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ABSTRACT

Introduction: Subclinical hypothyroidism is an endocrine disorder that can adversely affect cognitive performance. In this regard, patients with subclinical hypothyroidism may develop poor attention. However, the effects of this endocrine disorder on auditory attention is unclear. This study aims to evaluate the impact of subclinical hypothyroidism on auditory divided attention.

Materials and Methods: This cross-sectional, case control study was performed on 30 adult patients, aged 20-40 years, who met the study inclusion criteria. Based on their Thyroid Stimulating Hormone (TSH) level, the subjects were divided into two case groups (each with 15 patients) with a TSH level of 5<TSH≤10 and 10<TSH≤15. A healthy control group was recruited with matched age, sex, and education with the case groups. The participants were evaluated by the Persian version of the Bergen dichotic listening test. Among different experimental situations that included non-forced attention, forced right, and forced left, we chose the non-forced attention. Statistical analysis was done in SPSS V. 25.

Results: There was a significant difference between the subclinical hypothyroid ($5 < TSH \le 10$ group) and controls in the scores of the right and left ear. However, there was no significant difference between scores of right and left ear in subclinical hypothyroid ($10 < TSH \le 15$ group). No significant difference in scores of the right and left ear was found between men and women in three groups.

Conclusion: Subclinical hypothyroidism had no effect on auditory divided attention.

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1. Introduction

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ubclinical hypothyroidism is a common endocrine disorder that occurs when Thyroid Stimulating Hormone (TSH) levels are higher than normal range (0.27≤TSH≤5 µIU/mL) while the level of

T3 (0.92-2.33 nmol/L) and T4 (5.13-14.1 μ g/dL) hormones are within the normal range. However, in overt hypothyroidism, the levels of T3 and T4 hormones drop lower than normal [1, 2]. The prevalence of subclinical hypothyroidism is more than overt hypothyroidism, and it is more common in women [3, 4].

Different studies show that subclinical hypothyroidism adversely affects cognitive performance, causing memory impairment, low concentration, and poor attention [5-7]. Mcdermot et al. found that people with subclinical hypothyroidism showed poor memory and reasoning [3]. Akintola et al. reported that in adults subclinical hypothyroidism like overt hypothyroidism manifested with deficits in cognitive abilities like memory, attention, and concentration [8].

Attention is one of the cognitive functions that has an essential role in information processing [9]. It is the ability to filter the information to obtain the preferred data [10]. Attention can advocate visual, auditory, and tactile information [11]. Auditory attention is the ability to focus on preferred acoustic stimuli while simultaneously receiving other stimuli. Because acoustic and phonetic features of the language are affected by auditory attention, it may have essential roles in reading and writing [12].

Auditory attention has various types, including divided attention and selective attention. The divided attention refers to simultaneously focusing on two or more acoustic stimuli present in dichotic situation. Selective attention refers to focus selectively on preferred information while ignoring other competitive stimuli [10]. Compared to selective attention, divided attention is a more difficult task and demands more brain activity because of focusing on two or more stimuli at the same time [13].

For this reason, any possible processing disorder can affect divided attention. Divided attention is a practical and challenging skill needed in everyday life activities such as listening to the teacher and taking notes simultaneously or listening to music, and somebody's talking at the same time [14].

The dichotic listening test is one of the most valid behavioral tests that are used for evaluating the function of the brain hemispheres, transferring information between two hemispheres, and diagnosis of central auditory disorders [13]. Bergen dichotic listening test is a dichotic test that is used widely to investigate auditory attention and language laterality in healthy people, individuals with brain injury, and patients with attention deficits [15]. In dichotic listening, two different stimuli are presented to the right and left ear simultaneously. In this experiment utilizing speech stimuli, the right ear usually obtains higher scores than the left ear in healthy people. This result shows that left hemisphere is specialized for language understanding [16, 17].

Various studies have evaluated the effects of subclinical hypothyroidism on different kinds of attention, but their results are controversial [18-21]. Some studies reported that subclinical hypothyroidism had a negative impact on auditory attention. For example, Paladugo et al. compared cognitive functions between patients with subclinical hypothyroid and control groups using latencies of P300 auditory evoked potentials. They found that subclinical hypothyroid patients had P300 latency prolongation compared with the control group [7].

Quijano et al. assessed auditory attention with the Reaction Time Test (RT) in subclinical hypothyroid patients. Their results showed slowed reaction time in subclinical hypothyroid patients [22]. Nystrom et al. also noted a decrease in auditory and visual attention performance in elderly patients with subclinical hypothyroidism [23]. Baldini et al. evaluated neuropsychological functions in subclinical hypothyroid patients. They assessed auditory attention with RT. Their findings showed that RT was not prolonged in subclinical patients. Thus subclinical hypothyroidism had no effect on auditory attention [18].

Gussekloo et al. evaluated the impact of subclinical hypothyroidism on attention with Stroop test. They reported that attention did not affect patients with subclinical hypothyroid [19]. Park et al. used digit span test to evaluate attention in patients with subclinical hypothyroid. They demonstrated that subclinical hypothyroid had no adverse effect on auditory attention in general [24]. Jorde et al. also used digit span test to evaluate the impact of subclinical hypothyroidism on attention. Their results did not show any changes in sustained attention or attention in general [25].

It is well known that thyroid and its disorders such as overt hypothyroidism have a detrimental effect on central nervous system, but the impact of subclinical hypothyroidism on cognitive functions is inconclusive and needs to be determined [26, 27]. Furthermore, most of the previous studies have been conducted on older population (because hypothyroidism is more prevalent in this group) or have considered a wide range of age in their sample [18, 28, 29]. Also previous studies did not use dichotic tests for the evaluation of auditory attention. For these reasons, in the present study, we evaluated the effects of subclinical hypothyroidism on auditory divided attention among young patients with subclinical hypothyroid by the Persian version of the Bergen dichotic listening test.

2. Materials and Methods

This cross-sectional case control study was conducted on two groups of patients with subclinical hypothyroidism, and one group of healthy participants (age range: 20-40 years) with each group consisted of 15 participants. The first subclinical hypothyroid (5<TSH \leq 10 µIU/mL) group included nine women and six men, and the second subclinical hypothyroid (10<TSH \leq 15) group included 10 women and 5 men. This study was conducted in Audiology Clinic of Tehran University of Medical Sciences (TUMS).

The study participants were recruited from patients referring to Endocrine Clinic of Shariati Hospital affiliated to TUMS, and their diseases was approved by an endocrinologist. Their thyroid hormones levels were determined by blood tests. Four individuals in the first subclinical hypothyroid group and one individual in the second subclinical hypothyroid group had anti-thyroid peroxidase antibody (Anti-TPO Ab) higher than average, proving they had Hashimoto's thyroiditis, a kind of autoimmune disease.

The participants in three groups lacked any significant neurological and cognitive problems, did not use psychological drugs, alcohol or tobacco smoking, lacked history of head trauma, brain surgery and epilepsy audiological or neurological problems, had hearing threshold equal to or below 25 dB HL at octave frequencies from 250 to 8000 Hz and scored above 90% in speech discrimination score, had acoustic reflex at 0.5, 1, and 2 kHz and type A tympanometry in both ears.

They were right-handedness following the standard of Edinburgh. They were monolinguals (speaking Persian) and had no musical experience. They were not under treatment for hypothyroidism before the tests. Their thyroid test results were the 5<TSH \leq 10, 10<TSH \leq 15, in the case (subclinical hypothyroid) groups, and 0.27 \leq TSH \leq 5 µIU/mL in the control group.

The absence of any of the above criteria resulted in the exclusion of the subjects from the study. To recruit the participants, we carried several evaluations to observe the study inclusion criteria. To roll out any severe cognitive involvement, depression, and anxiety, mini-mental state examination, Beck depression inventory, and state-trait anxiety inventory were performed, respectively. The participants were asked to complete the written consent form and were compensated for their participation. Finally, the auditoryverbal memory test was performed via Lenovo Z500 laptop which connected to a calibrated AC40 audiometer with TDH39 headphones.

The materials of the Persian version of the Bergen dichotic listening test were three sets of 36 pairs of syllables. Each syllable is made up of an stop consonant (/b, d, g, p, t, k/) plus vowel /a/. Each set was used for different experimental situations, including Non-forced attention (NF), Forced Right (FR), and Forced Left (FL). At the most comfortable level in NF situation, the subjects would hear a duplicated presentation of six syllables with a CV (consonant-vowel) pattern at a comfortable listening level, and they should repeat the best syllable heard after each presentation.

They should tell only one syllable. At the end of presentation, calculation of the percentage of correct answers and laterality of each ear in each situation was determined. The formula, $LI = (R Ear - L Ear/R Ear + L Ear) \times 100$, was used to determine the laterality of each ear in each experimental situation. In this formula, LI refers to laterality index, R Ear is the score of attention to the right, and L Ear is the score of attention to the left ear. If the LI were positive, it would indicate the right ear laterality, and if it were negative, it would mean the left ear laterality.

One-way ANOVA was done to measure any statistical difference between the three groups. The Student t-test was used to measure the effect of sex in each group, and the paired samples t-test was used to measure ear effect in each group. Statistical analysis was fulfilled with SPSS V. 25.

3. Results

Table 1 shows the characteristics of case and control groups. Information is expressed in Mean±SD.

Table 2 shows the results of one-way ANOVA for auditory attention in each ear separately by Non-Forced attention situation (NF). There was no significant difference between the three groups in scores of right ears and left ears separately.

Table 3 shows the results of the Student t-test for the sex effect in each group. There was no significant dif-

Domosius Mariaklas	Subclinical Hy	Control		
Demographic variables	5 <tsh≤10< th=""><th>10<tsh≤15< th=""><th colspan="2">Control</th></tsh≤15<></th></tsh≤10<>	10 <tsh≤15< th=""><th colspan="2">Control</th></tsh≤15<>	Control	
Number	15	15	15	
Gender (male/female)	6/9	5/10	6/9	
Age, y	32.20±6.15	32.66±4.46	27.26±6.46	
PTA, min	14.33±2.58	13.33±3.08	14±3.38	
TSH (μIU/mL)	6.87±0.94	11.71±1.30	1.82±1.19	
T4 (μg/dL)	7.52±2.09	6.79±1.16	7.85±1.86	
T3 (nmol/L)	1.18±0.18	1.30±0.35	1.52±0.32	
Anti-TPO Ab (IU/mL)	55.50±109.31	16.61±0.58	-	
			JMR	

Table 1. Demographic and clinical characteristic of study groups (n=45)

Table 2. One-way ANOVA results for auditory attention between groups in each ear

Ear	Group	Sum of Squares	df	Mean Square	F	Sig.
Right ear	Between groups	107.16	2	53.58	0.71	0.49
Left ear	Between groups	179.65	2	89.82	1.28	0.28

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ference in scores of right and left ears between men and women in three groups.

ence between scores of right and left ears in control and subclinical hypothyroid 5<TSH≤10 groups. But there was no significant difference between scores of right and left ears in subclinical hypothyroid 10<TSH≤15 group.

Table 4 shows the results of paired samples t-test for ear effect in each group. There was a significant differ-

Table 3. The Independent samples t-test results comparing auditory attention in each group with sex- and ear-specific

Group	Ear	Sex	Mean±SD	df	t	Р
Control	Dight oor	Female	51.47±6.03	10	1 4 2	0.20
	Right ear	Male	57.77± 11.08	15	-1.45	0.50
	Left eer	Female	43.33±6.87	10	1.20	0.52
	Leitear	Male	37.21±10.62	13	1.30	0.53
Subclinical hypothy- roid 5 <tsh≤10< td=""><td>Dight oor</td><td>Female</td><td>49.25±6.40</td><td>10</td><td>0.22</td><td>0.22</td></tsh≤10<>	Dight oor	Female	49.25±6.40	10	0.22	0.22
	Right ear	Male	47.77±10.68	13	0.33	0.33
	Loft oor	Female	46.29±5.63	13	0.20	0.12
	Leitear	Male	47.21±12.18		-0.20	0.12
Subclinical hypothy- roid 10 <tsh≤15< td=""><td>Dight oor</td><td>Female</td><td>48.66±7.40</td><td>10</td><td>2 72</td><td>0.12</td></tsh≤15<>	Dight oor	Female	48.66±7.40	10	2 72	0.12
	Right ear	Male	43.99±11.44	15	2.72	0.12
	Loft oor	Female	43.33±8.77	10	0.72	0.41
	Leitedi	Male	41.99±15.54	12	0.72	0.41

Group	Number	Ear	Mean±SD	df	t	Р
Control	15	Right	53.99±8.65	14	3.09	0.001
		Left	40.88±8.77			
Subclinical hypothyroid 5 <tsh≤10< td=""><td rowspan="2">15</td><td>Right</td><td>48.66±8.04</td><td rowspan="2">14</td><td>0.48</td><td rowspan="2">0.000</td></tsh≤10<>	15	Right	48.66±8.04	14	0.48	0.000
		Left	46.66±8.45		0.10	
Subclinical hypothyroid 10 <tsh≤15< td=""><td rowspan="2">15</td><td>Right</td><td>47.55±10.72</td><td rowspan="2">14</td><td>0.42</td><td>0.07</td></tsh≤15<>	15	Right	47.55±10.72	14	0.42	0.07
		Left	45.53±10.89		0.12	0.07

Table 4. Paired samples t test results comparing auditory attention between groups in each ear

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Figure 1 shows the between-groups mean scores of auditory attention in the right ear and left ear. The mean scores of right ear in the control group is more than the mean scores of right ear in subclinical hypothyroid 5<TSH≤10 group, and the mean scores of right ear in the first subclinical hypothyroid 5<TSH≤10 group is more than the mean scores of right ear in the second subclinical hypothyroid 10<TSH≤15 group. But about left ear, there was not constant change in mean scores.

At first, the mean scores of left ear increased and then decreased. The laterality index was calculated in three groups. In the control group, 80% (12 people) had right ear advantage, 13.3% (2 people) had left ear advantage and 6.6% (1 person) had no ear advantage. In the first subclinical hypothyroid ($5 < TSH \le 10$) group, 66.66% (10 people) had right ear advantage, and 13.33% (2 people) had no ear advantages. In the second subclinical hypothyroid ($10 < TSH \le 15$) group, 73.33% (11 people) had right ear

advantage, 20% (3 people) had left ear advantage, and 6.66% (1 person) had no ear advantage.

4. Discussion

This study aimed to evaluate auditory divided attention in two subclinical hypothyroid ($5 < TSH \le 10$, $10 < TSH \le 15$) patient groups and a control group by the Persian version of the Bergen dichotic listening test. Findings indicated no significant difference among the three groups regarding the mean scores of right and left ear (Table 2). Gender did not affect-test results (Table 3). There were significant differences between scores of right and left ear between the first subclinical hypothyroid ($5 < TSH \le 10$) and the control groups, but there were no significant differences between scores of right and left ear in the second subclinical hypothyroid ($10 < TSH \le 15$) (Table 4).

Different studies showed different results about the effects of subclinical hypothyroidism on auditory atten-



Figure 1. The mean scores of auditory attention with ear separation between groups

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tion [18, 19, 25]. For example, park et al. showed that auditory attention performance did not change in subclinical hypothyroid patients [24]. Jorde et al. found no difference between control and subclinical hypothyroid groups in auditory attention performance [25]. Baldini et al. showed that auditory attention performance did not change in subclinical hypothyroid patients. However, few studies yield opposite results [18]. Nystrom et al., in their research, proved that subclinical hypothyroidism had adverse effects on auditory attention [23]. Quijano et al. showed auditory attention performance reduction in patients with subclinical hypothyroidism [22].

Different results in previous studies about the effect of subclinical hypothyroidism on auditory attention could be explained by hormone level disturbances, cognitive involvements, and anxiety. It was demonstrated that the lower levels of TSH hormone might suppress the effects of subclinical hypothyroidism on auditory attention because of a direct relationship between TSH hormone level and severity of the disorder [30].

Another reason is the effect of aging because older patients usually have some cognitive dysfunctions such as reduction of memory and attention performance and depression; these confounding factors can have adverse effects on the test results. So that changes of auditory attention could be due to the effects of aging and not the direct effect of subclinical hypothyroidism. Another reason is ignoring the neuropsychological factors such as anxiety before doing tests. Anxiety is one of the factors that may have adverse effects on the test results. Some studies disregard this factor, do the tests, and ignore the stress of their subjects [25].

In the present study, the Persian version of the Bergen dichotic listening test was used on patients with subclinical hypothyroid. Therefore, the method of presenting stimuli and calculating scores was different from other auditory attention tests. For this reason, we could not compare the results of this test with the results of other studies. As Bergen dichotic test is one of the tasks that require a lot of processing and mental demanding, the subclinical hypothyroidism in the present study could not sufficiently influence the processing centers that advocate for auditory divided attention [31].

One reason may be the less sensitivity of these areas to subclinical hypothyroidism. According to much evidence, in changing levels of thyroid hormones, different areas of the brain, including cognitive levels with different sensitivities, exhibit different functions [32, 33]. Probably the prefrontal cortex for processing of divided attention is not sensitive under these conditions [34]. These hypotheses, however, are beyond the scope of the current study; hence, further studies are needed to determine the reliability of the effect of subclinical hypothyroidism on auditory attention.

In the current study, there were significant differences between scores of right ear and left ear in the control and subclinical hypothyroid (5<TSH≤10) groups in the way that scores of the right ear were higher than scores of left ears. Jafari et al., in two studies, assessed scores of right and left ear in auditory attention utilizing Bergen dichotic test in healthy adults. They found that scores of the right ear were more than scores of left ear [13, 35].

The results of these two studies were consistent with the control and the subclinical hypothyroidism (5<TSH≤10) groups of the present study. Right ear advantage is a stable index that represents anatomical differences in right and left sides of the central auditory nervous system [35, 36]. It means that contralateral neural tracts compared with ipsilateral neural tracts are wider. Moreover, left hemisphere compared with right hemisphere has been specialized for language inputs. Therefore, speech stimuli presented to right ear are processed more faster than left ear [35].

In the subclinical hypothyroid ($10 < TSH \le 15$) group, although there was no significant difference between scores of right and left ears, the scores of the right ear were more than left ear. It seems that the non-significant difference was due to the higher standard deviation in subclinical hypothyroid ($10 < TSH \le 15$) group, compared with the two other groups. Because the obtained data were scattered, the possibility of a significant difference would be low. On the other hand, laterality index in the subclinical hypothyroid ($10 < TSH \le 15$) group is the same as two other groups, and most of the patients in this group had right ear advantage.

There was no significant difference between men and women in three groups regarding the scores of the right and left ear. Jafari et al., in their study, evaluated the effect of gender on the auditory attention by the Bergen dichotic test in 75 healthy adults. They found that gender did not affect auditory attention [13].

These results were similar to the results of the analysis of data from the Bergen dichotic test database, which was conducted on nearly 1000 people [37]. Sommer et al. assessed the effect of gender on auditory attention.

They reported no significant difference between men and women in auditory attention [38]. Hiscook et al. evaluated 477 healthy right-handed adults in five successive tests using dichotic language stimuli in which gender did not affect dichotic test results [39]. These studies were similar in their outcomes with the results of the present study.

The present study had a few limitations. First, our sample size was small due to difficulty in access to the appropriate participants, and second, we did not have enough time for conducting longitudinal study. Future studies are recommended to consider larger sample size and long-term follow-up to find out subtle changes in the subclinical hypothyroidism patients.

5. Conclusion

In the present study, subclinical hypothyroidism had no adverse effect on auditory divided attention. Also, subclinical hypothyroidism had no significant impact on right ear advantage in both case groups. Finally, gender did not affect dichotic test results.

Ethical Considerations

Compliance with ethical guidelines

The current study protocol was approved by the Ethics Committee of Tehran University of Medical Sciences with the Code: IR.TUMS.FNM.REC.1397.128.

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Authors contributions

Original manuscript draft: Fatemeh Khorasani; Designing and developing; The protocol: Nematollah Rouhbakhsh, Fatemeh Khorasani; Editing the manuscript: Nematollah Rouhbakhsh, Ahmad Geshani, Mohammad Reza Mohajeri Tehrani; Statistical analysis: Mostafa Qorbani.

Conflict of interest

The authors declared no conflict of interest.

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