The Value of Ultrasonography Using Thyroid Imaging Reporting and Data Systems (TIRADS) in the Diagnosis of Thyroid Cancer Among the Population of Ha’il, Saudi Arabia

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Abstract

Background: Thyroid cancer is one of the common malignant conditions of the head and neck region, and it is considered as one of the most common cancers among endocrine tumors. Ultrasonography is widely used in order to assess thyroid nodules, Therefore, the aim of our study is to determine the accuracy of ultrasonography and fine needle aspiration biopsy (FNAB)-guided by ultrasonography in the diagnosis of thyroid malignancy among the population in the Ha’il region of Saudi Arabia.

Methods: A retrospective evaluation of 137 patients was undertaken at King Salman Specialist Hospital, Ha’il, Kingdom of Saudi Arabia. Patients who presented with thyroid swellings underwent ultrasonography and FNAB.

Result: Our study results have concluded that the sensitivity of the ultrasonography in the confirmation of a malignant thyroid lesion is 59.4% and its specificity was found to be 74.3%.

Conclusion: The study showed that ultrasonography using Thyroid Imaging Reporting and Data Systems (TIRADS) is a significant step in evaluating a thyroid nodule. Also, it is highly sensitive and specific, cost-effective, and convenient for the patient.

Introduction

Thyroid cancer is the most common form of endocrine cancer worldwide, with an incidence of 56 cases per 100,000 [1]. Thyroid cancer is the fifth most common cancer in females and the second most common in women older than 50 years [2]. The incidence of thyroid cancer worldwide has increased at a higher rate than any other cancer in the past several decades, while mortality rates for thyroid cancer have remained almost stable. One of the most important risk factors is exposure to ionizing in some environments, along with female gender, age, and genetics [3,4]. The clinical manifestations of thyroid cancer include thyroid nodule, difficulty in swallowing, difficulty with breathing, and hoarseness of voice [5].

For investigation, ultrasound imaging and fine-needle aspiration cytology is needed. The features of ultrasonography that are suspicious of thyroid cancer include the presence of marked hypo-echogenicity, micro-lobulated or irregular margins, micro-calcifications, and a taller-than-wide shape [6].

Thyroid cancer is diagnosed histologically through fine needle aspiration biopsy (FNAB) and is categorized into four types: (i) papillary thyroid carcinoma, which is the most common type of thyroid cancer and is the least aggressive type of cancer, because it grows and metastasis slowly; (ii) follicular thyroid carcinoma, which is almost 14% of thyroid malignancies, is more aggressive than papillary thyroid carcinoma; (iii) medullary thyroid carcinoma, a cancer of non-thyroid cells that are normally present in the thyroid gland, medullary thyroid cancer is almost 3% of thyroid malignancies; (iv) anaplastic thyroid cancer, which is almost 2% of thyroid cancers and is the most dangerous type of thyroid malignancies because it metastasizes early to the surrounding lymph nodes and also distant organs [5].

FNAB is currently the gold standard for evaluating and assessment of thyroid nodules that could be cancerous [3].
Materials And Methods
This study is a retrospective evaluation of a prospectively maintained data set of patients at the King Salman Specialist Hospital, Ha'il, Saudi Arabia. Data were collected from the patient’s medical records and reviewed. This data collection process was focused on patients who were admitted to the hospital with thyroid-related diseases or outpatients who visited the hospital with similar complaints. In this study, the data collected was evaluated to determine the accuracy of ultrasonography compared to FNAB guided by ultrasonography in the diagnosis of thyroid malignancy among the population in the Ha'il region in the two-year period of 2019-2020.

The authors retrospectively evaluated the electronic medical records of 137 patient subjects during the study period of October to November, 2021, in King Salman Specialist Hospital. Of the subjects, 123 were females and 14 were males. Furthermore, an assessment of the health comorbidities was done to better evaluate the subjects. The last section of the study was designed to study and analyze the outcomes of the ultrasonographical method for evaluating a thyroid lesion along with the reports of the histopathology to later find out the sensitivity and the specificity of the ultrasound in confirming thyroid malignancy. A chi-square non-parametric test was implemented to find the statistical significance of the study categorical variables, and a p-value of <0.001 was accepted to be significant.

Imaging evaluation
An ultrasound session was conducted on each patient suspected to have a thyroid lesion after a complete clinical examination. Each of the subjects had an evaluation of whether the suspected thyroid lesion is considered benign or malignant, and a TIRADS score was given to each patient after reporting the findings.

Histopathological evaluation
Further, fine needle aspiration of the thyroid was also done for histopathological evaluation of the thyroid lesion after the ultrasonography for each subject to ultimately confirm the diagnosis of either a benign condition or a malignancy.

Results
Sociodemographic data
A total of 137 patient subjects’ electronic medical reports were reviewed and evaluated. Of the total, the majority were females (n= 123, 89.8%) and only 14 (10.2%) were males. The different comorbidities associated with the patients’ presentation were observed and it was found that 43.8% were medically free, 27.0% had diabetes, 16.8% had hypertension, and 15.3% had hypothyroidism. Other medical conditions that were found in lesser numbers included asthma, Hodgkin’s lymphoma, mammary duct ectasia, and previous medical history of atrial fibrillation. Table 1 describes the baseline sociodemographic information and the associated comorbidities.

| Gender         | N (%)   |
|----------------|---------|
| Female         | 123 (89.8%) |
| Male           | 14 (10.2%)  |
| Associated Comorbidities |         |
| Diabetes       | 38 (27.0%)  |
| Hypertension   | 23 (16.8%)  |
| Dyslipidemia   | 5 (3.6%)    |
| Hypothyroidism | 21 (15.3%)  |
| Hyperthyroidism| 4 (2.9%)    |
| Medically free | 60 (43.8%)  |

TABLE 1: Baseline sociodemographic information of the subjects

Ultrasonography, histopathological findings, and TIRAD scores
The authors considered any potential or confirmed thyroid malignancy on the ultrasound reports as "positive" and all the confirmed benign thyroid conditions as "negative". The majority (n=91, 66.4%) were
positive (malignant) based on the ultrasound findings, and 46 (33.6%) were negative (benign).

Of the total, most were diagnosed as benign follicular nodules with cystic degeneration based on the histopathological studies (n= 29, 21.2%). Twenty-five (18.2%) were diagnosed as papillary thyroid carcinoma, 21 (15.3%) were benign follicular nodules, 16 (11.7%) were confirmed as adenomatoid nodules with cystic degeneration, and 10 (7.3%) were chronic lymphocytic thyroiditis. However, it is necessary to acknowledge that 22 (16.1%) were negative for evidence of malignancy (Table 2). With the use of Chi-square non-parametric test, it was found that the difference between the histopathological findings were statistically significant (p <0.001).

| Histopathology Findings                                                                 | N (%)   |
|-----------------------------------------------------------------------------------------|---------|
| Adenomatoid nodule                                                                     | 1 (0.7%)|
| Adenomatoid nodule with cystic degeneration                                            | 16 (11.7%)|
| Adenomatoid nodule with cystic degeneration (suspicious for follicular cancer)         | 1 (0.7%)|
| Benign colloid nodule                                                                  | 5 (36%) |
| Benign follicular nodule                                                                | 21 (15.3%)|
| Benign follicular nodule with cystic degeneration                                      | 29 (21.2%)|
| Chronic lymphocytic thyroiditis                                                        | 10 (7.3%)|
| Follicular neoplasm                                                                    | 7 (5.1%) |
| Negative for malignancy                                                                | 22 (16.1%)|
| Papillary thyroid carcinoma                                                             | 25 (18.2%)|

**TABLE 2: The histopathological findings for the thyroid assessment**

Furthermore, the authors have investigated the TIRAD scores given for each confirmed thyroid condition falling in one of the TIRAD categories (TR1, TR2, TR3, TR4, and TR5). It was noted that the majority of them were scored as TR1 (n= 46, 33.6%), while 22 (16.1%) were reported as TR2, 24 (17.5%) were reported as TR3, 29 (21.2%) were reported as TR4, and 16 (11.7%) were reported as TR5 (p <0.001).

Comparing the histopathological findings and the given TIRAD score, the authors found that the majority of those who had benign findings on histopathology scored "TR1" (n=41, 39.0%), while the majority of those who had malignant findings on histopathology scored "TR5" (n=10, 31.3%) and to a lesser extent "TR4" (n=9, 28.1%), with all findings’ association between TIRAD and results of histopathology showing statistical significance (p<0.001). Table 3 describes the different scores given for the benign and malignant findings.

| Associations between Histopathology and the TIRADS score                           |
|-------------------------------------------------------------------------------------|
| TIRADS score                                                                         |
| TR1            | TR2            | TR3            | TR4            | TR5            | P<0.001          |
| Benign        | 41 (39.0%)    | 19 (18.1%)    | 19 (18.1%)    | 20 (19.0%)    | 6 (5.7%)         |
| Malignant     | 5 (15.6%)     | 3 (9.4%)      | 5 (15.6%)     | 9 (28.1%)     | 10 (31.3%)       |

**TABLE 3: The frequencies of the TIRAD scores given to each histopathological finding**

TIRAD: Thyroid Imaging Reporting and Data Systems

**Sensitivity and specificity of ultrasonography in thyroid malignancy**

The sensitivity, also known as the true positive rate, of ultrasonography to confirm the diagnosis of a malignant thyroid condition was found to be 59.4%. Moreover, the false negative rate, also known as the type two error, was found to be 40.6%. In contrast, the false positive rate, also known as type one error, was found...
to be 25.7%. However, the specificity of the ultrasonography, also known as the true negative rate, was found to be 74.3%.

**Discussion**

Thyroid cancer is a common malignant condition of the head and neck region. The incidence of this type of malignancy is considerably high, with an incidence of approximately 56 diagnosed cases in 100,000 individuals, ranking it as the topmost common among endocrine tumors [7,8]. In the last decade, the rate of occurrence of thyroid cancer has escalated with the general environmental deterioration, increasing its threat to public health [9].

In most clinical presentations, the early signs and symptoms of thyroid malignancy are similar to the manifestations of benign thyroid nodules. The presentation of these nodules usually includes difficulty in the natural process of breathing, neck swallowing, hoarseness of the voice, and obstruction. Therefore, a clinician should focus to diagnose and possibly exclude thyroid malignancy in a patient presenting with a thyroid nodule [10,11]. The most common thyroid cancer type is papillary adenocarcinoma, while the least common type is undifferentiated carcinoma [12]. Commonly, thyroid cancer occurs unilaterally, meaning at one side of the gland, often singularly [13]. At a pathologic level, the most commonly encountered thyroid nodule is of the papillary adenocarcinoma category, accounting for approximately 70% of all the pathological types [12].

In daily clinical practice, the ultrasound imaging modality is often utilized to diagnose thyroid cancer. However, overlaps in the images recorded by the ultrasound occur as well as frequent diversity. Thereby, a high rate of misdiagnosis is notable with the use of ultrasound imaging solely in diagnosing early as well as atypical types of thyroid cancer [14]. Fortunately, FNAB is considered a safe and simple technique, with higher diagnostic accuracy and sensitivity in the process of thyroid cancer detection [15]. Nowadays, patients presenting to the clinic with thyroid nodules are often initially examined by ultrasound. After the ultrasound examination, these patients are subjected to be examined by FNAB further in the assessment. In addition, FNAB with the guidance of ultrasonography is considered to be the current gold standard method of evaluating a thyroid lesion due to the high sensitivity associated with its findings [16]. Nevertheless, the final diagnosis of some of the thyroid lesions such as follicular adenocarcinoma for instance remains unclear [17]. The accuracy of FNAB diagnosis is based on multiple variables. These variables include the technical skills of the individual performing it and the skills of the pathologist who will process and interpret the sample biopsy.

Even though FNAB is the method of choice, the first step taken in evaluating any thyroid nodule is the use of ultrasonography. This is followed by FNAB, which decides whether the patient will undergo surgical management or not [18].

TIRADS is a criteria figured based on the radiological findings of the thyroid gland. Based on the results of the TIRADS, the decision of subjecting a lesion to FNAB is made [19]. This TIRADS algorithm used to report a thyroid lesion has clear and defined set objectives that are based on the ultrasonographic features, and the higher the TIRADS classification, the higher the thyroid malignancy risk [20,21].

In this study, the authors investigated ultrasound imaging with the use of the TIRADS criteria and FNAB method of diagnosis in relation to its diagnostic value and its role of contribution in establishing a diagnosis of thyroid malignancy. Our study results have concluded that the sensitivity of the ultrasonography in the confirmation of a malignant thyroid lesion in 59.4% and its specificity was 74.3%. In comparison, Youssef et al. have concluded in a similar study that the sensitivity of ultrasonography in detecting was 100% with a specificity of 94.12% [18].

Moreover, Wettasinghe and colleagues have noted in their published study that detecting microcalcifications as well as hypoechogenicity is considered to be the most necessary points of criteria in the prediction of thyroid malignancy, which matched our study findings [22].

Ram et al., in another study, have evaluated the sensitivity and specificity of ultrasonography in diagnosing thyroid nodules concluding that this radiological method in regard to calcifications is 80% sensitive and 68% specific, and based on hypoechogenicity are 80% and 52%, respectively [23].

Furthermore, and in comparison to our study findings, in a study conducted by Li et al. to figure the diagnostic value of FNAB in combination with the use of ultrasound, it was found that the sensitivity of the ultrasound in diagnosing thyroid cancer was 86.67%, the specificity was 28.57%, the accuracy was 80.60%, while the positive predictive value was 91.25%, and the negative predictive value was 25% [1]. Additionally, Yang et al. have noted in their study that the fine needle aspiration has an accuracy percentage of 100%; however, this accuracy is only implicated in the setting of medullary thyroid carcinoma specifically if the diameter is more than 1 cm. Moreover, if the diameter is less than 1 cm, the accuracy of the FNAB becomes 66.6% [24]. There is a published study conducted by Bahaj et al. who noted in a study on 314 subjects, the sensitivity of the fine needle aspiration is 79.8%, while the specificity was valued at 82.1%. Regarding the
positive predictive value, it was found to be 74.8%, the accuracy was 84.8%, and the negative predictive value was 85.9%, which were all generally similar to our study findings [25].

Conclusively, more clinical retrospective studies should be conducted to assess the diagnostic validity of ultrasonography in detecting thyroid malignancies in order to accurately compare the different study results and assess this radiological approach.

Conclusions
In conclusion, ultrasonography using TIRADS is a very significant step in evaluating a thyroid nodule before moving on to the next step of FNAB, despite the fact that it is the gold standard method of diagnosis. Ultrasonography is highly sensitive and specific, cost-effective, and convenient to the patient. Thereby, more retrospective studies should be conducted in the future to evaluate its diagnostic accuracy.

Additional Information

Disclosures
Human subjects: Consent was obtained or waived by all participants in this study. Institutional Review Board, General Directorate of Health Affairs, Ha’il Region, Kingdom of Saudi Arabia issued approval H-08-L074 dated April 15, 2021. Animal subjects: All authors have confirmed that this study did not involve animal subjects or tissue. Conflicts of interest: In compliance with the ICMJE uniform disclosure form, all authors declare the following: Payment/services info: All authors have declared that no financial support was received from any organization for the submitted work. Financial relationships: All authors have declared that they have no financial relationships at present or within the previous three years with any organizations that might have an interest in the submitted work. Other relationships: All authors have declared that there are no other relationships or activities that could appear to have influenced the submitted work.

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