Platelet indices and neutrophil to lymphocyte ratio in adults with acute appendicitis

I D Kostakis, N Machairas, C Damaskos, C Doula, P Tsaparas, P Charalampoudis, E Spartalis, G C Sotiropoulos, G Kouraklis

Second Department of Propedeutic Surgery, Laiko General Hospital, National and Kapodistrian University of Athens, Medical School, Athens, Greece

Corresponding author: Ioannis D Kostakis (i.d.kostakis@gmail.com)

Background: A study was performed in adults with acute appendicitis and matched controls to assess the utility of the platelet indices and neutrophil to lymphocyte ratio, as a diagnostic adjunct

Methods: Data were retrospectively collected from a complete blood count test of 155 adult patients (72 men and 83 women) with histologically proven acute appendicitis upon admission, and of 50 healthy adults (20 men and 30 women). The parameters for white blood cells and platelets were compared between the two groups, and for each gender separately. **Results:** A higher white blood cell count, neutrophil count, neutrophil percentage, neutrophil to lymphocyte ratio and lower lymphocyte percentage was reported in patients with acute appendicitis than that in the healthy controls, with high areas under the curve (AUC), sensitivities, specificities, positive predictive values (PPVs) and moderate negative predictive values (NPVs). The lymphocyte count was lower in patients than it was in the healthy controls. The platletcrit was lower in the female patients than that in the female controls, whereas a difference was not detected in the male participants. Differences were not detected with regard to platelet count, mean platelet volume and platelet distribution width for both genders.

Conclusion: The neutrophil to lymphocyte ratio increases and lymphocyte percentage decreases in acute appendicitis, and can be used as an additional diagnostic marker. Plateletcrit, and therefore total platelet mass, is reduced in women with acute appendicitis, indicating the involvement of platelets in its pathophysiology. However, it is neither a reliable predictor or excluder of the disease.

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Acute appendicitis is an inflammatory disease, constituting a common cause of acute abdominal pain, 1-4 and with an estimated lifetime risk of 8%. However, diagnosis is often difficult because many other clinical entities also cause right lower quadrant pain, and acute appendicitis can present with atypical symptoms and signs. Blood tests (leukocytosis, neutrophilia and increased C-reactive protein) and imaging studies (abdominal ultrasound and/or computed tomography) are adjuncts used to establish the diagnosis. 1,3,4

It is known that the neutrophil to lymphocyte ratio is increased in many inflammatory diseases. Thus, it is a marker of inflammatory response^{5–9} It has also been found that platelets are involved in inflammatory processes.¹⁰⁻¹⁶ Their involvement in various inflammatory diseases has been emphasised in several studies.^{10,11,17,18} However, the role of the aforementioned parameters in acute appendicitis has only been investigated in a few studies. The aim of the study was to examine the probable changes of the neutrophil to lymphocyte ratio and platelet indices in adults with acute appendicitis.

Method

Patients

One hundred and fifty-five patients (72 men and 83 women, with a mean age of 31 years, a median age of 27 years, and an age range of 15-77 years) with histologically proven acute appendicitis who had undergone appendectomy in our department from August 2008 until November 2013 were included in the study. Data were retrospectively collected on white blood cell count, neutrophil count, neutrophil percentage, lymphocyte count, lymphocyte percentage, neutrophil to lymphocyte ratio, platelet count, mean platelet volume, platelet distribution width and plateleterit from the complete blood count test performed on patients upon their admission to the emergency department of the hospital. The same data were collected from a complete blood count test of 50 healthy volunteers (20 men and 30 women, with a mean age of 34 years, median age of 34 years, and an age range of 18-58 years), who were used as controls. The study conformed to the World Medical Association's Declaration of Helsinki.

Complete blood count measurement

Three millilitres of blood were collected in EDTA Vacutainer® tubes with lavender stoppers, and the specimens processed within 20 minutes of the blood having been drawn, using the Sysmex® XT-4000i automated haematology analyser (Sysmex, Kobe, Japan).

Statistical analysis

The Shapiro-Wilk test was used to assess the normality of the data distribution. The t-test and Mann-Whitney U test was used for the two group comparisons when the values were, and were not, normally distributed, respectively. The receiver operating characteristic (ROC) curve was used to assess the optimal cut-off points of the parameters for which significant differences were found. Sensitivity, specificity, positive predictive values (PPVs) and negative predictive values (NPVs) were calculated for these parameters, taking these cut-off points into account. The tests were two-tailed. The results were considered to be statistically significant if $p = \le 0.050$.

Results

A comparison of the patients and the healthy controls

There were no significant differences between patients and healthy controls concerning age (p = 0.199) or gender (p =0.425). Higher values pertaining to white blood cell count $(p = \le 0.000)$, neutrophil count $(p = \le 0.000)$, neutrophil percentage $(p = \le 0.000)$ and neutrophil to lymphocyte ratio ($p = \le 0.000$) were reported in patients with acute appendicitis, when compared to the values reported for the healthy controls. Moreover, a lower lymphocyte count $(p = \le 0.000)$, lymphocyte percentage $(p = \le 0.000)$ and plateleterit (p = 0.003) was noted in patients with acute appendicitis, when compared with that in the healthy controls, whereas no significant differences concerning platelet count (p = 0.056), mean platelet volume (p = 0.058) and platelet distribution width (p = 0.558) were found. The mean values, median values, standard deviations (SDs) and ranges for each parameter of each group are shown in Table 1.

The ROC curve analysis provided the following optimal cut-off points with which to distinguish between cases of acute appendicitis and those of healthy controls:

- White blood cell count: $\geq 9~000/\mu l$ [AUC: 0.96, 95% confidence interval (CI): 0.94–0.99, SE: 0.01, $p = \leq 0.000$, sensitivity: 91% (141/155), specificity: 92% (46/50), PPV: 97% (141/145), NPV: 77% (46/60), accuracy: 91% (187/205)].
- Neutrophil count: $\geq 5500/\mu$ l [AUC: 0.97, 95% CI: 0.95–0.99, SE: 0.01, $p = \leq 0.000$, sensitivity: 92% (142/155), specificity: 92% (46/50), PPV: 97% (142/146), NPV: 78% (46/59), accuracy: 92% (188/205)].
- Neutrophil percentage: $\geq 70\%$ [AUC: 0.94, 95% CI: 0.91–0.97, SE: 0.02, $p = \leq 0.000$, sensitivity: 87%

- (135/155), specificity: 88% (44/50), PPV: 96% (135/141), NPV: 69% (44/64), accuracy: 87% (179/205)].
- Lymphocyte count: $< 2~000/\mu l$ [AUC: 0.71, 95% CI: 0.64–0.77, SE: 0.04, $p = \le 0.000$, sensitivity: 71% (110/155), specificity: 64% (32/50), PPV: 86% (110/128), NPV: 42% (32/77), accuracy: 69% (142/205)].
- Lymphocyte percentage: $\leq 24\%$ [AUC: 0.94, 95% CI: 0.92–0.97, SE: 0.02, $p = \leq 0.000$, sensitivity: 90% (140/155), specificity: 86% (43/50), PPV: 95% (140/147), NPV: 74% (43/58), accuracy: 89% (183/205)].
- Neutrophil to lymphocyte ratio: ≥ 3 [AUC: 0.94, 95% CI: 0.92-0.97, SE: 0.02, p = ≤ 0.000, sensitivity: 90% (139/155), specificity: 88% (44/50), PPV: 96% (139/145), NPV: 73% (44/60), accuracy: 89% (183/205)].
- Plateletcrit: ≤ 0.25 [AUC: 0.64, 95% CI: 0.55–0.72, SE: 0.04, p = 0.003, sensitivity: 62% (96/155), specificity: 56% (28/50), PPV: 81% (96/118), NPV: 32% (28/87), accuracy: 61% (124/205)].

A comparison of the male patients and the healthy male controls

There was an increased white blood cell count $(p = \leq 0.000)$, neutrophil count $(p \leq 0.000)$, neutrophil percentage $(p = \leq 0.000)$ and neutrophil to lymphocyte ratio $(p = \leq 0.000)$, in male patients with acute appendicitis, when compared with that in the healthy male controls. Furthermore, there was a decreased lymphocyte count (p = 0.014) and lymphocyte percentage $(p = \leq 0.000)$ in the male patients with acute appendicitis, when compared with that in the healthy male controls, whereas no significant differences were detected regarding platelet count (p = 0.576), mean platelet volume (p = 0.306), platelet distribution width (p = 0.160) and plateletcrit (p = 0.290). The mean values, median values, SDs and ranges for each parameter of each group are shown in Table 1.

The ROC curve analysis provided the following optimal cut-off points with which to distinguish between cases of acute appendicitis and those of healthy controls:

- White blood cell count: $\geq 9~000/\mu l$ [AUC: 0.98, 95% CI: 0.96–1.00, SE: 0.01, $p = \leq 0.000$, sensitivity: 94% (68/72), specificity: 95% (19/20), PPV: 99% (68/69), NPV: 83% (19/23), accuracy: 95% (87/92)].
- Neutrophil count: $\geq 5~600/\mu$ l [AUC: 0.98, 95% CI: 0.96–1.00, SE: 0.01, $p = \leq 0.000$, sensitivity: 97% (70/72), specificity: 100% (20/20), PPV: 100% (70/70), NPV: 91% (20/22), accuracy: 98% (90/92)].
- Neutrophil percentage: $\geq 73\%$ [AUC: 0.96, 95% CI: 0.93-1.00, SE: 0.01, $p = \leq 0.000$, sensitivity: 90% (65/72), specificity: 100% (20/20), PPV: 100% (65/65), NPV: 74% (20/27), accuracy: 92% (85/92)].
- Lymphocyte count: $\geq 1~900/\mu l$ [AUC: 0.68, 95% CI: 0.56–0.80, SE: 0.06, p = 0.014, sensitivity: 69% (50/72), specificity: 60% (12/20), PPV: 86% (50/58), NPV: 35% (12/34), accuracy: 67% (62/92)].
- Lymphocyte percentage: $\leq 21\%$ [AUC: 0.97, 95% CI: 0.94–1.00, SE: 0.02, $p = \leq 0.000$, sensitivity: 92% (66/72), specificity: 90% (18/20), PPV: 97% (66/68),

Table 1: The blood test results of the healthy controls and patients with acute appendicitis						
Blood test parameter	Healthy controls			Patients with acute appendicitis		
	All controls	Men	Women	All patients	Men	Women
White blood cell count (cells/μl)						
$Mean \pm SD$	$6\ 855 \pm 1\ 438$	6.733 ± 1.089	$6\ 936 \pm 1\ 625$	$14\ 186 \pm 4\ 034$	$14\ 878 \pm 3\ 750$	$13\ 586 \pm 4\ 172$
Median (minimum to maximum)	6 955 (3 750–10 070)	6 740 (4 210–9 010)	7 020 (3 750–10 070)	14 230 (5 160–25 800)	14 870 (5 200–24 380)	13 370 (5 160–25 800)
Neutrophil count (cells/µl)						
$Mean \pm SD$	3920 ± 1167	$3\ 935 \pm 989$	$3\ 910\pm 1\ 272$	$11\ 466 \pm 3\ 974$	$12\ 045 \pm 3\ 498$	$10\ 964 \pm 4\ 282$
Median (minimum to maximum)	3 650 (1 800-7 400)	3 600 (2 100-5 600)	3 900 (1 800-7 400)	11 800 (2 700–22 300)	12 300 (2 700–22 300)	10 800 (3 200–22 200)
Neutrophil percentage (%)						
$Mean \pm SD$	56.7 ± 8.8	58.3 ± 9.7	55.6 ± 7.9	79.4 ± 9.9	80.3 ± 8	78.6 ± 11.2
Median (minimum to maximum)	55.3 (38.3-73.7)	59 (38.3–72.7)	54.9 (42.4–73.7)	81.8 (48.1–94.7)	81.6 (48.1–93.7)	81.8 (48.6–94.7)
Lymphocyte count (cells/μl)						
$Mean \pm SD$	224 ± 626	$2\ 148 \pm 718$	$2\ 304 \pm 561$	1.753 ± 815	$1~700\pm768$	1799 ± 856
Median (minimum to maximum)	2 135 (1 290–4 180)	1 995 (1 310–4 180)	2 210 (1 290–3 370)	1 640 (320–4 550)	1 655 (320-4 280)	1 600 (530-4 550)
Lymphocyte percentage (%)						
$Mean \pm SD$	33.2 ± 7.5	32.1 ± 8.8	33.9 ± 6.6	13.5 ± 8.3	11.9 ± 6	14.9 ± 9.6
Median (minimum to maximum)	34.2 (18.4–49.4)	31.7 (18.4–49.4)	34.3 (19.4–44.5)	11.5 (1.7–43.3)	10.9 (2.7–37.8)	11.9 (1.7–43.3)
Neutrophil to lymphocyte ratio						
$Mean \pm SD$	1.9 ± 0.8	2.1 ± 0.9	1.8 ± 0.7	8.4 ± 5.6	8.8 ± 5.6	8 ± 5.6
Median (minimum to maximum)	1.6 (0.8–4.0)	1.9 (0.8-4.0)	1.6 (0.9–3.8)	7.2 (1.1–34.7)	7.4 (1.3–34.7)	6.8 (1.1–24.8)
Platelet count (cells/µl)						
$Mean \pm SD$	254 480 ± 58 574	232 200 ± 42 439	269 333 ± 62 976	237 794 ± 63 103	229 430 ± 59 553	245 048 ± 65 165
Median (minimum to maximum)	240 000 (174 000– 446 000)	224 000 (174 000–318 000)	246 000 (177 000– 446 000)	227 000 (120 000– 491 000)	218 500 (120 000- 398 000)	230 000 (144 000– 491 000)
Mean platelet volume (fl)						
$Mean \pm SD$	10.5 ± 0.9	10.4 ± 1.1	10.5 ± 0.7	10.1 ± 1.2	10.1 ± 1.2	10.2 ± 1.1
Median (minimum to maximum)	10.5 (8.3–12.0)	10.7 (8.3–12.0)	10.4 (9.2-11.9)	10.1 (6.5–14)	10.1 (7.2–14)	10.1 (6.5–13.6)
Platelet distribution width (%)						
$Mean \pm SD$	13.4 ± 1.9	14 ± 1.9	13 ± 1.8	13.4 ± 2.6	13.5 ± 2.8	13.2 ± 2.4
Median (minimum to maximum)	13.2 (10.1–16.8)	14.4 (10.5–16.8)	12.7 (10.1-16.5)	12.8 (9.2–25.1)	12.9 (9.8–25.1)	12.5 (9.2–21.1)
Plateletcrit (%)						
$Mean \pm SD$	0.3 ± 0.1	0.2 ± 0.0	0.3 ± 0.1	0.2 ± 0.1	0.2 ± 0.1	0.2 ± 0.1
Median (minimum to maximum)	0.3 (0.2-0.5)	0.2 (0.2-0.3)	0.3 (0.2-0.5)	0.2 (0.1-0.5)	0.2 (0.1-0.4)	0.2 (0.1-0.5)
SD: standard deviation						

- NPV: 75% (18/24), accuracy: 91% (84/92).
- Neutrophil to lymphocyte ratio: ≥ 3.5 [AUC: 0.97, 95% CI: 0.94–1.00, SE: 0.02, $p = \leq 0.000$, sensitivity: 90% (65/72), specificity: 90% (18/20), PPV: 97% (65/67), NPV: 72% (18/25), accuracy: 90% (83/92).
- Plateletcrit: ≤ 0.23 [AUC: 0.58, 95% CI: 0.44–0.71, SE: 0.07, p = 0.291, sensitivity: 53% (38/72), specificity: 60% (12/20), PPV: 83% (38/46), NPV: 26% (12/46), accuracy: 54% (50/92)].

A comparison of the female patients and the healthy female controls

There was an increased white blood cell count $(p=\le 0.000)$, neutrophil count $(p=\le 0.000)$, neutrophil percentage $(p=\le 0.000)$ and neutrophil to lymphocyte ratio $(p=\le 0.000)$ in the female patients with acute appendicitis, when compared with that in the healthy female controls. Moreover, there was a decreased lymphocyte count (p=0.000), lymphocyte percentage (p=< 0.000) and plateletcrit (p=0.004) in the female patients with acute appendicitis, when compared with that in the healthy female controls, whereas no significant differences concerning platelet count (p=0.058), mean platelet volume (p=0.054) and platelet distribution width (p=0.938) were detected. The mean values, median values, SDs and ranges for each parameter of each group are shown in Table 1.

The ROC curve analysis provided the following optimal cut-off points with which to distinguish between cases of acute appendicitis and those of healthy controls:

- White blood cell count: $\geq 9\ 100/\mu l$ [AUC: 0.94, 95% CI: 0.90–0.98, SE: 0.02, $p = \leq 0.000$, sensitivity: 87% (72/83), specificity: 93% (28/30), PPV: 97% (72/74), NPV: 72% (28/39), accuracy: 89% (100/113)].
- Neutrophil count: $\geq 5~200/\mu l$ [AUC: 0.95, 95% CI: 0.92–0.99, SE: 0.02, $p = \leq 0.000$, sensitivity: 88% (73/83), specificity: 90% (27/30), PPV: 96% (73/76), NPV: 73% (27/37), accuracy: 89% (100/113)].
- Neutrophil percentage: $\geq 67\%$ [AUC: 0.93, 95% CI: 0.88–0.98, SE: 0.02, p = < 0.000, sensitivity: 86% (71/83), specificity: 93% (28/30), PPV: 97% (71/73), NPV: 70% (28/40), accuracy: 88% (99/113)].
- Lymphocyte count: $\leq 2~000/\mu l$ [AUC: 0.72, 95% CI: 0.63–0.82, SE: 0.05, p=0.000, sensitivity: 66% (55/83), specificity: 73% (22/30), PPV: 87% (55/63), NPV: 44% (22/50), accuracy: 68% (77/113)].
- Lymphocyte percentage: ≤ 26% [AUC: 0.92, 95% CI: 0.88-0.98, SE: 0.02, p = < 0.000, sensitivity: 88% (73/83), specificity: 87% (26/30), PPV: 95% (73/77), NPV: 72% (26/36), accuracy: 88% (99/113)].
- Neutrophil to lymphocyte ratio: ≥ 2.5 [AUC: 0.93, 95% CI: 0.88–0.98, SE: 0.02, p = < 0.000, sensitivity: 88% (73/83), specificity: 87% (26/30), PPV: 95% (73/77), NPV: 72% (26/36), accuracy: 88% (99/113)].
- Plateletcrit: ≤ 0.27 [AUC: 0.68, 95% CI: 0.57–0.78, SE: 0.06, p = 0.004, sensitivity: 69% (57/83), specificity: 53% (16/30), PPV: 80% (57/71), NPV: 38% (16/42), accuracy: 65% (73/113)].

Discussion

There is increasing evidence in the literature that platelets are involved in inflammatory and immunological processes. 10-16 In particular, platelets interact with endothelial cells, and all categories of leukocytes, through numerous mechanisms, such as adhesive interactions via P-selectin, which result in leukocyte activation and recruitment, the production of inflammatory cytokines, such as interleukin-1\beta and chemokines, and the expression of various immunological modulators, such as toll-like receptors and the CD40 ligand. 10-16 Platelets also interact with the complement system¹² and microorganisms.^{10,13} Furthermore, platelets have been implicated in the pathophysiological processes of various disorders in which inflammation plays a major role, such as atherosclerosis, sepsis, infectious disease, autoimmune disease (rheumatoid arthritis, systemic lupus erythematosus and systemic sclerosis), neurological disorders (multiple sclerosis) and transfusion-related acute lung injury. 10,11,17,18

Changes in platelet indices have been detected in various disorders in which inflammatory processes are implicated. First of all, reactive thrombocytosis is found in chronic inflammatory and infectious diseases, such as connective tissue disorders, inflammatory bowel disease, temporal arteritis, chronic pneumonitis and tuberculosis. 19 Mean platelet volume is a marker of platelet activation, because larger platelets have more and larger pseudopodia, making them more reactive than smaller ones, resulting in an enhanced production rate.²⁰⁻²³ It increases in metabolic syndrome, myocardial infarction, peripheral artery disease, ischaemic stroke, obstructive sleep apnoea, rheumatoid arthritis, ankylosing spondylitis, psoriasis, endocarditis, celiac disease and pre-eclampsia. On the other hand, it decreases in inflammatory bowel disease, familial Mediterranean fever and chronic obstructive pulmonary disease.^{23–26} Platelet distribution width is the width of the size distribution curve of the platelets in flat the 20% level when assuming that the peak distribution curve is 80-100%. 23,24 It is higher during platelet activation and represents the variability of platelet size, probably owing to the higher production of larger platelets. 20,22,23 It is increased in many diseases, such as myocardial infarction,²² obstructive sleep apnoea,²⁵ pulmonary tuberculosis²⁷ and pre-eclampsia,²³ whereas it is reduced in various disorders, such as inflammatory bowel disease.26 Finally, plateletcrit reflects the total platelet mass because it indicates the volume percentage of platelets in the whole blood [pateletcrit (%) = (mean platelet volume (fl) x platelet count (cells/µl)/107].^{22,26} It is higher in various diseases, such as pulmonary tuberculosis,²⁷ autoimmune gastritis²¹ and pre-eclampsia, 23 whereas it is lower in some disorders, such as inflammatory bowel disease.²⁶

There are only a few studies on platelet indices in acute appendicitis. Albayrak et al. found that mean platelet volume decreased in adult patients with acute appendicitis, in comparison with that in healthy controls, ²⁸ whereas Yang et al. detected this reduction only in adult male patients with acute appendicitis. ²⁹ By contrast, Narci et al. reported that

mean platelet volume increased in adult patients with acute appendicitis, when compared with that in healthy controls.³⁰ Furthermore, Bilici et al. found that children with acute appendicitis had a lower platelet count and mean platelet volume than healthy children,³¹ whereas no significant difference regarding mean platelet volume was detected by Uyanik et al. between children with acute appendicitis and healthy children.³²

Systemic inflammatory response also causes neutrophilia and lymphocytopenia, thus resulting in an increase in the neutrophil to lymphocyte ratio, a marker of inflammation in many diseases.^{5–9,33} Critically ill patients with severe sepsis have a higher neutrophil to lymphocyte ratio, which relates to a worse prognosis.⁹ An increased neutrophil to lymphocyte ratio has also been reported in children with cystic fibrosis and with poor clinical status.⁷ Moreover, high values of this ratio have been found in many malignant diseases, such as breast, lung, gastric, colorectal and pancreatic cancer, and have been associated with a poor prognosis and more aggressive and/or more advanced disease.^{6,8} Higher values of this ratio relate to increased morbidity and mortality, and worse outcomes in patients with acute and chronic coronary artery syndromes.^{5,33}

Only a few studies have been conducted on the neutrophil to lymphocyte ratio in patients with acute appendicitis. It has been reported in some studies that there is an increased neutrophil to lymphocyte ratio in cases with histologically confirmed acute appendicitis, when compared with that in cases without it,34-37 and that this ratio is even higher in complicated acute appendicitis, than in uncomplicated appendicitis. 37-39 Guraya et al. also found that one in two patients with acute appendicitis had lymphopenia. 40 Fergusson et al. and Nordback and Harju reported that the lymphocyte count and percentage were lower in cases with histologically confirmed acute appendicitis, than that in cases with a normal appendix and clinical image mimicking acute appendicitis.41,42 Furthermore, Goulart et al. and Jahangiri and Wyllie reported a decreased lymphocyte count in patients with complicated acute appendicitis, compared to that in patients with uncomplicated appendicitis. 43,44

Interesting findings were reported in our study. Plateletcrit was the only platelet index found to be altered in acute appendicitis. Significant differences were not detected between the patients and healthy controls concerning the platelet count, mean platelet volume and platelet distribution width. In particular, a lower plateleterit was reported in adult patients with acute appendicitis, when compared with that in healthy adults. However, this difference only applied to female patients and not to male ones, when gender was taken into account. This finding leads to the conclusion that plateletcrit, and therefore total platelet mass, is reduced only in female adult patients with acute appendicitis, and not in male ones. However, plateletcrit was associated with a moderate AUC and intermediate to insufficient sensitivity, specificity and NPV. Only PPV was satisfactory, with values just above 80%. On the other hand, the neutrophil to lymphocyte ratio (reported to be higher in cases of acute appendicitis), lymphocyte percentage (reported to be lower in acute appendicitis),

and white blood cell count, neutrophil count and neutrophil percentage (widely used markers of inflammation and used in the diagnosis of acute appendicitis),^{1,3,4} were associated with a very good AUC, and high sensitivities, specificities and PPV. Only the NPV was moderate. However, the lymphocyte count, lower in patients with acute appendicitis, was associated with a moderate AUC and intermediate to insufficient sensitivity, specificity and NPV. Only the PPV was satisfactory, with values above 80%.

Conclusion

The neutrophil to lymphocyte ratio increases, and the lymphocyte percentage decreases, in acute appendicitis, and can be used as additional diagnostic markers which provide results equivalent to those of the usual inflammatory markers used. By contrast, plateletcrit, and therefore total platelet mass, is reduced in female adult patients with acute appendicitis, which while indicating the involvement of platelets in its pathophysiology, is not a sufficient diagnostic index of this disease.

Conflict of interest

The authors declare that there was no conflict of interest when writing this article.

REFERENCES

- 1. Hennelly KE, Bachur R. Appendicitis update. Curr Opin Pediatr. 2011;23(3):281–285.
- 2. Virmani V, Papadatos D, Fasih N, et al. Rule out appendicitis: "to be or not to be". Can Assoc Radiol J. 2012;63(1):47–60.
- 3. Petroianu A. Diagnosis of acute appendicitis. Int J Surg. 2012;10(3):115-119.
- Wray CJ, Kao LS, Millas SG, et al. Acute appendicitis: controversies in diagnosis and management. Curr Probl Surg. 2013;50(2):54–86.
- Bhat T, Teli S, Rijal J, et al. Neutrophil to lymphocyte ratio and cardiovascular diseases: a review. Expert Rev Cardiovasc Ther. 2013;11(1):55-59.
- Guthrie GJ, Charles KA, Roxburgh CS, et al. The systemic inflammation-based neutrophil-lymphocyte ratio: experience in patients in cancer. Crit Rev Oncol Hematol. 2013;88(1):218–230.
- O'Brien CE, Price ET. The blood neutrophil to lymphocyte ratio correlates with clinical status in children with cystic fibrosis: a retrospective study. PLoS One. 2013;8(10):e77420.
- Li MX, Liu XM, Zhang XF, et al. Prognostic role of neutrophilto-lymphocyte ratio in colorectal cancer: a systematic review and meta-analysis. Int J Cancer. 2014;134(10):2403–2413.
- Zahorec R. Ratio of neutrophil to lymphocyte counts rapid and simple parameter of systemic inflammation and stress in critically ill. Bratisl Lek Listy. 2001;102(1):5–14.
- Semple JW, Freedman J. Platelets and innate immunity. Cell Mol Life Sci. 2010;67(4):499–511.
- 11. Smith TL, Weyrich AS. Platelets as central mediators of systemic inflammatory responses. Thromb Res. 2011;127(5):391–394.
- Hamad OA, Bäck J, Nilsson PH, et al. Platelets, complement, and contact activation: partners in inflammation and thrombosis. Adv Exp Med Biol. 2012;946:185–205.

- 13. Vieira-de-Abreu A, Campbell RA, Weyrich AS, Zimmerman GA. Platelets: versatile effector cells in hemostasis, inflammation, and the immune continuum. Semin Immunopathol. 2012;34(1):5–30.
- Huang HS, Chang HH. Platelets in inflammation and immune modulations: functions beyond hemostasis. Arch Immunol Ther Exp (Warz). 2012;60(6):443–451.
- Amison R, Page C, Pitchford S. Pharmacological modulation of the inflammatory actions of platelets. Handb Exp Pharmacol. 2012;(210):447–468.
- Ghasemzadeh M, Hosseini E. Platelet-leukocyte crosstalk: linking proinflammatory responses to procoagulant state. Thromb Res. 2013;131(3):191–197.
- Ware J, Corken A, Khetpal R. Platelet function beyond hemostasis and thrombosis. Curr Opin Hematol. 2013;20(5):451–456.
- Habets KL, Huizinga TW, Toes RE. Platelets and autoimmunity. Eur J Clin Invest. 2013;43(7):746–757.
- Schafer AI. Trombocytosis and thrombocythemia. Blood Rev. 2001;15(4):159–166.
- 20. Vagdati E, Gounari E, Lazaridou E, et al. Platelet distribution width: a simple, practical and specific marker of activation of coagulation. Hippokratia. 2010;14(1):28–32.
- Tüzün A, Keskin O, Yakut M, et al. The predictive value of mean platelet volume, plateletcrit and red cell distribution width in the differentiation of autoimmune gastritis patients with and without type I gastric carcinoid tumors. Platelets. 2014;25(5):363-366.
- Cetin M, Bakirci EM, Baysal E, et al. Increased platelet distribution width is associated with ST-segment elevation myocardial infarction and thrombolysis failure. Angiology. 2014;65(8):737-743.
- Yang SW, Cho SH, Kwon HS, et al. Significance of the platelet distribution width as a severity marker for the development of preeclampsia. Eur J Obstet Gynecol Reprod Biol. 2014;175:107-111
- 24. Leader A, Pereg D, Lishner M. Are platelet volume indices of clinical use? A multidisciplinary review. Ann Med. 2012;44(8):805–816.
- 25. Nena E, Papanas N, Steiropoulos P, et al. Mean platelet volume and platelet distribution width in non-diabetic subjects with obstructive sleep apnoea syndrome: new indices of severity? Platelets. 2012;23(6):447–454.
- 26. Öztürk ZA, Dag MS, Kuyumcu ME, et al. Could platelet indices be new biomarkers for inflammatory bowel diseases? Eur Rev Med Pharmacol Sci. 2013;17(3):334–341.
- Sahin F, Yazar E, Yildiz P. Prominent features of platelet count, plateletcrit, mean platelet volume and platelet distribution width in pulmonary tuberculosis. Multidiscip Respir Med. 2012;7(1):38.
- 28. Albayrak Y, Albayrak A, Albayrak F, et al. Mean platelet volume: a new predictor in confirming acute appendicitis diagnosis. Clin Appl Thromb Hemost. 2011;17(4):362–366.

- 29. Yang JJ, Cho SY, Ahn HJ, et al. Mean platelet volume in acute appendicitis: a gender difference. Platelets. 2014;25(3):226–227.
- 30. Narci H, Turk E, Karagulle E, et al. The role of mean platelet volume in the diagnosis of acute appendicitis: a retrospective case-controlled study. Iran Red Crescent Med J. 2013;15(12):e11934.
- 31. Bilici S, Sekmenli T, Göksu M, et al. Mean platelet volume in diagnosis of acute appendicitis in children. Afr Health Sci. 2011;11(3):427–432.
- 32. Uyanik B, Kavalci C, Arslan ED, et al. Role of mean platelet volume in diagnosis of childhood acute appendicitis. Emerg Med Int. 2012;2012:823095.
- 33. Wang X, Zhang G, Jiang X, et al. Neutrophil to lymphocyte ratio in relation to risk of all-cause mortality and cardiovascular events among patients undergoing angiography or cardiac revascularization: a meta-analysis of observational studies. Atherosclerosis. 2014;234(1):206–213.
- 34. Goodman DA, Goodman CB, Monk JS. Use of the neutrophil:lymphocyte ratio in the diagnosis of appendicitis. Am Surg. 1995;61(3):257–259.
- 35. Yazici M, Ozkisacik S, Oztan MO, Gürsoy H. Neutrophil/ lymphocyte ratio in the diagnosis of childhood appendicitis. Turk J Pediatr. 2010;52(4):400–403.
- 36. Markar SR, Karthikesalingam A, Falzon A, Kan Y. The diagnostic value of neutrophil:lymphocyte ratio in adults with suspected acute appendicitis. Acta Chir Belg. 2010;110(5):543-547.
- 37. Kahramanca S, Ozgehan G, Seker D, et al. Neutrophil-to-lymphocyte ratio as a predictor of acute appendicitis. Ulus Travma Acil Cerrahi Derg. 2014;20(1):19–22.
- 38. Białas M, Taran K, Gryszkiewicz M, Modzelewski B. Evaluation of neutrophillymphocyte ratio usefulness in the diagnosis of appendicitis. Wiad Lek. 2006;59(9-10):601–606.
- Ishizuka M, Shimizu T, Kubota K. Neutrophil-tolymphocyte ratio has a close association with gangrenous appendicitis in patients undergoing appendectomy. Int Surg. 2012;97(4):299–304.
- 40. Guraya SY, Al-Tuwaijri TA, Khairy GA, Murshid KR. Validity of leukocyte count to predict the severity of acute appendicitis. Saudi Med J. 2005;26(12):1945–1947.
- 41. Fergusson JA, Hitos K, Simpson E. Utility of white cell count and ultrasound in the diagnosis of acute appendicitis. ANZ J Surg. 2002;72(11):781–785.
- 42. Nordback I, Harju E. Inflammation parameters in the diagnosis of acute appendicitis. Acta Chir Scand. 1988;154(1):43-48.
- 43. Goulart RN, Silvério Gde S, Moreira MB, Franzon O. Main findings in laboratory tests diagnosis of acute appendicitis: a prospective evaluation. Arq Bras Cir Dig. 2012;25(2):88–90.
- 44. Jahangiri M, Wyllie JH. Peripheral blood lymphopenia in gangrenous appendicitis. BMJ. 1990;301(6745):215.