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# Effects of Covid-19 on the audio-vestibular system

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ARTICLE INFO	A B S T R A C T
Keywords: COVID-19 Hearing Vertigo	<i>Purpose</i> : It was aimed to investigate the effects of COVID-19 infection on hearing and the vestibular system. <i>Methods</i> : Twenty-six patients whose treatment had been completed and who had no previous hearing or balance complaints were included in the study. Patients diagnosed with the disease by PCR were included in the study. Patients with at least one month of illness were included in the study. The hearing of patients was evaluated with transient evoked otoacoustic emissions (TEOAE) and pure-tone audiometry. Bedside tests, the European Evaluation of Vertigo scale (EEV), Video Head Impulse Test (vHIT), Ocular Vestibular Myogenic Evoked Potential (oVEMP), Cervical Vestibular Myogenic Evoked Potential (cVEMP) and Videonystagmography (VNG) tests were applied to evaluate the vestibular system. <i>Results</i> : A statistically significant difference was found between the COVID-19 positive and control groups according to the mean values of the 4000 Hz and 8000 Hz in both the right and left ears ( $p < 0.05$ ). No statistically significant difference was found between the COVID-19 positive and control groups according to the mean values of the 4000 Hz and 8000 Hz in both the right and left ears ( $p < 0.05$ ). No statistically significant difference was found between the COVID-19 positive and control groups in terms of their normal or pathological VNG saccade, optokinetic and spontaneous nystagmus values ( $p > 0.05$ ). The normal and pathological VNG head shake values were found to be significantly different between the COVID-19 positive and control groups ( $p < 0.05$ ). <i>Conclusion:</i> The high frequencies in audiometry in the COVID-19 positive group were worse than those in the control group. In the vestibular system, especially in oVEMP and cVEMP, asymmetric findings were obtained in comparison to the control group, and a low gain in vHIT was shown. This study shows that the audiovestibular system of people with COVID-19 infection may be affected.

## 1. Introduction

The novel coronavirus disease-2019 (COVID-19), which spreads rapidly around the world and appeared in the Wuhan city in the Hubei province of China in December 2019, was declared as a global pandemic by WHO in March 2020 [1]. The name of the virus that causes this disease has been determined as the severe acute respiratory syndromecoronavirus-2 (SARSCOV-2) [2]. 175 million cases and 3.7 million deaths had been reported worldwide by June 2021 [3]. Symptoms in this disease range from a mild upper respiratory tract infection to severe pneumonia. Common clinical symptoms include dry cough, fever, headache, sore throat, shortness of breath, diarrhea, vomiting and abdominal pain [4–6]. Anosmia and taste changes are common symptoms [7,8]. Studies have investigated whether the SARS-CoV-2 virus has an indirect or direct neurotrophic effect on the nervous system [9–10]. Various neurological symptoms such as loss of consciousness, headache and dizziness have been reported in COVID-19 patients [11–14]. Facial paralysis, sudden hearing loss and cases of vertigo among otologic symptoms have also been reported in relation to COVID-19 [15,16]. Whether COVID-19 has an effect on hearing and the vestibular system is still unclear. In this study, we planned to investigate whether there was any change in the hearing and vestibular system in patients with COVID-19 infection using a large battery of tests after recovery.

## 2. Material and method

This is a prospective study. The research was conducted in a Department of Audiology between April 2020 and June 2020. Twentysix patients whose treatment had been completed and who had no previous hearing or balance complaints were included in the study.

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Information was obtained about their complaints at the time of the disease, hospitalization and treatment that they received. Patients diagnosed with the disease by PCR were included in the study. Patients with at least 30 days of illness were included in the study. Those with severe disease or hospitalization at intensive care units were excluded from the study. Patients with previous hearing problems or balance disorders, those who had ear surgery, cardiovascular and circulatory problems, and patients using chloroquine were excluded from the study. The patients were asked whether or not they had any problems with hearing and balance before by taking a detailed anamnesis, and those who did not have any problems were included in the study. It was made sure to question the patients' histories of exposure to noise, and those with a history of working in a loud place were excluded. To conduct the study, approval was obtained from the Ethics Committee of University Institute of Health Sciences (Decision number: 2020/1268), and consent was obtained from all individuals participating in the study. Audiometry and transient evoked otoacoustic emissions (TEOAE) to assess patients' hearing, the European Evaluation of Vertigo scale (EEV) for vestibular system assessment, the video head impulse test (vHIT), ocular Vestibular Myogenic Evoked Potential (oVEMP), cervical Vestibular Myogenic Evoked Potential (cVEMP) and Videonystagmography (VNG) tests were performed.

### 2.1. Statistical analysis

The analysis of the data included in the study was carried out with the SPSS (Statistical Package for the Social Sciences) 25 program. The significance level (p) was taken as 0.05 for the comparison tests. Shapiro Wilk Test was used to check whether the data fit a normal distribution. Since the data were normally distributed, comparisons between the case and control (COVID (+), COVID (-)) groups were performed with the significance test (*t*-test) of the difference between the two means. The homogeneity of variances was checked by Levene's test to decide which test result to look for in the comparison (p > 0.05). The values of the variables are given as frequency, percentage, mean and standard deviation. In the analysis of the categorical data, cross-tables were created, and chi-squared ( $\chi^2$ ) analysis was carried out.

## 3. Results

A total of 53 people were included in the study, where 26 (49.1%) of these individuals were in the COVID-19 positive group, and 27 (50.9%) were in the control group. 36 (67.9%) of the participants were female, 17 (32.1%) were male, and the mean age of the participants was calculated as 28.98  $\pm$  10.91 years. There were 14 (53.8%) female and 12 (46.2%) male individuals in the COVID-19 positive group, and their mean age was calculated as  $34.20 \pm 12.63$ . There were 22 (81.5%) female and 5 (18.5%) male individuals in the control group, and their mean age was calculated as 23.96  $\pm$  5.92. The mean disease duration of the COVID-19 positive patients was found as 66.35  $\pm$  23.51 days. The symptoms seen during the illness of the patients were evaluated. Fever was observed in 8 (30.8%) of the patients, but not in 18 (69.2%). 5 (19.2%) had respiratory distress, but none of them needed oxygen support. 21 of the patients (80.8%) had no respiratory distress. While 11 (42.3%) had joint pain and headache, 15 (57.7%) did not. The numbers of the patients with and without dizziness were equal to 13 each (50%). While there was loss of taste in 4 (15.4%) patients, this was not observed in 22 (84.6%) patients. Oxygen support and hospitalized patients were not available, and fatigue was observed in all patients (26). Favipiravir was used for treatment in 22 patients (88.5%), and no treatment was given in 3 (11.5%) (Table 1).

## 3.1. Audiology results

The 125 Hz, 250 Hz, 500 Hz, 1000 Hz, 2000 Hz, 4000 Hz, 8000 Hz and PTO (average of the results for the 500, 1000, 2000 and 4000)

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#### Table 1

Characteristics of patients with covid.

Symptom		Number	%
Fever	+	8	30,8
	_	18	69,2
Respiratory distress	+	5	19,2
	-	21	80,8
Oxygen support	+	26	100,0
Hospitalization	-	26	100,0
Weakness	+	26	100,0
Joint pain	+	11	42,3
	-	15	57,7
Headache	+	11	42,3
	-	15	57,7
Dizziness	+	13	50,0
	-	13	50,0
Loss of the smell	+	4	15,4
	-	22	84,6
Loss of the sense	+	12	46,2
	-	14	53,8
Treatment	Favipiravir	23	88,5
	No treatment	3	11,5
Days of illness	$\text{Mean} \pm \text{ss}$	66,35 $\pm$ 23,51 day	,

ss; standard deviation.

average values of the participants were examined in both the right and left ears. No statistically significant difference was found between the COVID-19 positive and control groups in the mean values of the 125 Hz, 250 Hz, 500 Hz, 1000 Hz, 2000 Hz and PTO measurements in both the right and left ears (p > 0.05). A statistically significant difference was found between the COVID-19 positive and control groups according to the mean values of the 4000 Hz and 8000 Hz in both the right and left

#### Table 2

Comparison of groups according to audiological values.

	Group	Right ear			Left ear		
		$\frac{\text{Mean} \pm}{\text{ss}}$	Test <sup>a</sup>	p value	$\frac{\text{Mean} \pm}{\text{ss}}$	Test <sup>a</sup>	p value
125	Covid	10,38 $\pm$	1179	0,244	10,96 $\pm$	1125	0,266
Hz	(+)	5,99			6,33		
	Covid	8,33 $\pm$			9,07 $\pm$		
	(-)	6,65			5,89		
250	Covid	9,62 $\pm$	1497	0,141	10,77 $\pm$	0,953	0,345
Hz	(+)	7,34			6,59		
	Covid	6,85 $\pm$			9,07 $\pm$		
	(–)	6,07			6,36		
500	Covid	13,27 $\pm$	2447	0,018	9,81 $\pm$	1837	0,072
Hz	(+)	6,47			7,14		
	Covid	8,89 $\pm$			6,48 $\pm$		
	(-)	6,55			6,02		
1000	Covid	6,92 $\pm$	2090	0,052*	4,81 $\pm$	0,629	0,532
Hz	(+)	5,67			6,08		
	Covid	6,07 $\pm$			3,89 $\pm$		
	(–)	4,17			4,46		
2000	Covid	11,15 $\pm$	2737	0,059*	9,04 ±	1051	0,298
Hz	(+)	6,68			6,33		
	Covid	8,67 $\pm$			7,22 $\pm$		
	(-)	5,19			6,25		
4000	Covid	13,27 $\pm$	4034	0,001*	11,54 $\pm$	3862	0,001*
Hz	(+)	8,6			7,59		
	Covid	5,37 $\pm$			4,81 $\pm$		
	(–)	5,18			4,7		
8000	Covid	13,65 $\pm$	2970	0,005*	15,77 $\pm$	2031	0,047*
Hz	(+)	7,29			10,93		
	Covid	7,78 $\pm$			10,56 $\pm$		
	(–)	7,12			7,51		
PTO	Covid	11,12 $\pm$	3372	0,191*	8,58 $\pm$	2069	0,144*
	(+)	5,78			5,32		
	Covid	10,52 $\pm$			5,85 $\pm$		
	(–)	4,01			4,22		

ss; standard deviation, a; significance test of the difference between two means (*t*-test).

<sup>\*</sup> p < 0,05 there is a statistically significant difference between groups.

## ears (*p* < 0.05, Table 2).

#### 3.2. TEOAE results

The TEOAE results measured in both the right and left ears of the participants were examined at 1000 Hz, 1400 Hz, 2000 Hz, 2800 Hz and 4000 Hz. No statistically significant difference was found between the COVID-19 positive and control groups according to the mean values of the 1000 Hz, 1400 Hz, 2000 Hz, 2800 Hz and 4000 Hz measurements in both the right and left ears (p > 0.05). Although not statistically significant, the mean values of the 1000 Hz, 1400 Hz, 2800 Hz, 2800 Hz, 1400 Hz, 2800 Hz, and 4000 Hz measurements made in both the right and left ears were found to be lower in the COVID-19 positive group in comparison to the control group (p < 0.05, Table 3).

## 3.3. EEV results

The mean European Evaluation of Vertigo scale (EEV) score in the COVID-19 positive group was found to be 4.5. A statistically significant difference was calculated between the COVID-19 positive group and the control group according to EEV (p < 0.05).

### 3.4. Bedside test results

Chi-squared ( $\chi^2$ ) analysis was performed by creating cross-tables to compare the normal and pathological values of the bedside Romberg, tandem Romberg, Fukuda and tandem gait-tests between the COVID-19 positive and control groups. No statistically significant difference was found between the COVID-19 positive and control group in terms of the normal or pathological bedside Romberg values (p > 0.05, Table 4). A statistically significant correlation was found between the normal or pathological values in the tandem Romberg, Fukuda and tandem gaittests (p < 0.05, Table 4).

## 3.5. VNG results

Cross-tables were created, and Chi-squared ( $\chi^2$ ) analysis was performed to compare the normal and pathological values of VNG gaze vertical, gaze horizontal, saccade, pursuit, optokinetic, spontaneous

#### Table 3

Comparison of groups according to teoae values.

	Group	Right ear			Left rar		
		$\begin{array}{c} \text{Mean} \\ \pm \text{ ss} \end{array}$	Test	p value	$\begin{array}{c} \text{Mean} \\ \pm \text{ ss} \end{array}$	Test	p value
1000	Covid	$9 \pm$	-2085	0,142*	10,57	-1454	0,152
Hz	(+)	4,71			$\pm$ 4,09		
	Covid	12,19			12,49		
	(-)	$\pm$ 6,27			$\pm$ 5,4		
1400	Covid	15,16	-0,754	0,454	16,4 $\pm$	-0,624	0,536
Hz	(+)	$\pm$ 6,79			7,05		
	Covid	16,59			17,76		
	(-)	$\pm$ 7,02			$\pm$ 8,69		
2000	Covid	16,79	-1015	0,315	16,37	-0,993	0,326
Hz	(+)	$\pm$ 5,87			$\pm$ 7,38		
	Covid	18,56			18,36		
	(-)	$\pm$ 6,78			$\pm$ 7,22		
2800	Covid	18,08	-1091	0,280	16,63	-0,397	0,693
Hz	(+)	$\pm$ 6,89			$\pm$ 7,3		
	Covid	20,33			17,37		
	(-)	$\pm$ 8			$\pm$ 6,39		
4000	Covid	14,42	-1971	0,054	15,54	-0,895	0,375
Hz	(+)	$\pm$ 7,1			$\pm$ 6,2		
	Covid	18,67			17,29		
	(–)	$\pm$ 8,52			$\pm$ 7,9		

ss; standard deviation, a; significance test of the difference between two means (*t*-test).

 $^{\circ}$  p < 0,05 there is a statistically significant difference between groups.

nystagmus, head shake, dix Hallpike and head roll between the COVID-19 positive group and the control group. Since no pathological data group was found in the VNG gaze vertical, gaze horizontal, pursuit, dix Hallpike and head roll values, the comparison value could not be statistically calculated. No statistically significant difference was found between the COVID-19 positive and control groups in terms of their normal or pathological VNG saccade, optokinetic and spontaneous nystagmus values (p > 0.05). The normal and pathological VNG head shake values were found to be significantly different between the COVID-19 positive and control groups (p < 0.05, Table 5).

## 3.6. vHIT results

The mean values of vHIT lateral gain, LARP gain and RALP gain measured in both the right and left ears of the participants were examined. A statistically significant difference was found between the COVID-19 positive group and the control group in terms of the mean values of the vHIT lateral gain, LARP gain and RALP gain measurements in both the right and left ears (p < 0.05, Table 6). The mean values of vHIT lateral gain, LARP gain and RALP gain in both the right and left ears were found to be lower in the COVID-19 positive group than the control group (Table 6). No statistically significant difference was found between the COVID-19 positive group and the control group in terms of the mean value of the vHIT lateral gain measurement (p > 0.05, Table 7). A statistically significant difference was found between the COVID-19 positive group and the control group in the mean values of the vHIT LARP gain and RALP gain asymmetry measurements (p < 0.05, Table 7). The mean values of vHIT lateral gain, LARP gain and RALP gain asymmetry were calculated to be higher in the COVID-19 positive group in comparison to the control group (Table 7). Cross-tables were created, and Chi-squared ( $\chi^2$ ) analysis was performed to compare the values of vHIT lateral saccades, LARP saccades and RALP saccades between the COVID-19 positive and control groups. Since there was no data group with LARP saccades and RALP saccades in both the right and left ears, the comparison value could not be calculated statistically (Table 8). A statistically significant relationship was found between the presence of lateral saccades in the patients in both the right and left ears and the groups (p < 0.05, Table 8).

## 3.7. cVEMP results

The cVEMP P1 latency, N1 latency, P1-N1 latency and P1-N1 amplitude mean values were compared. A statistically significant difference was found between the COVID-19 positive and control groups according to the mean values of the left ear P1 latency and N1 latency measurements (p < 0.05, Table 9), but no statistically significant difference was found in the mean values of the P1 latency and N1 latency measurements in the right ear.

A statistically significant difference was found between the COVID positive and control groups in terms of the mean values of the right ear P1N1 amplitude measurement (p < 0.05, Table 9). According to the mean values of the P1N1 amplitude measurement in the left ear, no statistically significant difference was found between the COVID positive and control groups (p > 0.05, Table 9). A statistically significant difference was found between the COVID positive and control groups in the mean values of the P1N1 latency measurement in both the right and left ears (p < 0.05, Table 9).

The CVEMP N1 latency, P1N1 latency and P1N1 amplitude mean values were found to be lower in the COVID-19 positive patients than the control group. However, the P1 Latency measurement value was calculated to be higher in the right ear in the COVID-19 positive patients in comparison to the control group, while being lower in the left ear (Table 9). A statistically significant difference was found between the COVID-19 positive and control groups according to the mean value of CVEMP asymmetry (p < 0.05, Table 10). It was calculated that the mean CVEMP asymmetry value was higher in the COVID-19 positive patients

#### Table 4

Comparison of Bedside Values According to Groups.

	Group		Group	Group		Test value <sup>a</sup>	p value
			Covid (+)	Covid (–)			
Bedside Romberg	Normal	Number	25 <sub>a</sub>	27 <sub>a</sub>	52	1444	0,229
		%	96,2	100,0	98,1		
	Pathological	Number	1 <sub>a</sub>	0 <sub>a</sub>	1		
		%	3,8	0,0	1,9		
Bedside Tandem Romberg	Normal	Number	21 <sub>a</sub>	27 <sub>b</sub>	48	7665	0,006*
		%	80,8	100,0	90,6		
	Pathological	Number	5 <sub>a</sub>	0 <sub>b</sub>	5		
		%	19,2	0,0	9,4		
Bedside Fukuda	Normal	Number	23 <sub>a</sub>	27 <sub>b</sub>	50	4,46	0,035*
		%	88,5	100,0	94,3		
	Pathological	Number	3 <sub>a</sub>	0 <sub>b</sub>	3		
		%	11,5	0,0	5,7		
Bedside Tandem Gait	Normal	Number	21 <sub>a</sub>	27 <sub>b</sub>	48	7665	0,006*
		%	80,8	100,0	90,6		
	Pathological	Number	5 <sub>a</sub>	0 <sub>b</sub>	5		
	-	%	19,2	0,0	9,4		

 $^{*}$  p < 0.05; different letters in the rows show the difference between the two groups, while the same letters show that there is no difference a;  $\chi 2$  test value.

#### Table 5

Comparison of VNG values according to groups.

			Group	Group		Test value	p value
			Covid (+)	Covid (–)			
VNG gaze vertical	Normal	Number	26 <sub>a</sub>	27 <sub>a</sub>	53		
-		%	100,0	100,0	100,0		
VNG gaze horizontal	Normal	Number	26 <sub>a</sub>	27 <sub>a</sub>	53		
		%	100,0	100,0	100,0		
VNG saccade	Normal	Number	24 <sub>a</sub>	27 <sub>a</sub>	51	2930	0,087
		%	92,3	100,0	96,2		
	Pathological	Number	$2_a$	0 <sub>a</sub>	2		
		%	7,7	0,0	3,8		
VNG pursuit	Normal	Number	26 <sub>a</sub>	27 <sub>a</sub>	53		
		%	100,0	100,0	100,0		
VNG optokinetics	Normal	Number	23 <sub>a</sub>	27 <sub>a</sub>	50	3302	0,069
		%	88,5	100,0	94,3		
	Pathological	Number	3 <sub>a</sub>	0 <sub>a</sub>	3		
		%	11,5	0,0	5,7		
VNG spontaneous nystagmus	Normal	Number	23 <sub>a</sub>	27 <sub>a</sub>	50	3302	0,069
		%	88,5	100,0	94,3		
	Pathological	Number	3 <sub>a</sub>	0 <sub>a</sub>	3		
		%	11,5	0,0	5,7		
VNG head shake	Normal	Number	19 <sub>a</sub>	27 <sub>b</sub>	46	8735	0,001*
		%	73,1	100,0	86,8		
	Pathological	Number	7 <sub>a</sub>	0 <sub>b</sub>	7		
		%	26,9	0,0	13,2		
VNG dix hallpike	Normal	Number	26 <sub>a</sub>	27 <sub>a</sub>	53		
-		%	100,0	100,0	100,0		
VNG head roll	Normal	Number	26 <sub>a</sub>	27 <sub>a</sub>	53		
		%	100,0	100,0	100,0		

 $^{*}$  p < 0.05; different letters in the rows show the difference between the two groups, while the same letters show that there is no difference a;  $\chi 2$  test value.

## Table 6

Comparison of groups according to vHIT values.

	Group	Right ear	Right ear			Left ear		
		Mean $\pm$ ss	Test	p value	Mean $\pm$ ss	Test	p value	
vHIT Lateral Gain	Covid (+) Covid (–)	$\begin{array}{c} 0,\!88 \pm 0,\!13 \\ 0,\!96 \pm 0,\!06 \end{array}$	-2978	0,005*	$\begin{array}{c}\textbf{0,86}\pm\textbf{0,12}\\\textbf{1}\pm\textbf{0,06}\end{array}$	-5314	0,001*	
vHIT Larp Gain	Covid (+) Covid (-)	$\begin{array}{c} 0,\!79\pm0,\!16\\ 0,\!92\pm0,\!09 \end{array}$	-3450	0,001*	$\begin{array}{c} 0,83 \pm 0,16 \\ 0,94 \pm 0,09 \end{array}$	-3051	0,004*	
vHIT Ralp Gain	Covid (+) Covid (-)	$\begin{array}{c} \textbf{0,84} \pm \textbf{0,15} \\ \textbf{0,91} \pm \textbf{0,09} \end{array}$	-2093	0,043*	$\begin{array}{c}\textbf{0,73}\pm\textbf{0,18}\\\textbf{0,87}\pm\textbf{0,1}\end{array}$	-3564	0,001*	

ss; standard deviation, a; significance test of the difference between two means (t-test).

\* p < 0.05 there is a statistically significant difference between groups.

#### Table 7

Comparison of groups a	ccording to vHIT	asymmetry values.
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	Group	Asymmetry	Asymmetry		
		Mean $\pm$ ss	Test	p value	
vHIT lateral	Covid (+) Covid (–)	$4 \pm 3,42 \\ 3,15 \pm 2,28$	1071	0,289	
vHIT larp	Covid (+) Covid (-)	$5\pm4,04$ $3\pm2,81$	2098	0,041*	
vHIT ralp	Covid (+) Covid (-)	$\begin{array}{c} 8,54 \pm 6,75 \\ 3,67 \pm 3,77 \end{array}$	3227	0,003*	

ss; standard deviation, a; significance test of the difference between two means (*t*-test).

p < 0.05 there is a statistically significant difference between groups.

## (Table 10).

#### 3.8. oVEMP results

The mean values of the oVEMP P1 latency, N1 latency, P1N1 latency and P1N1 amplitude measurements made in both the right and left ears of the participants were evaluated. No statistically significant difference was found between the COVID-19 positive and control groups according to the mean values of the P1 latency, N1 latency, P1N1 latency and P1N1

#### Table 8

Comparison of vHIT saccade values according to groups

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amplitude measurements in both the right and left ears (p > 0.05, Table 9). It was found that the mean values of oVEMP P1 latency, N1 latency and P1N1 amplitude in both the right and left ears were higher in the COVID-19 positive group, and the P1N1 latency measurements were found to be higher in the control group (Table 9).

A statistically significant difference was found between the mean oVEMP asymmetry values of the COVID-19 positive group and the control group (p < 0.05). The mean OVEMP asymmetry value was found to be higher in the COVID-19 positive group in comparison to the control group (Table 10).

Table 10
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Comparison of c VEMP and oVEMP asymmetry values according to groups.

	Group	Asymmetry			
		Mean $\pm$ ss	Test	p value	
cVEMP	Covid (+) Covid (–)	$\begin{array}{c} 24,35 \pm 15,22 \\ 11,73 \pm 8,31 \end{array}$	3424	0,002*	
oVEMP	Covid (+) Covid (-)	$\begin{array}{c} \textbf{28,6} \pm \textbf{20,12} \\ \textbf{13,77} \pm \textbf{10} \end{array}$	3094	0,004*	

ss; standard deviation, a; significance test of the difference between two means (*t*-test).

p<0,05 there is a statistically significant difference between groups.

Ear		Group		Group	Group		Test <sup>a</sup>	p value
				Covid (+)	Covid (–)			
RİGHT	vHIT Lateral Saccade	+	Number	4 <sub>a</sub>	0 <sub>b</sub>	4	4496	0,034*
			%	15,4	0,0	7,5		
		-	Number	22 <sub>a</sub>	27 <sub>b</sub>	49		
			%	84,6	100,0	92,5		
	vHIT Larp Saccade	+	Number	26 <sub>a</sub>	27 <sub>a</sub>	53		
			%	100,0	100,0	100,0		
	vHIT Ralp Saccade	-	Number	26 <sub>a</sub>	27 <sub>a</sub>	53		
			%	100,0	100,0	100,0		
LEFT	vHIT Lateral Saccade	+	Number	5 <sub>a</sub>	0 <sub>b</sub>	5	7665	0,006*
			%	19,2	0,0	9,4		
		-	Number	21 <sub>a</sub>	27 <sub>b</sub>	48		
			%	80,8	100,0	90,6		
	vHIT Larp Saccade	_	Number	26 <sub>a</sub>	27 <sub>a</sub>	53		
			%	100,0	100,0	100,0		
	vHIT Ralp Saccade	-	Number	26 <sub>a</sub>	27 <sub>a</sub>	53		
	-		%	100,0	100,0	100,0		

 $^{*}$  p < 0.05; different letters in the rows show the difference between the two groups, while the same letters show that there is no difference a;  $\chi 2$  test value.

## Table 9

Comparison of groups according to cVEMP and oVEMP values.

	Group	Rigt ear			Left ear		
		Mean $\pm$ ss	Test	p value	Mean $\pm$ ss	Test	p value
P1 Latans	Covid (+)	$13{,}83\pm1{,}02$	0,585	0,562	$13{,}43 \pm 1{,}22$	-2780	0,008*
cVEMP	Covid (-)	$13{,}58\pm1{,}77$			$14,56 \pm 1,66$		
N1 Latans	Covid (+)	$\textbf{20,37} \pm \textbf{1,53}$	-2008	0,050	$\textbf{20,26} \pm \textbf{1,5}$	-4723	0,001*
cVEMP	Covid (-)	$\textbf{21,36} \pm \textbf{1,88}$			$22,55 \pm 1,91$		
P1N1 Latans cVEMP	Covid (+)	$\textbf{6,55} \pm \textbf{1,38}$	-3280	0,002*	$\textbf{6,83} \pm \textbf{1,53}$	-2833	0,007*
	Covid (-)	$\textbf{7,78} \pm \textbf{1,27}$			$\textbf{7,98} \pm \textbf{1,37}$		
P1N1 Amplitude	Covid (+)	$57,5 \pm 25,94$	-3363	0,002*	$73,23 \pm 19,45$	-1923	0,060
cVEMP	Covid (-)	$82,\!17 \pm 25,\!79$			$\textbf{86,3} \pm \textbf{27,78}$		
P1 Latans	Covid (+)	$10{,}25\pm0{,}91$	1981	0,053	$10{,}15\pm0{,}8$	1077	0,288
oVEMP	Covid (-)	$9{,}82\pm0{,}64$			$9{,}92\pm0{,}65$		
N1 Latans	Covid (+)	$\textbf{15,24} \pm \textbf{0,93}$	0,426	0,672	$15,46 \pm 1,05$	0,023	0,982
oVEMP	Covid (-)	$15,\!13 \pm 0,\!94$			$15,45 \pm 1,11$		
P1N1 Latans oVEMP	Covid (+)	$\textbf{4,98} \pm \textbf{0,63}$	-1738	0,089	$5,31\pm0,69$	-1125	0,266
	Covid (-)	$5{,}31\pm0{,}69$			$5,54\pm0,73$		
P1N1 Amplitude	Covid (+)	$\textbf{9,98} \pm \textbf{8,79}$	0,551	0,585	$10{,}88 \pm 7{,}65$	1100	0,281
oVEMP	Covid (–)	$\textbf{8,88} \pm \textbf{4,49}$			$\textbf{8,94} \pm \textbf{3,41}$		

ss; standard deviation, a; significance test of the difference between two means (t-test).

 $^{*}\,\,p<0,05$  there is a statistically significant difference between groups.

### 4. Discussion

The SARS-CoV-2 pandemic that started in December 2019 has caused the deaths of over 2 million people. Fever, cough, sore throat, respiratory failure, taste and smell disorders are common clinical symptoms in patients [16]. Neurotrophic features for the coronavirus family have been described so far [15]. Among otoneurologic symptoms, patients with hearing loss, tinnitus and dizziness have been reported. This situation may be directly related to neural tissue invasion or vasculitis [17]. Although the average of all frequencies was found within the normal limits in the hearing assessment of patients by audiological and TEOAE performed in patients who had had COVID-19 and recovered, a significant difference was found, especially at 4000 Hz and 8000 Hz, in comparison to the control group. Although the SSO value was 11.12  $\pm$ 5.78 dB for the right ear and 8.58  $\pm$  5.32 dB for the left ear within normal limits, it was calculated higher than the control group. In the TEOAE results, no significant difference was found in the control group in the mean frequency measurements of 1000 Hz, 1400 Hz, 2000 Hz, 2800 Hz and 4000 Hz, but the mean values were found to be low at all frequencies. When we look at the published literature, sudden hearing loss was shown in a 67-year-old COVID-19 patient, and it was treated with steroids [18]. In another study, audiometry and TEOAE were performed on patients with asymptomatic COVID-19. Significant results were obtained at 4000 Hz, 6000 Hz and 8000 Hz in audiometry. The decrease in TEOAE was found to be significant in comparison to the control group [19]. These studies similarly supported the information that cochlear hair cells may be affected by COVID-19 infection. COVID patients may have disturbances in the vestibular system. In the VNG tests of the COVID-19 positive patients, pathological findings were found in 3 patients in the head shake test. In the vHIT test, the gain was found to be lower in all channels in comparison to the control group. The RALP and LARP asymmetry among the vHIT asymmetry values were found to be significantly different in comparison to the control group. When we look at the saccades in vHIT, a significant difference between the groups was found only in the lateral canal. The fact that saccades were seen together with the losses in the VOR gains suggested that there was a significant influence in the semicircular canals or afferent ways [20]. In cVEMP, a significant difference was found based on the value of the control group in the asymmetry values. Additionally, a statistically significant difference was found between the groups' P1, N1 and P1-N1 latencies and p1-n1 amplitudes. A significant difference was found in the asymmetry values in oVEMP, but no significant changes were observed in the amplitude and latency values. Changes in the cVEMP latencies and amplitudes of the patients with COVID suggested that COVID affects the brainstem and vestibulocollic arc and slows down the communication on the arc. These impacts were shown to occur in the retro-labyrinth lesions in the vestibulospinal pathway in the literature. The lack of large differences in the gain asymmetries in VEMPs revealed the importance of the compensation mechanism of the central vestibular system [21,22]. Moreover, the difference in the patients with COVID in the bedside tests drew attention to the effect of COVID on the vestibulospinal arc and postural balance. When we look at COVID-19 and vestibular system evaluation in the literature, in a study evaluating the vestibular system in 185 patients through online questions, balance disorders were detected in 34 patients after their diagnosis of COVID-19. Among these patients, 32 patients reported dizziness (94.1%), and 2 (5.9%) reported acute vertigo attacks. For balance disorders, the mean VAS score was calculated as 5 [17]. In another case report, a case of vestibular neuritis that could be associated with COVID-19 with nausea and vomiting was presented [23]. In another review article, vestibular complaints were reported in 7 patients diagnosed with COVID-19 infection, but a direct vestibular origin was not mentioned [24]. Although scientific knowledge on COVID-19 is increasing, the information in the audiovestibular literature is still limited. Researchers have not been able to address the life-threatening symptoms of COVID-19, as well as associated hearing and balance problems. The former members

of the coronavirus family (MERS and SARS) have had effects on the hearing and balance system. SARS-CoV-2 may have direct neurological involvement or inner ear involvement due to widespread hypercoagulation recently seen in COVID-19 patients. Vascular involvement may be one of the clinical signs of COVID-19 such as various viral infections including hepatitis B and C vasculitis [25]. Audiovestibular system disorders may occur due to vascular damage because the inner ear is very sensitive to ischemia [26]. Primary and secondary vasculatures may be associated with hearing and vestibular symptoms [27]. This study is a comprehensive study which evaluated both the auditory and vestibular systems and included a control group. The vestibular system was evaluated with a large battery of tests. In terms of hearing, statistically significant results were obtained at higher frequencies in comparison to the control group. In the vestibular system, especially in oVEMP and cVEMP, asymmetric findings were obtained in comparison to the control group, and a low gain in vHIT was shown.

## 5. Conclusion

The high frequencies in audiometry in the COVID-19 positive group were worse than those in the control group. In the vestibular system, especially in oVEMP and cVEMP, asymmetric findings were obtained in comparison to the control group, and a low gain in vHIT was shown. This study shows that the audiovestibular system of people with COVID-19 infection may be affected. However, the support of this study with series with higher numbers of patients will be thanks to future studies.

## CRediT authorship contribution statement

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