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## ACCURACY OF ULTRASOUND GUIDED FINE NEEDLE ASPIRATION CYTOLOGY IN THE EVALUATION OF THYROID NODULES

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#### ABSTRACT

**Background:** Thyroid nodules are a common problem which requires an accurate cytological diagnosis for proper management. An essential investigation modality is fine needle aspiration cytology (FNAC) with a sensitivity ranging between 80-90%. Recommendations vary regarding the use of ultrasound in improving the accuracy of free hand fine needle aspiration biopsy.

**Objective:** The aim of our study was to investigate the role and accuracy of ultrasound guided fine needle aspiration cytology in the evaluation of patients with thyroid nodules.

**Methods:** Forty four patients with nodular thyroid disease who were assessed by ultrasound guided fine needle aspiration cytology (US-FNAC) followed by thyroidectomy were identified. Retrospective analysis of data on the cytology specimens and histological results was carried out.

**Results:** The sensitivity and specificity of US-FNAC for diagnosing thyroid neoplasms were 91.7% and 93.3% respectively. The diagnostic accuracy was 92.8%, with the rate of inadequate smears as 4.6%. **Conclusions:** Ultrasound guided fine needle cytology of thyroid nodules yields more accurate results, with a decrease in the number of inadequate sample category. Precise targeting of lesions in both palpable and non palpable thyroid nodules by the FNA needle under ultrasound guidance will increase the diagnostic accuracy, thereby avoiding or reducing the number of unnecessary thyroid surgeries.

**Keywords:** thyroid nodule, ultrasound guided, FNAC, neoplasm

#### INTRODUCTION

Nodular thyroid disease found is in approximately 5% of the population at palpation screening. Routine ultrasound screening of thyroid in clinically normal individuals suggests a prevalence rate of 50% for thyroid nodules<sup>1, 2</sup>. Fine needle aspiration cytology (FNAC) of the thyroid gland is over 50 years old<sup>3</sup> and is the single most important method of establishing a diagnosis in both adults and children. Recent guidelines published by the British Thyroid Association/Royal College of Physicians and the American Association of Clinical Endocrinologists (AACE)/Association Medici Endocrinologi(AME) emphasize that FNAC should be considered as the early investigation of choice for thyroid nodules and specifically for thyroid cancer<sup>4,5</sup>.

FNAC requires careful aspiration technique and meticulous interpretation of the findings by a skilled cytologist. Studies have suggested a sensitivity and specificity for predicting thyroid 83% malignancies averaging and 92% respectively<sup>5, 6-9</sup>. Unfortunately, FNAC can miss malignancies with a false-negative rate of approximately 5%<sup>5</sup>. Current guidelines recommend the usage of ultrasound guided FNAC (US-FNAC) to reduce this error<sup>1,5</sup>.

Studies have shown that US-FNAC significantly reduces the number of inadequate fine-needle aspirates and improves the diagnostic yield for malignancies<sup>1,10</sup>. US-FNAC also improves the yield of cancer found at surgery<sup>11</sup>.

The aim of our study was to investigate the role and accuracy of ultrasound guided fine needle aspiration cytology in the management of patients with thyroid nodules.

### PATIENTS AND METHODS

Two hundred and eighty two FNACs of thyroid nodules were carried out at Mahatma Gandhi Medical College and Research Institute and the results reviewed. Out of this, 44 patients had US-FNAC for assessment of thyroid nodules followed by thyroidectomy. A retrospective analysis of the data was carried out that included patient demographics, US-FNAC results and the final histopathological reports. All patients had baseline thyroid function tests followed by ultrasound examination and guided FNAC of the thyroid nodules. A high resolution 7.5 MHz probe was used to guide a 23/25 gauge needle attached to a syringe to obtain cellular aspirates from the lesions. Cytological diagnosis was correlated with histopathology and the accuracy of ultrasound guided FNAC in diagnosing thyroid neoplasms was calculated.

### **Study parameters:**

- Demographic data
- Sensitivity, specificity
- Positive and negative predictive values
- Diagnostic accuracy rate
- Inadequate sample category

### RESULTS

Data regarding 44 patients who had undergone US-FNAC of the thyroid nodules followed by thyroidectomy was retrospectively analyzed. There were 3 men and 41 women in the study group, with age ranging from 18 to 51 years. The mean age was 39.6 years with male to female ratio of 1:13.7(Table 1). 29(63.6%) patients had non-neoplastic lesions (nodular/colloid goiter, thyroiditis) and 14(31.8%) had neoplastic lesions on cytological examination. Overall, US-FNAC yielded satisfactory cell samples in 42(95.4%) patients and inadequate material in 2(4.6%) patients (Table 2). An aspirate was considered adequate by the presence of 5 to 6 clusters of cells with each cluster containing 10 or more cells.

Nodular goiter constituted the majority (55.2%) in the non-neoplastic category and papillary carcinoma was the commonest (61.5%) cytological diagnosis in the neoplastic group as shown in Tables 3 and 4. On comparing the US-FNAC reports of the 29 non-neoplastic nodules with the final histological diagnosis, it was observed that 26 patients had same cytological diagnosis, 1 had papillary carcinoma (false negative) and 2 had Hashimoto's thyroiditis (Table 5).

The US-FNAC results for the 13 neoplastic lesions were compared with the corresponding histological diagnoses. 8 patients had papillary carcinoma confirmed by histological study and out of 4 cases of follicular neoplasm, 2 were confirmed as follicular adenoma and 1 each of colloid goiter (false positive) and follicular carcinoma. One patient diagnosed as Hurthle cell neoplasm by US-FNAC turned out to be Hashimoto's thyroiditis on final histological study (false positive), Table 6.

The sensitivity and specificity of US-FNAC for diagnosing neoplastic lesions were 91.7% and 93.3% respectively. The diagnostic accuracy was 92.8%, positive predictive value (PPV) was 86.5% and negative predictive value (NPV) 96.5% (Table 7 and Table 8).

## DISCUSSION

FNAC is considered as the gold standard in the evaluation of thyroid nodules. Major problems associated with free hand FNAC are significant false negatives (missed neoplasms) and difficulties in accurate identification of follicular lesions leading to wrong FNAC diagnoses<sup>12</sup>. Possible reasons cited for this include the technique of aspiration, smear preparation and cytopathologist. experience of the The AACE/AME has suggested that ultrasound should be employed routinely to detect features suspicious of malignancy in thyroid nodules and to identify lesions that would be suitable for  $FNAC^{5}$ . Ultrasound (US) is useful to guide the needle for fine needle aspiration of a thyroid nodule; it improves the quality of diagnosis in pre operative assessment of thyroid nodules and to appropriately select patients for thyroidectomy. Ultrasound guidance allows continuous visualization of the needle during insertion and sampling, which results in pinpoint accuracy with a high - level of safety. Some of the potential advantages of US-FNAC in the evaluation of nodular thyroid disease are selecting the suspicious nodule for biopsy in a multinodular goiter and guiding the needle to take samples from nonpalpable nodules.

A recent review of studies that evaluated the role of US-FNAC in the detection of thyroid cancer revealed a sensitivity of 76%-98%, specificity of 71%-100%, false-negative rate of 0%-5%, falsepositive rate of 0-5.7%, and overall accuracy of 69%–97%<sup>13-23</sup>. Another report based on a systematic review of 12 studies had shown a median sensitivity and specificity of 88% and 90.5%<sup>24</sup>. A large prospective cohort study had reported a marked increase in the diagnostic accuracy of FNAC from 85% to 95% when FNAC was combined with US guidance<sup>25</sup>. Our results have shown a sensitivity and specificity of US-FNAC for diagnosing neoplastic lesions as 91.7% and 93.3% respectively. The false negative rate was 2.3%, PPV 86.5%, NPV 96.5% and the diagnostic accuracy was 92.8%.

The accuracy of FNAC depends crucially on the technique, operator performing the aspiration and the cytopathologist analyzing it. Even under optimal conditions, the false negative rate for thyroid neoplasms can vary from 1% to 6% due

to wrong diagnosis or sampling errors<sup>26</sup>. Danese et al in their study have observed the false negative rate much lower for US-FNAC when compared with free hand FNAC<sup>27</sup>. In our analysis, there was one patient who had cytological picture of nodular goiter whereas the histological study showed a small focus of papillary carcinoma in a background of multinodular goiter(false negative - 2.3%).

The majority of thyroid FNAC is now performed under US guidance and this has become mandatory at some of the centers. The major benefit of this technique is accurate sampling of small or multiple nodules<sup>28-30</sup>. Certain sonologic features of thyroid lesions are predictive of malignancy and hence such lesions can be accurately biopsied<sup>31</sup>. Another crucial benefit of using US-FNAC is significant reduction in the number of inadequate aspirates<sup>32</sup>. Case series have reported rates of inadequate samples in the order of 15-30% for free hand FNAC. Many published studies have shown US-FNAC to decrease the rate of inadequate samples from 15% to 3% when compared to free hand FNAC<sup>17,33,34</sup>. In a recent study, the authors have noted a prevalence rate of 8.5% for thyroid cancer in patients with inadequate samples and they recommended repeat aspirations by US-FNAC technique<sup>35</sup>. It is possible that reduced number of inadequate aspirates may potentially lead to increase in the yield of cancer and the diagnostic accuracy of FNAC. The rate of inadequate specimens in our retrospective study was 4.5 % (2 out of 44 samples), which is comparable to the above quoted studies.

### CONCLUSION

In conclusion, US-FNAC greatly facilitates the precise and adequate sampling of tissues from thyroid lesions thus enabling the cytopathologist to make an accurate diagnosis. This may lead to the elimination of the cost and potential morbidity of unnecessary thyroidectomy.

Ultrasound guidance helps in the selection of the most suspicious focus within the nodule especially with a background of multinodularity. It is therefore desirable that practice guidelines recommend universal use of ultrasound guided FNAC for all nodular thyroid lesions, especially for inadequate smears from free hand FNAC, for small or nonpalpable nodules and multinodular goiters. US- FNAC is the most accurate and costeffective method for diagnostic evaluation of thyroid nodules.

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Table	1:	Baseline	characteristics	of	the	patients
(n=44)						

Age (mean in years)	36.9
Men (%)	3(6.8)
Women (%)	41(93.2)
Male: Female	1:13.6
Solitary nodule (%)	32(72.7)
Multinodular goiter (%)	12(27.3)

## Table 2: Distribution of thyroid lesions based on cytological study

Lesions	No. of Patients	Percentage	
Non-neoplastic	29	65.9	
Neoplastic	13	29.5	
Inadequate	2	4.6	
Total	44	100	

#### Table 3: US-FNAC report of non-neoplastic lesions

Cytological report	No. of Patients	Percentage
Colloid goiter	13	44.8
Nodular goiter	16	55.2
Total	29	100

#### Table 4: US-FNAC report of neoplastic lesions

Cytological report	No. of Patients	Percentage
Papillary carcinoma	8	61.5
Follicular neoplasm	4	30.8
Hurthle cell neoplasm	1	7.7
Total	13	100

## Table 5: Correlation of US-FNAC with

Histopathological (HPE) report for non-neoplastic nodules

Cytological	Histopathological report				
report	Nodular/coll	Papillary	Hashimoto's		
report	oid goiter	carcinoma	thyroiditis		
Nodular/					
Colloid	26	1	2		
goiter (29)					

#### Table 6: Correlation of US-FNAC and HPE for neoplastic lesions

	Histopathological report					
Cytological report	Papillary	Follicular	Hashimoto	Follicular	Colloid	
	carcinoma	adenoma	thyroiditis	carcinoma	nodule	
Papillary carcinoma(8)	8	-	-	-	-	
Follicular neoplasm(4)	-	2	-	1	1	
Hurthle cell neoplasm(1)	-	-	1	-	-	

# Table7:ComparisonofUS-FNACwithhistopathological diagnosis for neoplasms

FNAC	Histopa (neop	Total	
(neoplasm)	Positive	Negative	
Positive	11	2	13
Negative	1	28	29
Total	12	30	42

True Positives – 11 False Negatives – 1 False Positives- 2 True Negatives – 28

# Table 8: Statistical parameters for US-FNACdiagnosis of neoplastic lesions

Parameters	Value
Sensitivity	91.7
Specificity	93.3
Positive predictive value	84.6
Negative predictive value	96.5
Diagnostic accuracy	92.8