

## The Importance of neutrophil-lymphocyte ratio, monocyte-lymphocyte ratio, and thrombocyte-lymphocyte ratio in addition to abnormal laboratory parameters in COVID-19 pneumonia

*Anormal laboratuvar parametrelerine ilave olarak, nötrofil-lenfosit, monosit-lenfosit ve trombosit-lenfosit oranlarının COVID-19 pnömonisindeki önemi*

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### ABSTRACT

**Aim:** We aimed to examine the importance of the neutrophil-lymphocyte ratio (NLR), monocyte-lymphocyte ratio (MLR) and thrombocyte-lymphocyte ratio (TLR) in addition to laboratory parameters in terms of pneumonia between patients with and without pneumonia diagnosed with COVID-19 infection.

**Materials and Methods:** In our retrospective study, 506 patients diagnosed with COVID-19 who were hospitalized and treated between April 2020 and July 2020 were included. While 50.6% (n= 256) of the patients had pneumonia, 49.4% (n= 250) did not have. File informations were scanned; the demographic data, hospitalization and discharge laboratory parameters were recorded.

**Results:** The median age of the patients was 44.5 (16-89) years, 55.5% (n= 281) of them were male. The median age of patients with pneumonia was higher than those without pneumonia, and the frequency of female gender was higher (p <0.05 for both conditions). The lymphocyte count at hospitalization and discharge was significantly lower in patients with pneumonia (p <0.001, 0.007, respectively). The count of eosinophils at hospitalization was significantly lower in patients with pneumonia (p <0.001). NLR and TLR values were significantly higher in patients with pneumonia at admission and discharge (p <0.008, <0.007, respectively for NLR and p <0.001, p<0.001 respectively for TLR). MLR values at discharge were higher in patients with pneumonia (p = 0.001). In ROC analyzes made in terms of predicting pneumonia; hospitalization CRP level, eosinophil, monocyte counts and TLR values were found to be important variables.

**Conclusion:** In this study, the importance of monocyte, eosinophil counts and TLR values in COVID-19 pneumonia was demonstrated.

**Keywords:** COVID-19 pnömonisi, monosit, lenfosit, eozinofil, NLO, MLO, TLO.

### ÖZ

**Amaç:** COVID-19 pnömonisi olan ve olmayan hastalar arasındaki klinik ve hematolojik parametrelerdeki farklılıkların irdelenmesi, yatış ve taburculuk değerlerinin kıyaslanması, laboratuvar parametrelerinin yanında nötrofil-lenfosit oranı (NLO), monosit-lenfosit oranı (MLO) ve trombosit-lenfosit oranı (TLO)' nın pnömoni açısından öneminin karşılaştırılması amaçlandı.

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**Gereç ve Yöntem:** Retrospektif yaptığımız çalışmada Enfeksiyon Hastalıkları servisinde Nisan 2020-Temmuz 2020 tarihleri arasında yatarak tedavi alan COVID-19 tanılı 506 hasta dahil edildi. Hastaların 256'sı pnömoni tablosu olan (%50,6), 250'si pnömoni tablosu olmayan (%49,4) COVID-19 tanılı olgular idi. Her iki hasta grubunda dosya bilgileri taranarak demografik veriler, yatış ve taburculuklarındaki laboratuvar parametreleri, NLO, MLO ve TLO irdelenip karşılaştırıldı.

**Bulgular:** Hastaların medyan yaşı 44,5 (16-89) yıl olup, %55,5 (n= 281)'i erkek idi. Pnömonisi olan hastaların medyan yaşı pnömonisi olmayanlara göre daha yüksekti ve bu grupta kadın cinsiyet sıklığının daha fazla olduğu görüldü (her iki durumda  $p<0,05$ ). Pnömonisi olanların hastanede ortanca kalış süresi pnömoni olmayanlara göre daha yüksekti ( $p:0,009$ ). Pnömonisi olanlarda olmayanlara göre yatış ve taburculuk lenfosit sayısı anlamlı olarak daha düşük idi ( $p$  değeri sırasıyla:  $<0,001$ ,  $0,007$ ). Pnömonisi olanlarda olmayanlara göre yatış eozinofil sayısı anlamlı olarak daha düşük gözlemlendi ( $p<0,001$ ). Pnömonisi olan hastaların yatış ve taburculuk NLO pnömonisi olmayan hastalara göre anlamlı olarak daha yüksek gözlemlendi ( $p$  sırasıyla  $<0,008$ ,  $<0,007$ ). Her iki grupta taburculuk NLO değeri yatış değerine göre düşük saptandı (her iki durumda  $p$ ;  $<0,001$ ). TLO pnömonisi olanlarda olmayanlara göre hem yatış hem de taburculukta daha yüksek görüldü (her iki durumda  $p <0,001$ ). MLO incelendiğinde taburculuk MLO değerleri pnömonisi olanlarda daha yüksek gözlemlendi ( $p=0,001$ ). Pnömoniyi öngörme açısından yapılan ROC analizinde; yatış eozinofil değerinin  $\leq 0,02/\mu\text{L}$  olmasının AUC:0,668, %56,3 sensitivite, %71,2 spesifiteye, monosit değerinin  $\leq 0,5/\mu\text{L}$  olmasının AUC:0,631, %55,1 sensitivite, %67,6 spesifiteye, TLO değerinin  $>105,7$  olmasının AUC:0,611, %75,4 sensitivite, %46,0 spesifiteye sahip olduğu gözlemlendi.

**Sonuç:** Bu çalışmada özellikle monosit, eozinofil sayıları ve TLO oranlarının COVID-19 pnömonisindeki önemi gösterilmiş oldu.

**Anahtar Sözcükler:** COVID-19 pneumonia, monocyte, eosinophil counts, NLR, TLR, MLR.

## INTRODUCTION

While the rapid course and increased mortality of COVID-19 infection has been remarkable, the new type of coronavirus called as SARS-CoV2, which has different characteristics from other coronaviruses (SARS-CoV and MERS-CoV) was discovered as the causative agent of the disease. Worldwide more than 265.000.000 people have been infected with the virus and more than 5.000.000 deaths have been experienced since December 2019 due to the COVID-19 infection, which was accepted as a pandemic by the WHO on March 12, 2020.

The inflammatory and pro-thrombotic process, which starts with the damage of the virus in the endotelium leads to multisystem circulatory disorders especially in the pulmonary circulation, finally multipl tissue and organ disfunctions occur [1].

The virus affects natural and acquired immune system cells by directly cell damage or indirectly through cytokines and chemokines. As a result, while lymphocytes and eosinophils decrease, an increase in neutrophil and monocyte counts can be observed (1, 2).

In many studies on COVID-19 elevated neutrophil-lymphocyte ratio (NLR) has been reported which is used in the follow-up of many solid tumors, bacterial infections, acute

pancreatitis and cardiovascular diseases as an inflammatory marker and prognostic factor (3). In addition, there are COVID-19 studies showing an increase in monocyte-lymphocyte ratio (MLR) and thrombocyte-lymphocyte ratio (TLR) (4-6).

However, there are not enough studies about COVID-19 pandemia comparing the laboratory parameters between patients with and without pnemonia. We aimed to investigate whether there are differences in laboratory parameters, NLR, MLR, and TLR values between patients with and without pneumonia in the course of COVID-19 infection.

## MATERIALS and METHODS

In this retrospective study 506 patients diagnosed with COVID-19 were included who received inpatient treatment between April 2020 and July 2020 in the Infection Service of Konya Training and Research Hospital. Among the patients hospitalized between the dates mentioned, those with confirmed diagnosis of Covid-19 were included in the study. The demographic characteristics (age and gender) of the patients on the hospitalization day, as well as the laboratory and radiologic examinations results were recorded. These examinations included COVID-19 RT-PCR test, thorax CT, hemogram, CRP, ferritin, kidney and liver function tests. In addition, the hemogram results at discharge were also recorded.

In patients whose COVID-19 infection was confirmed with a positive COVID-19 RT-PCR test, the pneumonia status was confirmed by chest CT in addition to clinical and laboratory findings. From the hemogram parameters, NLR obtained by dividing the neutrophil count ( $\times 10^3/\mu\text{L}$ ) by the count of lymphocytes ( $\times 10^3/\mu\text{L}$ ), MLR obtained by dividing the monocyte count ( $\times 10^3/\mu\text{L}$ ) by the count of lymphocytes ( $\times 10^3/\mu\text{L}$ ) and the TLR values obtained by dividing the platelet count ( $\times 10^3/\mu\text{L}$ ) by the count of lymphocytes ( $\times 10^3/\mu\text{L}$ ) were calculated. Primarily, we have received approval from the Ministry of Health of the Republic of Turkey established scientific studies. In addition, data collection procedures started after obtaining approval from the Medical Specialization Training Board (with the letter dated 02.07.2020 and numbered 48929119/774) of Konya Training and Research Hospital.

### Statistical Analysis

Analytical and graphical methods were used to evaluate the data in terms of normal distribution. The normal distribution of the data was determined using skewness-kurtosis values, Kolmogorov-Smirnov test and variance coefficient from analytical methods, and graphical methods with histogram and detrended Q-Q plot graphs. Mann Whitney U test was used to compare numerical variables between two groups. It was observed that all numerical parameters in the study did not show normal distribution and numerical parameters were shown with median (min-max). In comparison of categorical variables; Fisher's exact test or Chi-square test was used according to suitability. Wilcoxon test was used to evaluate the changes in the patients' admission and discharge laboratory values within the groups. MedCalc program was used to perform ROC analysis to evaluate the cut-off values, sensitivity and specificity of biochemical and hemogram parameters that best predict the status of COVID pneumonia in the course of infection. The Statistical Package for the Social Sciences software version 25.0 (IBM SPSS Corp.; Armonk, NY, USA) was used for statistical calculations. A p value of  $<0.05$  was accepted for statistical significance.

## RESULTS

### Demographic Characteristics of Patients:

A total of 506 patients diagnosed with COVID-19 treated in the infection service, 50.6% of whom

with pneumonia and 49.4% without pneumonia were included. While 55.5% ( $n = 281$ ) of the patients were male, 44.5% ( $n = 225$ ) were female and 52% of those with pneumonia were women. The median age was 44.5 (16-89) years, and it was 51 (16-84) years in patients with pneumonia ( $p < 0.001$ ). While additional comorbid diseases were observed more in patients with pneumonia, complaints of fever, dry cough, shortness of breath, sputum and weakness were significantly higher in this group ( $p < 0.001$  for both).

### Clinical and Laboratory Parameters:

While the median hospitalization day of all patients was 5 (4-26) days, it was observed longer in patients with pneumonia than those without pneumonia ( $p = 0.009$ ).

The leukocyte count ( $\times 10^3/\mu\text{L}$ ) at hospitalization was significantly lower in those with pneumonia ( $p = 0.006$ ). The lymphocyte count ( $\times 10^3/\mu\text{L}$ ) at hospitalization and discharge was significantly lower in patients with pneumonia ( $p < 0.001$ ,  $0.007$ , respectively). The median value of eosinophil count ( $\times 10^3/\mu\text{L}$ ) at hospitalization was lower in patients with pneumonia ( $p < 0.001$ ), and a significant increase at discharge was observed in all patients ( $p < 0.001$ ). Monocyte counts ( $\times 10^3/\mu\text{L}$ ) at hospitalization were significantly lower in those with pneumonia ( $p < 0.001$ ). In all patients the RBC values ( $\times 10^6/\mu\text{L}$ ) at hospitalization were significantly higher than the values measured at discharge ( $p < 0.001$ ). In patients with pneumonia both hospitalization and discharge Hb values (gr/dL) were lower (for both  $p < 0.001$ ). The comparison of hemogram parameters has been shown in (Table-1).

It was observed that both NLR and TLR values were higher in patients with pneumonia at both hospitalization and discharge (respectively  $p < 0.001$ ,  $p < 0.001$  for both values and both conditions). In all patients MLR decreased significantly at discharge ( $p < 0.001$ ). The important values of NLR, MLR and TLR have been compared in (Table-2).

Patients with pneumonia had significantly higher CRP and ferritin levels (respectively  $p < 0.001$ ,  $p = 0.003$ ) (Table-3).

The comparison of important laboratory levels at hospitalization and discharge of patients with and without pneumonia has been shown in (Table-4).

**Table-1.** Comparison of the hemogram parameters of COVID patients according to pneumonia status.

	<b>Total N=506</b>	<b>Without pneumonia N=250</b>	<b>With pneumonia N=256</b>	<b>p value</b>
<b>leukocyte at admission</b>	5.8 (1.9-48.8)	6 (1.9-48.8)	5.6 (1.9-20.1)	0.006
<b>leukocyte at discharge</b>	6.1 (2.3-44.6)	6.2 (2.5-44.6)	6 (2.3-29.5)	0.471
<b>neutrophil at admission</b>	3.4 (0.7-41.8)	3.5 (1.1-41.8)	3.4 (0.7-17.6)	0.447
<b>neutrophil at discharge</b>	3.3 (0.7-39.8)	3.3 (0.7-39.8)	3.4 (0.9-27.1)	0.509
<b>lymphocyte at admission</b>	1.6 (0.2-5.1)	1.7 (0.2-5.1)	1.4 (0.4-4.5)	<0.001
<b>lymphocyte at discharge</b>	2 (0.2-6)	2.1 (0.2-6)	1.9 (0.4-4.9)	0,007
<b>eosinophil at admission</b>	0 (0-0.8)	0.1 (0-0.8)	0 (0-0.4)	<0.001
<b>eosinophil at discharge</b>	0.1 (0-2.5)	0.1 (0-2.5)	0.1 (0-0.5)	0.090
<b>RBC at admission</b>	4.8 (2.2-6.3)	4.8 (3.5-6.2)	4.7 (2.2-6.3)	0.233
<b>RBC at discharge</b>	4.6 (1.7-6.4)	4.7 (2.9-6.4)	4.5 (1.7-6.1)	0.004
<b>Hb at admission</b>	13.9 (7.7-18.3)	14.4 (8-18.3)	13.5 (7.7-17.8)	<0.001
<b>Hb at discharge</b>	13.4 (5.9-18.4)	13.9 (8.3-18.4)	12.9 (5.9-17.1)	<0.001
<b>thrombocyte at admission</b>	202.5 (43-644)	202 (61-644)	203 (43-480)	0.940
<b>thrombocyte at discharge</b>	246 (52-697)	228 (53-697)	268.5 (52-589)	<0.001
<b>monocyte at admission</b>	0.5 (0.1-8.4)	0.6 (0.1-2.3)	0.5 (0.2-8.4)	<0.001
<b>monocyte at discharge</b>	0.6 (0-2)	0.5 (0-2)	0.6 (0.1-1.9)	0.279

Leukocyte, neutrophil, lymphocyte, eosinophil, monocyte and thrombocyte units: ( $\times 10^3 /\mu\text{L}$ ), RBC unit: ( $\times 10^6 /\mu\text{L}$ ), Hb unit: gr/dL

**Table-2.** Comparison of the NLR, MLR and TLR levels of COVID patients according to pneumonia status.

	<b>Total N=506</b>	<b>Without pneumonia N=250</b>	<b>With pneumonia N=256</b>	<b>p value</b>
<b>NLR at admission</b>	2 (0.4-26.7)	1.9 (0.5-26.7)	2.3 (0.4-23.8)	0.008
<b>NLR at discharge</b>	1.7 (0.5-56.5)	1.5 (0.5-17.5)	1.8 (0.5-56.5)	0.007
<b>TLR at admission</b>	128.9 (14.1-1160)	114.5 (37.9-1160)	143.1 (14.1-516.2)	<0.001
<b>TLR at discharge</b>	125.9 (15.3-663.8)	106.2 (31.5-663.8)	138 (15.3-559.3)	<0.001
<b>MLR at admission</b>	0.33 (0.1-3.9)	0.34 (0.1-3.4)	0.33 (0.1-3.9)	0.769
<b>MLR at discharge</b>	0.28 (0-3.9)	0.27 (0-2.2)	0.29 (0.1-3.9)	0.001

NLR: neutrophil count/lymphocyte count, TLR: thrombocyte count/lymphocyte count , MLR: monocyte count/lymphocyte count

**Table-3.** Biochemistry parameters of COVID patients according to pneumonia status.

	Total N=506	Without pneumonia N=250	With pneumonia N=256	p value
<b>AST</b>	26 (10-104)	25 (10-104)	27 (11-90)	0.011
<b>ALT</b>	20 (4-268)	20 (4-268)	21 (6-91)	0.495
<b>Creatinine</b>	0.9 (0.4-8)	0.9 (0.4-3.6)	0.9 (0,5-8)	0.235
<b>Urea</b>	27 (10.3-120)	25 (10.3-104)	29 (12.2-120)	<0.001
<b>CRP</b>	7 (3-271)	4.3 (3-236.7)	10.6 (3.1-271)	<0.001
<b>Ferritin</b>	77 (3.4-3927)	67 (4.6-3927)	98.5 (3.4-1650)	0.003

AST, ALT units: U/L, ürea and creatinine unit: mg/dL, CRP unit: mg/L, ferritin unit: ng/mL

**Table-4.** Differences between important laboratory values of COVID patients at admission and discharge.

	Total N=506	p value	Without pneumonia N=250	p value	With pneumonia N=256	p value
<b>Leukocyte at admission</b>	5.8 (1.9-48.8)	0.001	6 (1.9-48.8)	0.665	5.6 (1.9-20.1)	p<0.001
<b>Leukocyte at discharge</b>	6.1 (2.3-44.6)		6.2 (2.5-44.6)		6 (2.3-29.5)	
<b>Neutrophil at admission</b>	3.4 (0.7-41.8)	0.308	3.5 (1.1-41.8)	0.033	3.4 (0.7-17.6)	0.552
<b>Neutrophil at discharge</b>	3.3 (0.7-39.8)		3.3 (0.7-39.8)		3.4 (0.9-27.1)	
<b>Lymphocyte at admission</b>	1.6 (0.2-5.1)	p<0.001	1.7 (0.2-5.1)	p<0.001	1.4 (0.4-4.5)	p<0.001
<b>Lymphocyte at discharge</b>	2 (0.2-6)		2.1 (0.2-6)		1.9 (0.4-4.9)	
<b>Eosinophil at admission</b>	0 (0-0.8)	p<0.001	0.1 (0-0.8)	p<0.001	0 (0-0.4)	p<0.001
<b>Eosinophil at discharge</b>	0.1 (0-2.5)		0.1 (0-2.5)		0.1 (0-0.5)	
<b>RBC at admission</b>	4.8 (2.2-6.3)	p<0.001	4.8 (3.5-6.2)	p<0.001	4.7 (2.2-6.3)	p<0.001
<b>RBC at discharge</b>	4.6 (1.7-6.4)		4.7 (2.9-6.4)		4.5 (1.7-6.1)	
<b>Hb at admission</b>	13.9 (7.7-18.3)	p<0.001	14.4 (8-18.3)	p<0.001	13.5 (7.7-17.8)	p<0.001
<b>Hb at discharge</b>	13.4 (5.9-18.4)		13.9 (8.3-18.4)		12.9 (5.9-17.1)	
<b>Thrombocyte at admission</b>	202.5 (43-644)	p<0.001	202 (61-644)	p<0.001	203 (43-480)	p<0.001
<b>Thrombocyte at discharge</b>	246 (52-697)		228 (53-697)		268.5 (52-589)	
<b>Monocyte at admission</b>	0.5 (0.1-8.4)	0.915	0.6 (0.1-2.3)	p<0.001	0.5 (0.2-8.4)	p<0.001
<b>Monocyte at discharge</b>	0.6 (0-2)		0.5 (0-2)		0.6 (0.1-1.9)	
<b>NLR at admission</b>	2 (0.4-26.7)	p<0.001	1.9 (0.5-26.7)	p<0.001	2.3 (0.4-23.8)	p<0.001
<b>NLR at discharge</b>	1.7 (0.5-56.5)		1.5 (0.5-17.5)		1.8 (0.5-56.5)	
<b>TLR at admission</b>	128.9 (14.1-1160)	0.299	114.5 (37.9-1160)	0.293	143.1 (14.1-516.2)	0.746
<b>TLR at discharge</b>	125.9 (15.3-663.8)		106.2 (31.5-663.8)		138 (15.3-559.3)	
<b>MLR at admission</b>	0.33 (0.1-3.9)	p<0.001	0.34 (0.1-3.4)	p<0.001	0.33 (0.1-3.9)	p<0.001
<b>MLR at discharge</b>	0.28 (0-3.9)		0.27 (0-2.2)		0.29 (0.1-3.9)	

Leukocyte, neutrophil, lymphocyte, eosinophil, monocyte and thrombocyte units: ( $\times 10^3 /\mu\text{L}$ ), RBC unit: ( $\times 10^9 /\mu\text{L}$ ), Hb unit: gr/dL, NLR: neutrophil count/lymphocyte count, TLR: thrombocyte count/lymphocyte count, MLR: monocyte count/lymphocyte count

## ROC Analysis Results

In ROC analysis made in terms of predicting pneumonia in COVID-19 infection; CRP value of >3.93 mg/dL at hospitalization had AUC: 0.675 with 87.9% sensitivity and 47.6% specificity, the eosinophil value of  $\leq 0.02/\mu\text{L}$  had AUC: 0.668 with 56.3% sensitivity and 71.2% specificity, monocyte value of  $\leq 0.5/\mu\text{L}$  had AUC: 0.631 with 55.1% sensitivity and 67.6% specificity. TLR value of >105.7 had AUC: 0.611 with 75.4% sensitivity and 46% specificity, NLR value of > 1.59 had AUC: 0.569 with 73.8% sensitivity and 40% specificity. CRP, eosinophil, monocyte and TLR were determined as markers with high positive predictive parameters in favor of COVID-19 pneumonia.

## DISCUSSION

In this study, it has been shown that especially lymphocyte, monocyte, eosinophil counts and and TLR values may be important in COVID-19 pneumonia. It is thought that the comparison of the clinical and laboratory data of patients with and without pneumonia in the course of COVID-19 disease, as well as the determination of pneumonia predictive laboratory parameters will contribute to science. The lack of sufficient studies comparing these two groups during COVID-19 infection, reveals the significance and difference of the current study.

In a meta-analysis advanced age, obesity, and high fever were found to be independent variables predicting COVID-19 pneumonia (7). The importance of NLR in terms of prognosis and mortality has been frequently emphasized in studies (5, 6). The incidence of pneumonia in patients in the study was 50.6% (n = 256). The median age of patients with pneumonia was higher (p <0.05).

Many significant changes develop during the course of COVID-19 disease in hemogram and biochemical parameters. In most of the studies since the beginning of the epidemic, variables in predicting organ involvement and disease prognosis have been reported (8, 9). Leukopenia, leukocytosis and lymphopenia are the most frequently reported changes. Although leukocytosis is more common in severe disease, it has been reported to be around 25% (10, 1). Lymphopenia and neutrophilia have been reported as prognostic factors for severe disease in many studies (11). Eosinopenia is frequently reported in the course of COVID-19 and is has

been used as a prognostic marker for severe disease (9,12). Although the median leukocyte, lymphocyte, neutrophil and eosinophil counts of all patients diagnosed with COVID-19 in the study were mostly in the normal range, in the group with pneumonia the eosinophils and lymphocytes which frequently decrease in the course of infection especially in severe pneumonia, had been observed to be lower (p <0.05). Although the monocyte count generally increases during the inflammatory process, some studies comparing the healthy population with COVID-19 patients reported a slight increase in the patient group. However, in studies with large patient groups, monocyte count has been reported generally within normal limits (13, 14). In studies on COVID-19 thrombocytopenia has also been reported as a poor prognostic marker (15). In our study, the monocyte count of all patients was mostly within the normal range and it was compatible with the literature. Median monocyte count at hospitalization of patients with pneumonia were lower.

Although studies have reported that the platelet count decreases due to consumption and low values are associated with severe disease in the prothrombotic process which starts as a result of the damage by the virus in the endothelium and there are also studies reporting that reactive thrombocytosis develops as a result of inflammation (12, 16). While thrombocyte levels were within normal range in our patients, significant increment in thrombocyte count at discharge in both groups (p <0.001) can be explained by reactive thrombocytosis. The fact that the patients in the study were followed up in the clinic and the intensive care patients were not included may explain that laboratory parameters were mostly within the normal range in our patients.

It has been reported that a gradual decrease in RBC and Hb values has been observed within days in the course of the infection due to the damage developed in erythrocytes and suppression of erythropoiesis in bone marrow by the toxic effects of the virus on both mature and precursor cells (16, 17). During the recovery period of the infection, RBC and Hb values were reported as parameters that recovered at the latest (17). In the study, in all patients the lower RBC and Hb values at discharge (p <0.001 for both) were consistent with the literature.

Although NLR usually increases, especially in severe infection, it has been reported as one of the most important factors predicting disease prognosis and mortality (18). Although the normal NLR value was detected in the range of 0.78-3.53 in a healthy adult population in one study, there is no known universally accepted reference range (19). It has also been reported that the NLR varies according to gender and age. The median admission NLR value of the patients included in the study was 2 (0.4-26.7).

In addition, there are studies on COVID-19 showing that MLR and TLR values which are among the new inflammation markers, increase in the course of infection, especially in severe disease. These markers can be used as significant predictive variables in terms of mortality and prognosis in COVID-19 infection (20, 21). In a multi-center study, it has been determined that MLR is the most important marker to predict COVID-19 infection [21]. In the study, it was found that NLR and TLR values at hospitalization were higher in pneumonia group (p: 0.008, <0.001, respectively).

In studies, different cut-off values with predictive importance for NLR, TLR and MLR on poor prognosis and mortality of COVID-19 infection have been reported. In the comparison of the laboratory parameters of the two groups; lymphocyte, eosinophil, monocyte, CRP, NLR and TLR values were taken into consideration to predict Covid-19 pneumonia. It was found that CRP, eosinophil, monocyte and TLR were significant predictive markers in favor of Covid-19 pneumonia.

#### **Limitations of the study:**

Since the mortality rate was low in our patient population, the importance of these variables in terms of mortality and prognosis could not be evaluated.

Multi-center studies with larger patient populations are needed to clarify the course of COVID-19 pneumonia and to determine the important laboratory parameters in the diagnosis and follow-up.

**Conflict of interest:** There is no conflict of interest between the authors.

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