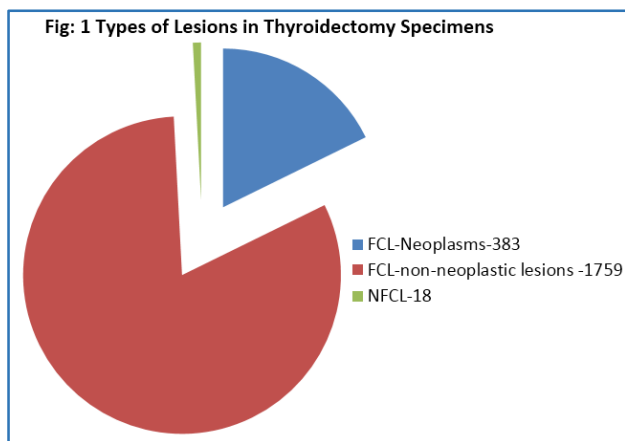
Number of Cases	Age in Years										Total
	< 10	10-19	29-29	30-39	40-49	50-59	60-69	70-79	80-89	90-99	
	2	33	153	567	739	390	220	48	7	1	2160
%	0.1	1.52	7.08	26.25	34.21	18.06	10.19	2.22	0.32	0.05	100

Table 2. Age Distribution of Thyroidectomy Cases

Among the 2160 thyroid specimens majority were (79%) total thyroidectomy specimens, followed by hemithyroidectomy 13%, near total thyroidectomy 6%, subtotal thyroidectomy and others together 2%. Histopathological study showed follicular cell derived lesions (FCL) in 2142 (99.2%) thyroid specimens that include 383 neoplastic and 1759 non-neoplastic lesions (Figure 1). Non-follicular cell derived lesions (NFCL) were observed in 18 (0.8%) thyroid specimens which include 10 para-follicular cell derived lesions, 4 lymphomas, 1 metastasis and 3 parathyroid lesions.

of 40 - 49 years, then starts decreasing in the age group of 50 - 59 years.

In our study, 60.6% of the total 2142 cases of follicular cell derived lesions belong to the age group of 30 to 49 years with a peak in the age group of 40 to 49 years. This value was found to be statistically significant with a Chi-square value of 99.327 at P < 0.001 (Table 3).



It is observed that prevalence rate of neoplastic FCL category starts increasing from 10 - 19 years of age group, then reaches maximum in the age group of 40 - 49 years and starts decreasing in the age group of 50 - 59 years, while prevalence rate of non-neoplastic FCL category starts increasing from 20-29 years of age group and reaches maximum in the age group

Age in Years	Neoplasm			Non-Neoplastic Lesions			Total No.
	No.	HP Group %	Age Group %	No.	HP Group %	Age Group %	
<10	1	0.2	50	1	0.1	50	2
10-19	16	4.2	50	16	0.9	50	32
20-29	52	13.6	34.4	99	5.6	65.6	151
30-39	91	23.8	16.1	473	26.9	83.9	564
40-49	96	25.1	13.1	639	36.3	86.9	735
50-59	56	14.6	14.5	331	18.8	85.5	387
60-69	46	12.0	21.2	171	9.7	78.8	217
70-79	22	5.7	46.8	25	1.4	53.2	47
80-89	3	0.8	50	3	0.2	50	6
90-99	0	0	0	1	0.1	100	1
Total	383	100		1759	100		2142

Table 3. Age Distribution of Follicular Cell Derived Lesions of Thyroid

Prevalence of follicular cell derived lesions was significantly higher among females than males (86.6% vs. 13.4%, $\chi^2 = 4.56, P = 0.33$). It is also noted that male-to-female ratio was 1:4.9 and 1:6.9 in neoplastic and non-neoplastic categories respectively.

Among the thirteen different morphological types of follicular cell derived lesions observed in our study, the most

common thyroid pathology was multinodular colloid goitre (63.6%), which was followed by papillary carcinoma (Table 4).

	Category	HP Type of FCL	Number	Category %	HP Group %
1	Neoplasms (n = 383)	Adenoma	107	28	5
2		Papillary Carcinoma	217	56.7	10.1
3		Follicular Carcinoma	46	12	2.1
4		Poorly Differentiated Carcinoma	8	2.1	0.4
5		Undifferentiated Carcinoma	4	1	0.2
6		Squamous Cell Carcinoma	1	0.2	0.05
7	Non-Neoplastic Lesions (n=1759)	Inflammatory Lesions	81	4.6	3.8
8		Diffuse Hyperplasia	1	0.1	0.05
9		Diffuse Colloid Goitre	4	0.2	0.2
10		Multi-Nodular Colloid Goitre	1362	77.4	63.6
11		Nodular Hyperplasia with Cellular Nodule	102	5.8	4.8
12	Solitary Cellular Nodule	48	2.7	2.2	
13	Solitary Colloid Nodule	161	9.2	7.5	
Total	2142		2142		100

Table 4. Histopathological Spectrum of Thyroid Follicular Cell Derived Lesions

Multicentricity of papillary carcinoma within the thyroid gland was seen macroscopically in 7 cases and microscopically in 24 cases. There were 8 cases of widely invasive follicular carcinoma and 38 cases of minimally invasive follicular carcinoma, four of which had morphology of minimally invasive Hurthle cell carcinoma. We observed 4 cases of undifferentiated carcinoma and 8 cases of poorly differentiated carcinoma in our study.

Second commonest lesion among non-neoplastic category was solitary colloid nodule (161 cases) and majority of which were managed by hemithyroidectomy. In 150 cases thyroid showed cellular nodule, either as a solitary lesion (48 cases) or as a part of nodular hyperplasia; 11 cases of cellular nodules had histopathological features of Hurthle cell nodule.

Inflammatory lesions were the indication for thyroidectomy in 81 cases which include 1 case of granulomatous thyroiditis (1.2%), 17 cases of lymphocytic thyroiditis (21%) and 63 cases of Hashimoto thyroiditis (77.8%). One of the Hashimoto thyroiditis case was clinically presented as multinodular goitre in a 47-year-old female. No developmental lesions were observed in our study.

DISCUSSION

Our retrospective study was performed using a database of 2160 thyroidectomy cases. Thyroid specimens constituted 3.6% of total surgical specimens received in the Department of Pathology of our centre during the study period. On comparing

observations, the prevalence of thyroid lesions was found to be higher in our study (3.6%) than that of Zulfikar et al^[1] in Karnataka and Bharathidhasan et al^[5] in Pondicherry (Table 5). This may be due to the influence of nutritional and environmental factors. We observed that majority of the patients in our study who underwent thyroidectomy were in the age group of 40-49 years, while in other studies^[1,5] it was in the age group of 30 - 39 years.

Authors		Zulfikar et al ^[1]	Bharathidhasan et al ^[5]	Present Study
Surgical Specimens	Total	6200	22676	60310
	Thyroid	1.6%	1.8%	3.6%
Thyroidectomy	Peak Age	30-39	30-39	40-49
	M:F	1:5.9	1:5.4	1:6.4
FCL	Developmental	2%	4.8%	Excluded
	Hyperplasia	57%	48%	78%
	Inflammatory	10%	11.4%	4%
	Neoplasia	31%	35.8%	18%

Table 5. Comparison of Follicular Cell Derived Lesions in Thyroidectomy Specimens with Other Studies

In the present study follicular cell derived lesions outstand (99.2%) among the different types of thyroid lesions. Prevalence of FCL was high in the age group of 40 to 49 years, while it was in the age group of 30 to 39 years in other studies.^[1,5] The prevalence of FCL was significantly higher among females. This corroborate the findings of Zulfikar et al^[1] and Bharathidhasan et al^[5] who have also reported similar finding.

Hyperplastic lesions predominate among the non-neoplastic category of FCL. Consistent with the observations of Zulfikar et al^[1] and Bharathidhasan et al,^[5] multi-nodular colloid goitre was the commonest (63.6%) follicular cell derived lesion in our study. Various factors such as iodine deficiency, goitrogens, radiation, genetic factors and certain food items have been implicated for the development of multinodular goitre.^[2,3 & 8]

The second commonest FCL (10.1%) in thyroid was papillary carcinoma, accounting for 57% of neoplastic category and 79% of malignant group. Deepa et al^[9] in 2014 and Bharathidhasan et al^[5] in 2015 reported that commonest thyroid malignancy in their study was papillary carcinoma, which was consistent with our observation. Papillary carcinoma often develops as multicentric tumour and such tumours are associated with increased risk of central and lateral neck lymph node involvement.^[10] In our study multicentricity of papillary carcinoma within the thyroid gland was seen macroscopically in 7 cases. It is also documented that microscopic multicentricity of papillary carcinoma are found in about 20% of cases if a few random sections are taken and in over 75% if step sections of entire thyroid gland is examined.^[8] In our study, microscopic multicentricity of papillary carcinoma within the thyroid gland was found in 24 cases (11%). Controversy still exists as to whether this represents multicentricity or intrathyroidal lymphatic permeation.^[8]

Follicular carcinoma contributed 2.1% of follicular cell derived lesions and 12% (46 cases) of the neoplastic lesions of thyroid. This group included 8 widely invasive neoplasms and rest were minimally invasive. Undifferentiated and poorly differentiated carcinomas of thyroid are considered as

tumours of older age group.^[8] In our study also majority of these tumours were observed in older age group.

Inflammatory lesions accounted for 3.8% of follicular cell derived lesions. Similar to other studies,^[1&5] among the different morphological types of thyroiditis the commonest indication for thyroidectomy was Hashimoto thyroiditis. In contrast to the usual pattern of diffuse symmetrical enlargement of thyroid gland, in some instances Hashimoto thyroiditis has a distinctly multinodular quality.^[8] In our study, one of the 63 cases of Hashimoto thyroiditis had multinodular pattern of thyroid enlargement.

As lesions involving ectopic thyroid tissue were excluded, no cases belonging to developmental lesions were observed in our study. It is not possible to predict on the basis of morphologic appearance alone whether the patient has clinical evidence of hyperthyroidism,^[8] so functional status of thyroid lesions in the current study was not known.

CONCLUSION

It was a retrospective study approved by Scientific Review Committee and Institutional Review Board and carried out in a tertiary care teaching centre, which serves both urban and rural population of central Kerala. In this study, thyroid specimens accounted for 3.6% (n = 2160) of total surgical specimens received over a period of five years. The commonest indication for thyroidectomy in all age group was follicular cell derived lesions of thyroid. Frequency of follicular cell derived lesions was 99.2% among thyroidectomy cases. Most common follicular cell derived lesion of thyroid was multinodular colloid goitre (63.6%).^(1,5,11-13) We observed a low incidence of thyroid neoplasms compared to other South Indian studies^(1,5); however, consistent with other reports^(5,9,11,14,15) most common thyroid malignancy was (79%) papillary carcinoma. Multicentricity of papillary carcinoma within the thyroid gland was seen macroscopically in 7 cases and microscopically in 24 cases. Frequency of follicular cell derived lesions was significantly higher in female gender with a male-to-female ratio of 1:6.5. Statistical analysis showed that in our study, thyroid lesions had a higher prevalence rate and the peak incidence of follicular cell derived lesions was in higher age group when compared to other studies.^[1,5,12] This difference may be due to the influence of nutritional and environmental factors, so more observations and studies are warranted in this aspect.

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REFERENCES

1. Zulfikar A, Ritica C, Umaru N. Study of prevalence of thyroid lesions in coastal regions of Karnataka. *Journal of Evolution of Medical and Dental Sciences* 2013;2:6995-7002.
2. Unnikrishnan AG, Menon UV. Thyroid disorders in India. An epidemiological perspective. *Indian J Endocrinol Metab* 2011;15(2):S78-S81.
3. Park K. Iodine deficiency disorders. In: Park K, editor. *Park's text book of Preventive and Social Medicine*, 23rd edn. Jabalpur:Banarsidas Bhanot; 2015. p. 643-644.
4. Menon UV, Sundaram KR, Unnikrishnan AG, et al. High prevalence of undetected thyroid disorders in an iodine sufficient adult south Indian population. *Journal of Indian Medical Association* 2009;107(2):72-7.
5. Bharathidhasan I, Goneppanavar M, Dhaka RS. Changing trends in the incidence of thyroid lesions in coastal regions of south India. *Ind J Health Sci Res* 2015;5(6):134-141.
6. DeLellis RA, Williams ED. Tumours of the thyroid and parathyroid. In: DeLellis RA, Lloyd RV, Heitz PU, et al. eds. *World Health Organization Classification of Tumours. Pathology and genetics of tumours of endocrine organs*. Lyon: IARC Press 2004:49-133.
7. Champa S, Tariq WK, Imrana Z, et al. Histopathological pattern of diagnoses in patients undergoing thyroid operations. *Rawal Med J* 2009;34(1):14-16.
8. Rosai J, Tallini G. Thyroid gland. In: Rosai J, editor. *Rosai and Ackerman's surgical pathology*, Volume 1, 10th edn. China:Elsevier; 2011. p. 500-515.
9. Kunjumon DT, Upadhyaya K. A clinicopathological audit of thyroid malignancies. A 2 year study. *International Journal of Biomedical Research* 2014;5(1):38-42.
10. Ju-Yeon K, Eun-Jung J, Sang-Ho J, et al. Clinical characteristics and prognosis of multifocal papillary thyroid carcinoma. *J Korean Surg Soc* 2010;79(6):442-446.
11. Pradeepkumar NS, Singh R, Joseph NM. Emerging trends in thyroid diseases in Tsunami hit coastal areas of Puduchery and Cuddalore, India. *Journal of Evaluation of Medical and Dental Sciences* 2012;1(5):857-863.
12. Prajapati VP, Nayak JC, Desai KS, et al. Histological study of adenomatous goitre. *National Journal of Integrated Research in Medicine* 2012;3(2):65-8.
13. Antony J, Celine TM, Chacko M. Spectrum of thyroid disorders: a retrospective study at a medical college hospital. *Thyroid Res Pract* 2014;11(2):55-9.
14. Vanderpump MP. The epidemiology of thyroid disease. *Br Med Bull* 2011;99(1):39-51.
15. Nadeem K, Akhtar N, Tarar JM. Thyroid malignancy in multi nodular goiter; incidence, a retrospective study in southern Punjab. *Professional Med J* 2013;20(4):587-90.