INTRODUCTION

Thyroid dysfunction is very common in the population and thyroid function tests (TFT) are one of the frequently requested tests within the hormone panel. Thyroid-stimulating hormone (TSH), free triiodothyronine (FT3) and free thyroxine (FT4) tests are used to evaluate thyroid functions.[1] Although it may seem useful to measure both TSH and FT4 for the diagnosis of thyroid dysfunction, even small changes in FT4 levels may cause very large changes in TSH levels.[1,2] Therefore, most guidelines and algorithms propose a two-step approach (thyroid cascade) in the TFT request. In this two-step approach, after TSH measurement is performed, if the TSH result is outside the reference range, then, FT4 measurement is recommended.[3–7]

Although there are algorithms and guidelines for requesting these tests, in general, usually, all three tests are requested in combination, which leads to unnecessary financial burden.[1, 8, 9] Studies have found that TSH has higher sensitivity and specificity compared to other tests in cases of thyroid dysfunction (hypothyroidism and hyperthyroidism).[7]
METHOD

TSH, fT3 and fT4 tests of patients aged 18-65 years (n=61,422) who were accepted to Amasya Central Public Health Laboratory between 01.01.2018-31.12.2018 were retrospectively analyzed and included in this study; patients with positive pregnancy test were excluded from this study. In this study, the first group consisted of patients for whom only TSH was requested. The second group included patients for whom only fT4 and TSH, the third group included patients for whom only fT3 and TSH, and the fourth group included patients for whom all three of fT4, fT3 and TSH were requested. In addition, patient groups were evaluated by being grouped as euthyroid, hypothyroidism and hyperthyroidism according to their clinical diagnoses (Table 1). TSH, fT3 and fT4 tests were performed by electrochemiluminescence method in module e601 of Roche Cobas 6000. Reference ranges of tests were TSH (0.27-4.2 µIU/mL), fT3 (2.0-4.4 pg/mL), fT4 (0.93-1.7 ng/dL). Unnecessary test request procedure was applied for groups except for the TSH alone group (group 1), which were the 2nd Group, 3rd Group and 4th Group. Tests other than the TSH test in these groups were considered unnecessary test requests. In the cost analysis of the impact on health expenditures, the unit price of TSH, fT3 and fT4 was determined as 4.28 Turkish Liras (TL), based on the updated 2013 Health Practices Communiqué (HPC) with the 01.02.2019 amendment request entered. The total cost was calculated by multiplying the specified unit price with the number of tests in the groups where unnecessary test request is thought to have been made.

Statistical Methods

For the statistical analysis, SPSS 15 for windows version was used. Descriptive statistics were presented for as number and percentage.

RESULTS

At the end of our retrospective analysis, we reached the number of 61,422 patients recorded in the system with the TFT requests. Among these patient results, the number of the patients who were requested only TSH, in accordance with the algorithms, was 12,522 (20.2%); the number of the patients who were requested fT3 and TSH together was 703 (1.1%); the number of the patients who were requested fT4 and TSH was 6,343 (11.2%); and the number of the patients who were requested all three (TSH, fT4 and fT3) was 41,854 (67.5%). In the first step, when only test requests other than TSH request are considered unnecessary, costs were 3,008 TL for fT3 and TSH request, 27,148 TL for fT4 and TSH and 358,270 TL for fT3, fT4, while the total cost of the unnecessary test request to our laboratory was found to be 388,426 TL per year as the monetary financial equivalent of reimbursement to the institutions according to the HPC principles.

DISCUSSION

In this study, the extent to which the correct test request procedures are applied in the algorithms specified in the thyroid guidelines and the financial burden of the unnecessary test requests of the samples that the correct test request procedure was not applied are investigated. According to the results, it was determined that the correct test request procedures in the algorithms specified in the guidelines were applied only at a 20.2% rate. In addition, the monetary financial equivalent of reimbursement to the institutions according to the HPC principles for the unnecessary tests on samples not conforming to the correct test request procedures laid out in the thyroid guidelines and algorithms was found to be 388,426 TL per year.

As stated in the diagnostic guidelines, it is generally recommended to use 2nd and 3rd generation methods in the laboratories to have high sensitivity and specificity of the TSH test, which should be measured in the first step. [7, 10-12] In studies with outpatients with no expectancy of pituitary/hypothalamic disease, it is often stated that it is usually unnecessary to measure fT4 when TSH is normal. [7, 13] Therefore, when performing TFT measurement, it is recommended by the thyroid guides that only the TSH measurement should be performed in the first step and

<table>
<thead>
<tr>
<th>Group 1 (TSH)</th>
<th>Group 2 (fT4+TSH)</th>
<th>Group 3 (fT3+TSH)</th>
<th>Group 4 (fT3+fT4+TSH)</th>
</tr>
</thead>
<tbody>
<tr>
<td>Euthyroid</td>
<td>10,526 (84.1)</td>
<td>5,022 (79.2)</td>
<td>692 (98.4)</td>
</tr>
<tr>
<td>Hypothyroid</td>
<td>861 (6.9)</td>
<td>741 (11.7)</td>
<td>8 (1.1)</td>
</tr>
<tr>
<td>Hyperthyroid</td>
<td>1,135 (9.0)</td>
<td>580 (9.1)</td>
<td>3 (0.5)</td>
</tr>
</tbody>
</table>

fT3: free triiodothyronine; fT4: free thyroxine; TSH: Thyroid-stimulating hormone.

The data were presented as n (%).
that fT4 measurement should be performed in the second step (using 2-step thyroid cascade) if the TSH measurement is out of the specified reference range. Therefore, TSH measurement is very important given that TSH measurement decides how the thyroid measurement cascade will progress in this case.

In previous studies, it was found that TFT were performed in compliantly with the algorithms and unnecessary test request rate was high. Demirci et al. investigating the suitability of diagnostic algorithms have reported the rate of inappropriate test requests for internal polyclinics as 40.6%; for all non-internal clinics as 52.4%; Yıldırım et al. reported the same as 36.6%, and Tekçe et al. as 76%. In our study, the rate of inappropriate test requests for TFT tests was 79.8%. Such unnecessary requests of the tests lead to an increase in laboratory load and loss of time as well as an unnecessary increase in financial burden. It is quite difficult to change existing habits regarding the test request. Test requests made according to guidelines and algorithms should be used effectively to reduce costs and achieve an accurate clinical diagnosis of the disease. In this way, health expenditures will be positively contributed as well as reliable diagnostic methods will be supported with reliable results. Kim et al. reported that instead of the classical algorithm, the addition of quick-pick procedures to the test selection program in accordance with the rules specified in the guidelines reduced the unnecessary test requirement by 50%. The use of quick-pick test procedures by clinicians according to reflex test rules can prevent both confusions when making a test request and unnecessary testing. If certain test panels are identified and used according to the priorities specified in the guidelines, the applicability of the algorithm may become more effective and practical for the clinician. Demirci et al. found that the unnecessary test request decreased significantly when the reflex test was started and interpreted the decrease in the unnecessary test request, especially in the units where the TFT request was intensified, such as internal diseases, as a change in the traditional habits.

A study reported that 46% of fT4 and TSH (2-step test request) requests were performed together. Another study found that the rate of 2-step testing was 47%. This rate was found to be 11.2% in our study. The low rate compared to other studies is generally attributable to the higher rate of all three tests being requested together (TSH, fT4 and fT3: 67.4%).

CONCLUSION

According to the results we obtained, it was found that the algorithms related to the request of thyroid tests were not followed and the financial implications of this situation were high. We believe that the financial burden will be reduced by the use of applications, such as reflex testing to reduce unnecessary test requests, informing about the algorithms specified in the guidelines, and widespread implementation of test requests in an accurate and effective manner.

Disclosures

Ethics Committee Approval: This study was approved by the Nigde Omer Halisdemir University Faculty of Medicine Ethics Committee (No: 2020/04 Date: 25 Jun 2020). This study was conducted in accordance with the Declaration of Helsinki.

Peer-review: Externally peer-reviewed.

Conflict of Interest: None declared.

REFERENCES

8. Kın Tekçe B, Dikbas O, Tekçe H, Tosun M. Evaluation of the Requests for Thyroid Function Test According Algorithms and
13. Viera AJ. Thyroid function testing in outpatients: are both sensitive thyrotropin (sTSH) and free thyroxine (FT4) necessary? Fam Med 2003;35:408–10.