



Platelet to Lymphocyte Ratio and Acute Appendicitis

Platelet Lenfosit Oranı ve Akut Apandisit

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ABSTRACT

Aim: Acute appendicitis (AA) is a frequent condition for general surgery practice. However, sometimes there may be difficulties in accurate diagnosis. The aim of this study is to research the predictive value of platelet to lymphocyte ratio (PLR) in diagnosis of AA.

Material and Method: Hospital records were investigated retrospectively for 569 patients who underwent operation for AA diagnosis. According to postoperative histopathological examination of specimens, patients were divided into two groups: acute appendicitis group (G1) and normal appendix group (G2). Demographic analyses and PLR calculations on hospital admission were compared intergroup.

Results: There were 475 patients in G1 whereas 94 in G2. The cut-off value of PLR for diagnosis of AA was 136.5 ($p=0.036$). The sensitivity and specificity were 56.3% and 55.3% respectively. Positive predictive value and negative predictive value were found as 86.2% and 19.6% respectively.

Conclusion: PLR may be a valuable parameter supporting clinical evaluation for diagnosis and management of AA.

Key words: acute appendicitis; platelet to lymphocyte ratio

ÖZET

Amaç: Akut apandisit (AA) genel cerrahi pratiğinde sık karşılaşılan bir durumdur. Ancak doğru tanıya ulaşmada bazen zorluklar olabilmektedir. Bu çalışmanın amacı AA tanısında platelet lenfosit oranının (PLO) belirleyiciliğini araştırmaktır.

Materyal ve Metot: Akut apandisit tanısı ile ameliyat edilmiş 569 hastaya ait hastane kayıtları geriye dönük olarak incelendi. Postoperatif histopatolojik inceleme sonuçlarına göre hastalar iki gruba ayrıldı: Akut apandisit grubu (G1) ve normal apandiks grubu (G2). Gruplar arasında demografik analizler ve PLO hesaplamaları karşılaştırıldı.

Bulgular: G1 de 475, G2 de 94 hasta vardı. AA tanısında PLO için sınır değer 136,5 ($p=0,036$) olarak hesaplandı. Duyarlılık ve seçicilik sırası ile %56,3 ve %55,3 idi. Pozitif prediktif ve negatif prediktif değerler sırası ile %86,2 ve %19,6 olarak bulundu.

Sonuç: AA tanısında ve tedavinin yönlendirilmesinde PLO değeri bir ölçüt olabilir.

Anahtar kelimeler: akut apandisit; platelet lenfosit oranı

Introduction

Surgical management of acute appendicitis (AA) is one of the most commonly performed operations worldwide since its first description by Reginald Fitz in 1886¹. However, there are some recent studies about medical treatment of the disease². The lifetime occurrence of AA is 8%; clinical presentation varies from mild to severe and mortal condition. Fortunately, mortality is less than 1%³. Historically, management of the patient is based on typical history, physical findings, laboratory tests and imaging studies. Traditionally, negative appendectomy within reasonable limits is acceptable for fear of morbidity and mortality due to perforation. But today, it is not acceptable and in order to improve pre-operative diagnosis and to avoid negative laparotomy many scoring systems were described⁴⁻⁶. The neutrophil to lymphocyte ratio (NLR) and platelet to lymphocyte ratio (PLR) are associated with worse outcome in various diseases. And NLR is one of the newest parameters for AA^{7,8}. However we didn't find any other study that investigated platelet to lymphocyte ratio (PLR) for this purpose in English literature.

Material and Method

After approval of the hospital ethics committee a retrospective study was designed. The patients with AA

diagnosis in preoperative period were evaluated from the hospital database. The demographic characteristics of patients, laboratory test results on admission to the hospital, operative findings and postoperative pathology reports were enrolled. The study was performed in 33 months period from January 2011 to September 2013. Initially there were 645 patients with these criteria. But 76 patients were excluded from the study because of comorbidities such as malign or haematological disorders, recent blood transfusion history and missing files. We determined that they had different problems except AA. Thus 569 patients were included to the study. The diagnoses of AA were obtained with traditional patient history, physical examination, routine laboratory and radiologic tests and sometimes imaging studies such as ultrasonography and computed tomography. The platelet and lymphocyte counts were measured by an automated hematologic analyzer (*Coulter LH 780 Hematologic Analyzer, Beckman Coulter Inc. Brea, USA*). The calculations of PLRs were done from these data at the time of patient admission.

Totally, data of 569 patients were analyzed. Pathology reports were used to determine whether the appendix was inflamed (catarrhal, suppurative or gangrenous) or normal. Normal appendix has not any pathologic change in mucosa. Catarrhal appendicitis has raised neutrophil in mucosa. Inflammatory changes involve muscular and subserosal layers in suppurative appendicitis. In gangrenous appendicitis, there are vessel thrombosis and necrotic areas in appendix wall. Patients were divided in two groups according to histopathology of the specimens: positive appendectomy group (G1) and negative appendectomy group (G2).

Data analysis was performed by using SPSS for Windows, version 17.0 (SPSS Inc, Chicago, Illinois, USA). Whether the distributions of continuous variables were normally or not was determined by Kolmogorov-Smirnov Z test. Data were shown as

mean±standard deviation or median (min-max), where applicable. Categorical data were analyzed by Pearson's chi-square test, where appropriate. The differences between groups were compared by using Mann Whitney U test or Student's T, where appropriate. The cut-off values of parameters for discrimination of the groups were determined using the ROC analysis. At each value, the sensitivity and specificity for each outcome under study was plotted, thus generating an ROC curve. A p value less than 0.05 was considered statistically significant.

Results

Data of 569 patients were analyzed. Negative appendectomy ratio was 94/569 (16.5%). Median age of analyzed group was 29 (17–85) and 30.6% of the patients were female. There were 475 patients in G1 whereas 94 patients in G2. There were no significant difference between the groups according to age ($p=0.658$). In gender analyses female/male ratio was 128/347 in G1 whereas 46/48 in G2. Thus negative appendectomy rate was statistically higher in females ($p<0.001$). The findings were summarised in Table 1. The median PLR values in G1 and G2 were 146.5 (59.7–975.0) and 123 (28.4–497.8) respectively. PLR values were significantly different intergroups ($p=0.036$) (Fig. 1).

The recommended cut-off value of the PLR for positive and negative appendectomies was decided using receiver operating characteristic curve analyses. The recommended cut-off value of the PLR was based on the most prominent point on the receiver operating characteristic curve. This was 0.563 for sensitivity, 0.553 for specificity, 0.196 for negative predictive value (NPV) and 0.862 for positive predictive value (PPV). According to these four parameters the recommended PLR cut-off value was defined as 136.5. The area under the receiver operating characteristic curve was 0.568 (95% CI 0.508–0.628 $p: 0.036$) (Fig. 2).

Table 1. Patients demographic characteristics

	Group 1 (positive appendicitis)	Group 2 (negative appendicitis)	Total	p value
Age (year) (median) (min-max)	29 (17–85)	29 (17–80)	29 (17–85)	$p=0.658$
Gender (F/M) (n)	128/347	46/48	174/395	$p<0.001$
Patients (n)	475	94	569	

F; female, M; male

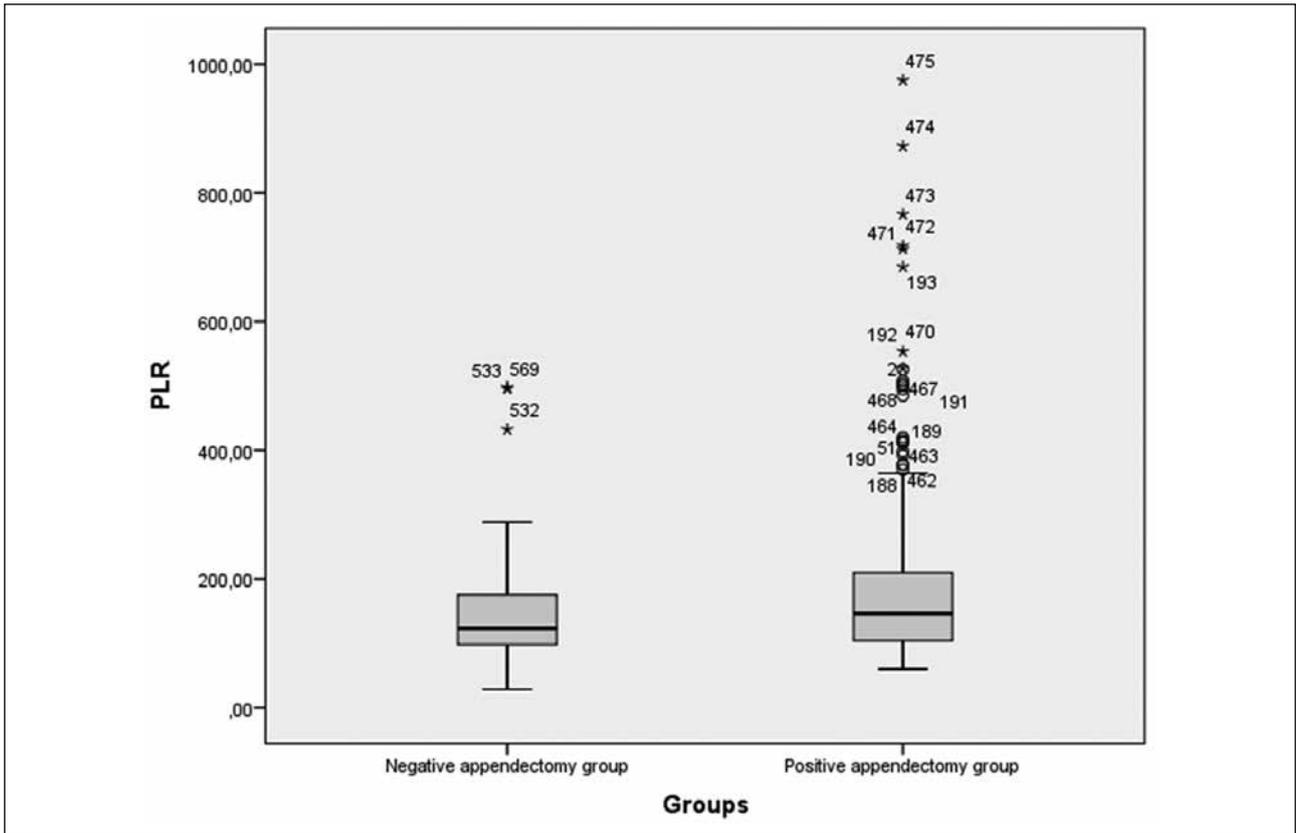


Figure 1. PLR distributions intergroups [PLR Group 1:146.5 (59.7–975.0), PLR Group 2:123.0 (28.4–497.8) and $p=0.036$].

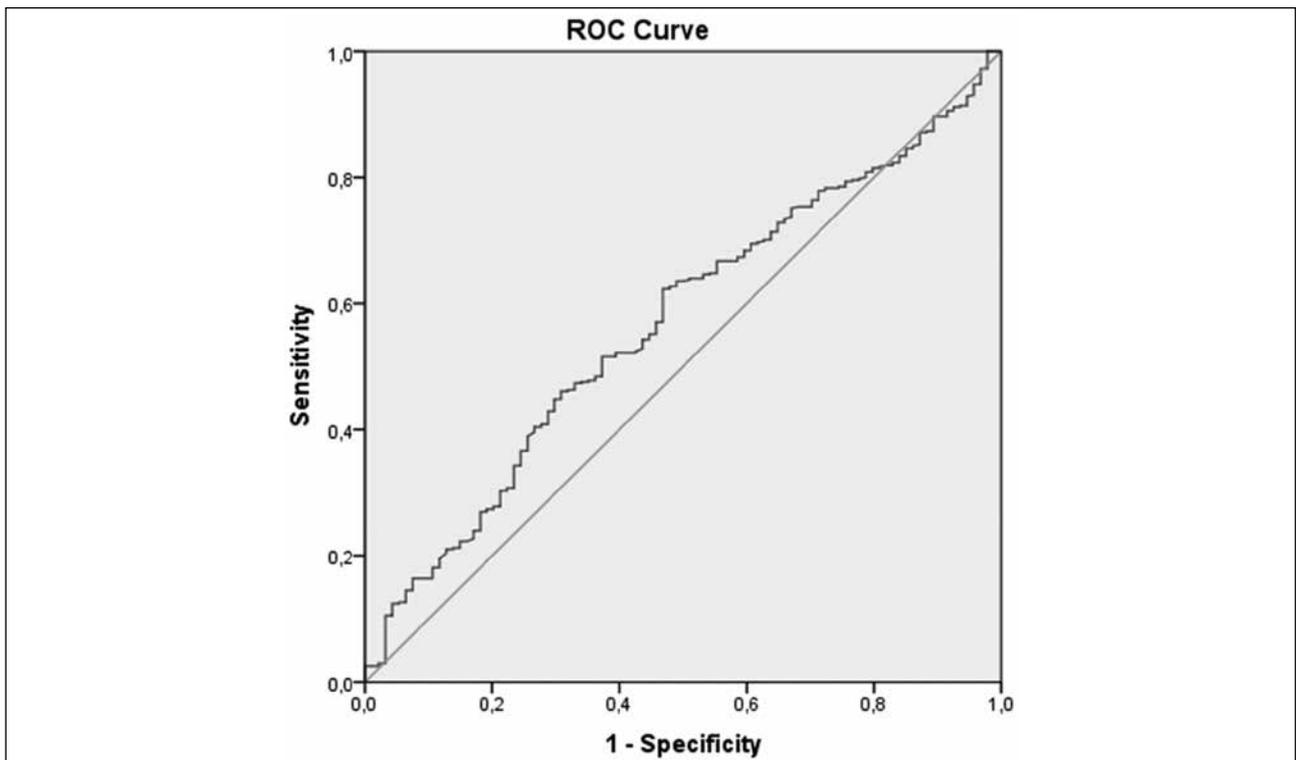


Figure 2. PLR ROC curve intergroups [Cut off value: 136.5, Area under the curve: 0.568; $p=0.036$ (CI% 95:0.508–0.628)].

Table 2. Multivariate COX regression analysis (Age, gender and PLR)

Independent factors	p values
Age	p=0.295
Gender	p<0.001
PLR	p=0.011

PLR; Platelet lymphocyte ratio

We found either of gender and PLO were independent factors to predict the complicated appendicitis when we investigated the joint role of them (Table 2).

Discussion

Appendicitis is one of the most frequent acute surgical conditions of the abdomen, and appendectomy is one of the most commonly performed operations in the world. Lee et al. reported that in their South Korean epidemiological study, the overall incidences of appendicitis, total appendectomy, and perforated appendectomy were 22.71, 13.56, and 2.91 per 10 000 populations per year, respectively⁹. Today traditional management of AA is changed from strict surgical intervention to conservative therapies such as antibiotic use². No longer are negative appendectomies acceptable because operation itself is a cause of morbidity and mortality. In recent years there have been many efforts to improve preoperative diagnosis¹⁰. We need some simple, easy accessible and cheap extra methods in addition to conventionally obtained patient history, physical examination results and laboratory tests. A lot of scoring systems derived from these simple findings were defined⁴⁻⁶. Advanced imaging studies may be more helpful but they are expensive and difficult to supply especially in rural areas.

Lately NLR is defined and used for these purposes and as a marker of serious inflammation. Positive results were reported in the literature^{7,8,11}. PLR is another marker like this and it was used in various clinical conditions to determine the degree of seriousness. In many types of cancer and inflammatory processes, the release of proinflammatory cytokines promotes magakaryocytes' proliferation. The activation of platelets is a hallmark in the natural course of cancer, by promoting neoangiogenesis, degradation of extracellular matrix, release of adhesion molecules, and growth factors^{12,13}. Lee et al. demonstrated that NLR or PLR were independent prognostic factors for overall survival with

advanced gastric cancer treated with chemotherapy¹⁴. Similarly PLR was found valuable for determination of prognosis in colorectal cancer as strong as NLR^{15,16}. But He W. et al. found no NLR was superior to PLR in patients with metastatic colorectal cancer¹⁷. There are some studies about PLR and periampullary cancer in the literature but reports are not satisfactory^{18,19}. There are valuable results in the studies about the role of PLR in terms of prognosis of the patients with breast and gynaecological malignancies^{12,13,20,21}. The predictive values of PLR in patients with myocardial infarction, peripheral arterial occlusive disease, and end-stage renal diseases were also studied²²⁻²⁴. But we didn't find any study about PLR and AA in English Literature.

In our study we found significantly higher PLR values in positive appendectomy group (G1) than negative appendectomy group (G2) (p=0.036). The cut-off value of PLR was 136.5; sensitivity, specificity, NPV and PPV were 56.3%, 55.3%, 19.6% and 86.2%, respectively. The low values may be explained by that only the operated patients were included in this study, data about other suspected cases which were not operated or medically treated were unknown. In gender analyses female/male ratio was 128/347 in G1 whereas 46/48 in G2. Thus negative appendectomy rate was statistically higher in females (p<0.001). This may be because of gynaecological diseases commonly mimicking acute appendicitis.

In this study, PLR was found a reliable parameter to distinguish AA when its value was higher than the cut-off value (136.5). There is a need for more prospective randomised studies to find ideal PLR cut-off values. It seems that quests for the ideal test should continue and the surgeon's experience will still have the biggest importance.

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