General Surgery

A Study on Clinicopathological Corelation and Management of Solitary Thyroid Nodule

Dr. Sandhya N¹, Dr. Rohith Yadav G^{2*}

¹Associate Professor, Dept. of Gen Surgery, KIMS Hubli, Karnataka 580022, India
²Post graduate Student, Dept. of Gen Surgery, KIMS Hubli, Karnataka 580022, India

DOI: 10.36347/sasjs.2023.v09i02.005

| **Received:** 02.01.2022 | **Accepted:** 06.02.2023 | **Published:** 10.02.2023

*Corresponding author: Dr. Rohith Yadav G Post graduate Student, Dept of Gen Surgery, KIMS Hubli, Karnataka 580022, India

Abstract

Original Research Article

Background: A solitary nodule is a goiter which, on clinical examination appears to be a single nodule in one lobe of the thyroid with no palpable abnormality elsewhere in the gland. The incidence of solitary nodule in general population in South India is 9%. This study aims to assess the presence of malignancy in patients presenting with solitary thyroid nodule and to evaluate and compare results of various investigative modalities used to diagnose a solitary thyroid nodule and the outcome of management modalities in solitary thyroid nodule. Methods: The present study on clinicopathological correlation and management of solitary thyroid nodule has been conducted on cases admitted and managed in the Department of Surgery at KIMS medical college, Hubli. Prospective analysis of 50 cases of solitary nodule thyroid in the specified period was done. These cases were selected by random sampling method and studied in detail clinically and recorded as per the proforma. Routine investigations and specific investigations including FNAC of the nodule, Thyroid profile, IDL, Plain X-ray neck, USG neck were done in all cases. Special investigations like radio-isotope scanning were not performed as the facilities were not available. All the patients were managed by surgery and diagnosis was confirmed by histo- pathological examination. The patients were grouped according to different variables like age, sex, size of the nodule, site of the nodule, functional thyroid status, FNAC reports and histo-pathological examination reports, then analyzed and compared with the previous similar studies conducted elsewhere. Results: Total of 50 cases of solitary nodule of thyroid studied. The mean age of presentation is 38.5 years. Cases in 3rd to 5th decades constitutes 72% of the cases studied. Out of 50 cases studied 46 were females and 4 were males, and the ratio comes to M:F = 1: 11.5. Out of 50 cases along with swelling in front of the neck, 3 cases had pain, 3 cases had discomfort and another 2 had dysphagia. Duration of onset symptoms varied from 15 days to 8 years. Out of 50 cases studied, 26 cases presented with nodule in right lobe of the thyroid gland and the remainder in the left lobe of thyroid. On clinical examination size of the nodule, in its largest dimension, varies from 2cm to 12cm. Out of 50 cases, two presented with features of thyrotoxicosis, one with hypothyroidism and rest all were in euthyroid state. In the study, out of 7 carcinomas, 5 were papillary and 2 follicular. Conclusions: Based on the data and results obtained in the present study, Solitary nodule of thyroid is more common in females. Solitary nodule of thyroid is more common the age group of 20-50 years. Most of the patients with solitary nodule of thyroid present with swelling, in euthyroid state. FNAC is the investigation of choice in the evaluation of solitary nodule of thyroid. Papillary carcinoma is the most common malignancy of thyroid.

Keywords: solitary nodule, Clinicopathological Corelation, Thyroid profile.

Copyright © 2023 The Author(s): This is an open-access article distributed under the terms of the Creative Commons Attribution 4.0 International License (CC BY-NC 4.0) which permits unrestricted use, distribution, and reproduction in any medium for non-commercial use provided the original author and source are credited.

INTRODUCTION

A solitary nodule is a goiter which, on clinical examination appears to be a single nodule in one lobe of the thyroid with no palpable abnormality elsewhere in the gland [1]. The solitary thyroid nodule is a topic of interest for surgeons and clinicians was studied by Warren H Cole in 1949. This study found that incidence of malignancy is higher when compared with multinodular goitre [2]. Thyroid nodules are very common entities, though varying in incidence in different geographical regions. The incidence of solitary nodule in general population in South India is 9% [3]. Thyroid nodules are very common with estimated prevalence that ranges from 4% [4] bypalpation to 67% [5] by ultrasonography. Autopsy studies reveal that 50% [6] of adults had nodules, the majority of which are impalpable. Thyroid nodules are 4 times more common

Citation: Sandhya N & Rohith Yadav G. A Study on Clinicopathological Corelation and Management of Solitary Thyroid Nodule. SAS J Surg, 2023 Feb 9(2): 77-87. in females than in men [7].

Overall incidence of malignancy in solitary thyroid nodule ranges from 10-30%. A single nodule in the thyroid is a definite clinical entity with important pathological significance. It is necessary to consider the status of opposite lobe when considering the nodule as solitary.

Another factor that influences the ultimate histopathological outcome of solitary nodule of solitary nodule thyroid is whether the definition of a solitary nodule is entirely clinical or proved by investigations like ultrasound, radio iodine scan etc. in general a solitary nodule is defined as "a palpable single clinically detected nodule in the thyroid gland that is otherwise normal." Visibility or palpability of opposite thyroid lobe precludes inclusion of such cases in this group. The usual presentation of a thyroid nodule is an asymptomatic swelling that is discovered by either the patient or the clinician. Nodules of at least 0.5cm to 1cm can be usually be detected by palpation, although estimates of nodule size vary from physician to physician. It can be difficult to palpate any nodule in patient with a thick, short neck.

The thyroid nodule has been subject of controversy with divergent opinions expressed by those who had wide experience in this field. The optimal management of thyroid nodule continues to be a source of controversy and the operative intervention recommended by most of surgeons is not always accepted by some physicians advocating either observation or suppression.

A solitary nodule is a clinical diagnosis and not a pathological diagnosis. Almost all conditions of the thyroid may present clinically as a solitary nodule. Diagnostic possibilities in case of solitary nodules are adenoma, carcinoma, thyroid cyst and palpable nodule in an evolving multinodular colloid goiter. Other rare causes of solitary nodules include inflammatory thyroid lesions and developmental abnormalities such as dermoid cyst, teratoma etc. Because of possibility of malignancy, some clinicians especially those in surgical subspecialties recommend that all nodules have to be removed. The solitary nodule in the thyroid gland may have a malignant potential and possibility of toxicity in the nodule and also complications like pressure effects and hemorrhage. Thyroid cancer has a much lower incidence compared to that of solitary thyroid nodule. Therefore, it is of utmost importance to determine which patients with solitary thyroid nodule would benefit from surgery.

As the clinical diagnosis may not correlate with the diagnosis during the time of management, this study is undertaken as it generates curiosity among the operating surgeons.

METHODS

The present study on clinicopathological correlation and management of solitary thyroid nodule has been conducted on cases admitted and managed in the Department of Surgery at KIMS medical college, Hubli.

Prospective analysis of 50 cases of solitary nodule thyroid in the specified period was done. These cases were selected by random sampling method and studied in detail clinically and recorded as per the proforma. Routine investigations and specific investigations including FNAC of the nodule, Thyroid profile, IDL, Plain X-ray neck, USG neck were done in all cases. Special investigations like radio-isotope scanning was not performed as the facilities were not available. All the patients were managed by surgery and diagnosis was confirmed by histo-pathological examination.

The patients were grouped according to different variables like age, sex, size of the nodule, site of the nodule, functional thyroid status, FNAC reports and histo-pathological examination reports, then analyzed and compared with the previous similar studies conducted elsewhere. Finally, conclusions were drawn accordingly.

RESULTS

Total of 50 cases of solitary nodule of thyroid studied and following conclusions were drawn:

Age Incidence:

The age of the patients ranges from 18 years to 66 years, with peaks being in 3^{rd} to 5^{th} decades. The mean age of presentation is 38.5 years. Cases in 3^{rd} to 5^{th} decades constitutes 72% of the cases studied.

Age in Years	No. of Patients
0-9	0
10-19	2
20-29	13
30-39	9
40-49	14
50-59	9
60-69	3
TOTAL	50



Sex Incidence:

Solitary nodule of thyroid are much more common in females. Out of 50 cases studied 46 were females and 4 were males, and the ratio comes to M:F

= 1 : 11.5. Also the malignant nodules are common in females. Out of 7 cases of malignancy in the study, 6 were females.

Sex	no. of patients
Males	4
Females	46
Total	50



Clinical Features:

All the cases in the present study presented complaint of swelling in the region of the thyroid. Only few patients presented with pain, discomfort and dysphagia. All the mentioned additional symptoms were of mild degree. Out of 50 cases, 3 cases had pain, 3 cases had discomfort and another 2 had dysphagia. Also none of the patient had lymphadenopathy which was confirmed by ultrasonographic examination. Two patients had symptoms of thyrotoxicosis, and one had features of hypothyroidism. The latter patients' thyroid profile confirmed the functional status.

Duration of symptoms:

In our study, duration of onset symptoms varied from 15 days to 8 years. Also, duration of malignant nodules extend from 1 month to 4 years.

Duration of Symptoms	No. of patients	
<1mon	1	
1-3 mon	7	
3-6 mon	10	
6-12 mon	7	
1-2yrs	12	
2-5yrs	11	
>5yrs	2	



Site of the nodule:

Out of 50 cases studied, 26 cases presented with nodule in right lobe of the thyroid gland and the remainder in the left lobe of thyroid. One patient among left sided solitary nodule had undergone right lobectomy 30 years back and presented with recurrent nodule in the rest of the lobe.

Site of the nodule	no. of patients	
Right	28	
Left	22	
Total	50	

Size of the nodule:

In the present study, on clinical examination size of the nodule, in its largest dimension, varies from 2cm to 12cm. Most of the patients presented with the size of about 3 to 5 cm. in the study, as such there is no correlation between the size of the nodule and the occurrence malignant nodule.

Size of the Nodule	No. of patients
<1 cm	0
1-2 cm	2
2-3 cm	8
3-4 cm	15
4-5 cm	13



Thyroid functional status:

Out of 50 cases, two presented with features of thyrotoxicosis, one with hypothyroidism and rest all were in euthyroid state. Patients with thyrotoxicosis were made euthyroid using antithyroid drugs and operated and both cases turned out to be toxic follicular adenoma. Patient with hypothyroidism was treated with thyroxine, USG neck revealed multiple nodules and managed by subtotal thyroidectomy, histopathological examination confirmed the diagnosis of multi-nodular goiter.

© 2023 SAS Journal of Surgery | Published by SAS Publishers, India

Thyroid functional status	No. of patients	
Euthyroid	46	
Hyperthyroid	3	
Hypothyroid	1	
Total	50	



FNAC Reports:

Fine Needle Aspiration Cytology is the important investigation in the evaluation of solitary nodule of thyroid. All 50 cases were subjected to FNAC during the course of evaluation. FNAC reports are mainly categorized into 6 entities- Benign, follicular neoplasm, suspicious (of malignancy), malignant, lymphocytic thyroiditis, cysts. In our study, out of 11 follicular neoplasms, two turned out to be follicular carcinoma. One suspicious (of papillary carcinoma) case confirmed papillary carcinoma on histopathological examination. Four cases of papillary carcinoma were diagnosed pre- operatively by FNAC alone.

Three cases diagnosed as cysts by FNAC confirmed to be simple cysts on histopathological examination.

FNAC reports	No. of patients
Benign	30
Follicular Neoplasm	11
Suspicious	1
Malignant	4
Lymphocytic Thyroiditis	1
Cysts	3



81

Aetiological incidence of solitary nodule of thyroid:

Out of 50 cases studied, common causes of solitary nodule are MNG, follicular adenoma and adenomatous goiter; the most common being MNG which constitutes about 34% of cases. Follicular adenomas have 22% and adenomatous goiters have 24% incidences.

Out of 50 cases, seven were malignant -5 papillary carcinoma and 2 follicular carcinomas. Ultrasonography detected suspicious findings in three

cases among seven malignant cases -2 papillary and 1 follicular.

Three cases of papillary carcinoma were diagnosed with certainty by FNAC, one case was suspicious which turned out to be papillary CA on histopathological examination.

Two cases of follicular carcinoma were diagnosed follicular neoplasm, one of them showed suspicious features on ultrasonographic examination.

HPE Reports	No. of patients
Follicular adenoma	11
Adenomatous Goitre	12
MNG	17
Carcinoma	7
Lymphocytic thyroiditis	1
Simple cyst of Thyroid	2
Total	50



Type of carcinoma:

From the study, out of 7 carcinomas, 5 were papillary and 2 follicular: no case of medullary or

anaplastic or lymphoma was detected. Papillary carcinoma accounts to 71% and follicular carcinoma accounts to 29%.

Carcinoma	No of cases	Percentage
Papillary	5	71
Follicular	2	29
medullary	0	0
anaplastic	0	0
lymphoma	0	0
Total	6	100



Surgery / operative procedure done:

Depending upon the clinical diagnosis and FNAC features, all the 50 patients undergone surgery. Among them, 25 patients had undergone hemithyroidectomy, 12 cases undergone sub-total thyroidectomy and 13 cases undergone total thyroidectomy.

Post-operatively, suppressive dose of thyroxine was started for patients who had undergone total thyroidectomy. Three cases out of 13 cases of total thyroidectomy showed features of hypocalcemia on 2-4 post-operative day, hence, they are supplemented with oral calcium and vitamin D3.

All the cases were followed up for 6 months, two cases had husky voice without any change in vocal cord movements.

DISCUSSION

The observations and results of the present study were compared with the available previous similar studies.

MEAN AGE AT PRESENTATION:

Authors	Mean Age In Years
Das DK (1999) [31]	35
Talepoor M (2005) [32]	38.6
Quari F. (2005) [33]	36.7
Rehman A.U. (2009) [34]	34.7
Khurshid Anwar (2012) [35]	37
Present study	38.5

In the study done by Quari F and Talepoor M separately in 2005, reported the mean age at presentation as 36.7 years and 38.6 years respectively. Khurshid Anwar reported, in 2012, the mean age of presentation as 37 years. From the present study, the mean age at presentation found to be 37.27 years, correlates with the previous studies.

Most of the earlier series reported peak incidence of solitary nodule thyroid in the 3^{rd} and 4^{th} decades. Bhansali S. K (1982) [36], in his similar study, reported the peak incidence in 4^{th} and 5^{th} decade. In the present study, the peak incidence found to be 3^{rd} to 5^{th} decades, which constitutes about 70% of the cases studied.

Sex Distribution:

Authours	Sex Incidence (M: F)
Dorairajan (1996) [26]	1:9
Das DK (1999) [31]	1:5.39
Gupta C (2001) [37]	1:5
Present study	1:11.5

In the study done by Dorairajan (1996) and Das DK (1999) reported ratio of sex incidence as 1:9 and 1:5.39 respectively. In the present study, it's found to be 1:11.5, which correlates with previous studies.

Because of periods of fluctuations in the demands of the hormonal requirement in female in their

life cycle (puberty, menstrual cycles, pregnancy, menopause), the chances of thyroid nodule formation are very high as compared with male counterparts.

Distribution of non-neoplastic and neoplastic lesions diagnosed by FNAC:

Authours	Non- Neoplastic	Neoplastic	Ratio
Sarda AK (1997) [39]	487	59	8.25:1
Das DK (1999) [31]	346	85	4.07:1
Gupta C (2001) [37]	470	30	15.66
Talepoor M (2005) [32]	325	70	4.33:1
Hurtado Lopez M (2005) [38]	80	50	1.6:1
Nagada (2006)	51	18	2.83:1
Chao CT (2007)	276	264	1.04:1
Present study	34	16	2.12:1

In the present study, neoplastic conditions include adenomas and all malignant lesions. From the study, the ratio of non-neoplastic to neoplastic cases is about 2.12:1, which is comparable to the studies done

earlier like Karur (2002), Hurtado Lopez M (2005), Nagada (2006), Chao CT (2007).

Distribution of malignancies by FNAC:

Authours	Percentage
Sarda A. K et al., (1997)	10.8
Karur K et al., (2002)	18
Mundsad B et al., (2006) [40]	4.16
Present study	8

In the present study, among 5 cases of papillary CA, 4 were diagnosed with certainty by FNAC and the rest one was suspicious of malignancy. But both the follicular CA were initially reported as follicular neoplasm. From the study, distribution of malignancy is about 8, which is comparable with the earlier studies.

Aetiological incidence (in percentage):

Series	MNG	Adenoma Follicular	Carcinoma	Others	Total No. of cases
Zaman & Bhagbati (1971)	83	9	8	-	2221
Ananth Krishnan (1983)	12	47	2	2	104
Bhansali (1982)	71	20	9	-	449
Fenn (1980)	22	55	12	11	342
Kapur (1982)	28	50	11	11	221
Present series	34	22	14	30	50

From the present study, commonest cause of solitary nodule is MNG, which is comparable with the studies done by Fenn (1980), Kapur (1982), Bhansali (1982). The common causes are follicular adenoma and

adenomatous goiter.

Incidence of carcinoma:

Study	Year	Percen tage
C.Leigh et al., [41]	1969	20.9%
A S Fenn et al., [42]	1980	12.0%
Bhansali S K	1982	9.0%
Kapur <i>et al.</i> ,	1982	11.0%
A K Sarda et al., [39].	1997	10.8%
Mazafferi et al., [43]	1998	11-12%
Wagana <i>et al.</i> ,	2002	16%
G.A Khairy et al., [44]	2004	13.9%
Talepoor et al., [32]	2005	15.8%
CatrherineIhre Lundgreen [45]	2007	20.9%

© 2023 SAS Journal of Surgery | Published by SAS Publishers, India

Study	Year	Percen tage
Judy Jin <i>et al.</i> , [46]	2009	15%
Rehman A U [34]	2009	11.4%
Salim Ahmed et al., [47]	2011	12.3%
Md. Abul Hossain et al., [48]	2014	28%
Rameshbabu et al., [49]	2015	10.83%.
Fernando et al., [50]	2015	12%
Present study	2018	14.0%

From the literature, the incidence of malignancy in thyroid nodule ranges from 5% to 30%. From the present study, the incidence found to be 14%, which is comparable with the previous studies.

CONCLUSIONS

The present study is a prospective analysis of 50 cases of solitary nodule of thyroid, admitted in KIMS medical college. Though a large number of patients are required to come to better conclusions, based on the data and results obtained in the present study, the following conclusions can be drawn:

- Solitary nodule of thyroid is more common in females.
- Solitary nodule of thyroid is more common the age group of 20-50 years.
- Most of the patients with solitary nodule of thyroid present with swelling alone.
- Most of the patients with solitary nodule of thyroid are in euthyroid state and only few present with toxicity and hypothyroidism.
- Incidence of malignancy in male patients presenting with solitary nodule thyroid is more when compared to female patients presenting with the same.
- Commonest cause of solitary nodule of thyroid is multi-nodular goitre.
- USG can be used to detect multi-nodular goitre in patients presenting with solitary nodule thyroid.
- FNAC is the investigation of choice in the evaluation of solitary nodule of thyroid. It has few pitfalls. In such situations, only histopathology can confirm the exact pathology. It detects papillary carcinoma in a solitary nodule with high sensitivity and specificity.
- Papillary carcinoma is the most common malignancy of thyroid, followed by follicular carcinoma.

REFERENCES

- Zygmunt, H. K. The thyroid and the thyroglossal tract. Chapter 53 in Bailey and Love's short practice of Surgery, 24th edition, p 771.
- Cole, W. H., Majarakis, J. D., & Slaughter, D. P. (1949). Incidence of carcinoma of the thyroid in nodular goiter. *The Journal of Clinical Endocrinology*, 9(10), 1007-1011.
- 3. Ananthakrishnan, N., Rao, K. M., Narasimhan, R.,

© 2023 SAS Journal of Surgery | Published by SAS Publishers, India

Veliath, A. J., Smile, S. R., & Jagadish, S. (1993). The single thyroid nodule: a south Indian profile of 503 patients with special reference to incidence of malignancy. *Ind J Surg*, *55*(10), 487-492.

- 4. Vander, J. B., Gaston, E. A., & Dawber, T. R. (1968). The significance of nontoxic thyroid nodules: final report of a 15-year study of the incidence of thyroid malignancy. *Annals of internal medicine*, *69*(3), 537-540.
- Ezzat, S., Sarti, D. A., Cain, D. R., & Braunstein, G. D. (1994). Thyroid incidentalomas: prevalence by palpation and ultrasonography. *Archives of internal medicine*, 154(16), 1838-1840.
- Paul, R. M., Malcolm, H. W. (1997). Approach to thyroid nodules. Chapter 9, textbook of Endocrine Surgery, Orlo H. Clark, Philadelphia, WB Saunders; p688.
- 7. Mazzaferri, E. L. (1993). Management of solitary thyroid nodule. *N Engl J Med*, 328 553-559.
- Beaugie, M. J. (1975). Principles of thyroid surgery. Kent: Pitman Medical Publishing Company, pp2.
- Chaurasia, B. D. (2004). Deep structures in neck. Chapter 12 in Human anatomy regional and applied, 4th edition: CBS publishers, 165-171.
- Fielding, H. G. (1960). An Introduction to History of Medicine, 4th edition, WB Saunders Company; pp 554-563, 695, 729, 730-872.
- John, B. H. (2001). Thyroid Chapter 32, Sabiston Textbook of Surgery, 16th edition. Townsend Jr. Harcourt Asia Pvt. Ltd, India, pp 603-628.
- 12. Sing, I. (1996). Human Embryology. The Pharyngeal Arches. Chapter 9, 6th edition, McMillan India Ltd., pp 119-122.
- 13. Decker Gag, Plessus, D. J. D. (1995). The thyroid Thymus and parathyroid glands Chapter 17, Lee Mc Gregors Synopsis of Surgical Anatomy; 10th edition, Verghese Publishing House, Bombay, India; pp 198-205.
- Dyson, M. (2000). Endocrine system Thyroid gland, Chapter 15, Williams Peter L. Gray's Anatomy. 38th edition Churchill Livingstone London, UK, pp 1891-1897.
- Ellis, H. (1997). The Head and Neck, Parts of Clinical Anatomy; 9th edition, Blackwell Science, UK; pp 284-287.
- Sinnathambi Chummy, S. (1999). Head and Neck and Spine. Chapter 6, Lasts Anatomy, 10th edition, Churchill Living stone, London UK, pp 330-332.
- 17. Williams, A. C. (1962). Thyroid cancer in 1330 cases of sporadic goitre. Am J Surg, 104: p 672-

676.

- Guyton Arthur, C., & Hall, J. (2001). The Thyroid Metabolic Hormones, Chapter 76, Textbook of Medical Physiology, 10th edition, Harcort Asia Pvt. Ltd. India, pp 858-868.
- Gould, E. A., Hirsch, E., & Brecher, I. R. A. (1965). Complications arising in the course of thyroidectomy. *Archives of Surgery*, 90(1), 81-85.
- 20. Burch, H. B. (1995). Evaluation and Management of Solitary Thyroid Nodule in Thyroid Cancer I, Burman, Kenneth D, Philadelphia, WB Saunders, Endocrinology and Metabolism. Clinics of North America, 24(4), 668.
- Woolner, L. B., Beahrs, O. H., Black, B. M., McConahey, W. M., & Keating Jr, F. R. (1961). Classification and prognosis of thyroid carcinoma: A study of 885 cases observed in a thirty year period. *The American Journal of Surgery*, 102(3), 354-387.
- 22. Das, S. Examination of thyroid gland Chapter 27, Manual of Clinical Surgery, 5th edition, Dr. S. Das, Calcutta, India, pp 284-297.
- Lumley, J. S. P. (1997). The thyroid and parathyroids. Chapter 17, Hamilton Baileys Physical Signs. 18th edition, Butterworth Heinemann, Oxford UK, pp 223-226.
- Matheson, M. A. (1986). Diagnosis of Thyroid Swelling. Recent Advances in Surgery, Russel RCG. Vol 12, Churchill Livingstone, UK; PP 179-197.
- Bouvet, M., Feldman, J. I., Nahum, A. M., Robbins, K. T., Gill, G. N., Dillmann, W. H., & Russack, V. (1992). Surgical management of the thyroid nodule: Patient selection based on the results of fine-needle aspiration cytology. *The Laryngoscope*, 102(12), 1353-1356.
- 26. Dorairajan, N., & Jayashree, N. (1996). Solitary thyroid nodules and role of fine needle aspiration cytology in diagnosis. *J Indian Med Assoc*, 94(2), 50-52.
- Russel, C. F. J. (1994). Management of solitary thyroid nodule. Recent advances in surgery. CD Johnson, I Taylor (ed), 17(1), p1-6.
- Joseph, N. A. (1998). Crucial controversies in surgery. Mosch Schien, Brooklyn N: (ed) Karger Landes system (B), 215-324.
- Shirish, K. B. (2002). Current management of Solitary nodule of thyroid, Roshanlal Gupta, 8, 197-204.
- 30. Das, S. (2001). The thyroid and parathyroids. Chapter 37, In: A concise textbook of surgery, 3rd edition, Dr. S. Das publisher: p648-670.
- Das, D. K., Khanna, C. M., Tripathi, R. P., Pant, C. S., Mandal, A. K., Chandra, S., ... & Thusoo, T. K. (1999). Solitary nodular goiter. Review of cytomorphologic features in 441 cases. *Acta cytologica*, 43(4), 563-574.
- Talepoor, M., Karbankhsh, M., & Mirzali, F. A. (2005). Management of Solitary Thyroid Nodules: The Dilemma of Multinodular Goiter as False-

Positive Cases. Medicine On-Line; 2005, January. Available from: http://www.priory.com/ med/thyroidnodule.htm

33. Quari, F. (2005). Unnecessary Tests and Delay in the Diagnosis of Solitary Thyroid Nodules at the University Hospital. April; 2005. Available from: http://

www.bhj.org/journal/april2005/htm/original_unnec essary_138.htm

- Rehman, A. U., Lodhi, S., & Anwar, M. I. (2009). Histopathological evaluation of 432 cases of goiter. *Annals of King Edward Medical* University, 15(2), 54-54.
- Anwar, K., Din, G., Zada, B., & Shahabi, I. (2012). Single center study. *J Postgrad Med Inst*, 26(1), 96-101.
- Bhansali, S. K. (1982). Solitary nodule in the thyroid gland; experience with 600 cases. *Ind J* Surg, 44, 547-61.
- Gupta, C., Sharma, V. K., Agarwal, A. K., & Bisht, D. (2001). Fine needle aspiration cytology of solitary nodule of thyroid and its histopathological correlation. *Journal of cytology*, *18*(3), 151-156.
- 38. Hurtado-López, L. M., Arellano-Montaño, S., Torres-Acosta, E. M., Zaldivar-Ramirez, F. R., Duarte-Torres, R. M., Alonso-de-Ruiz, P., ... & Martínez-Duncker, C. (2004). Combined use of fine-needle aspiration biopsy, MIBI scans and frozen section biopsy offers the best diagnostic accuracy in the assessment of the hypofunctioning solitary thyroid nodule. *European journal of nuclear medicine and molecular imaging*, 31, 1273-1279.
- Sarda, A. K., Gupta, A., Jain, P. K., & Prasad, S. (1997). Management options for solitary thyroid nodules in an endemic goitrous area. *Postgraduate medical journal*, 73(863), 560-564.
- 40. Mundasad, B. M., Mcallister, I., Carson, J., & Pyper, P. C. (2006). Accuracy of fine needle aspiration cytology in diagnosis of thyroid swellings. In: The Internet Journal of Endocrinology 30th July; 2006. http://www. ispub.com/ostia/index.php?xmlfilepath=journals/ije n/vol2n2/fna.xml.
- 41. Kendall, L., & Condon, R. (1969). Prediction of malignancy in solitary thyroid nodules. *The Lancet*, 293(7605), 1071-1073.
- 42. Fenn, A. S. (1980). Solitary nodule of thyroid gland. *Ind J Surg*, 42, 171-175.
- Messaris, G. K. P. C., Kyriakou, K., Vasilopoulos, P., & Tountas, C. (1974). The single thyroid nodule and carcinoma. *British Journal of Surgery*, 61(12), 943-944.
- 44. Khairy, G. H. (2004). Solitary thyroid nodule: the risk of cancer and the extent of surgical therapy. *East African medical journal*, 81(9), 459-462.
- 45. Lundgren, C. I., Stalberg, P., Grodski, S., Sidhu, S., Sywak, M., & Delbridge, L. (2007). Minimally invasive thyroid surgery for diagnostic excision of

solitary thyroid nodules. *Asian journal of surgery*, 30(4), 250-254.

- Jin, J., Wilhelm, S. M., & McHenry, C. R. (2009). Incidental thyroid nodule: patterns of diagnosis and rate of malignancy. *The American journal of surgery*, 197(3), 320-324.
- Soomro, S. A., Memon, M. R., Abro, H., & Mahar, M. (2011). Frequency of Malignancy in Cold Nodule Thyroid". *Medical Channel*, *17*(1), 44-49.
- 48. Hossain, M. A., Sarkar, M. Z., Dutta, U. K., Karim, M. A., & Alam, M. Z. (2014). Frequency of

Malignancy in solitary Thyroid nodule and Multinodular Goitre. *Bangladesh Journal of Otorhinolaryngology*, 20(2), 55-59.

- 49. Ramesh, B. M. (2015). Malignant incidence in solitary nodule thyroid a clinical study, *IJAR*, 5
- Fernando, J. R., Raj, S. E. K., Kumar, A. M., & Anandan, H. (2017). Clinical study of incidence of malignancy in solitary nodule of thyroid. *International Journal of Scientific Study*, 5(4), 232-236.