Mortality in Acute Arterial Mesenteric Ischemia: A Single-Center Experience

Akut Arteryel Mezenterik İskemide Mortalite: Tek Merkez Deneyimi

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Öz	
Objective	Acute mesenteric ischemia (AMI) is a life-threatening pathology that often requires urgent surgical intervention. It is frequently faced in elderly patients. Despite advances in the diagnosis of AMI morbidity, and mortality rates are still high. The presence of comorbidities delayed surgical intervention due to diagnostic difficulties, and in most cases, older patients with heart problems may be remarkable factors for higher mortality rates. Diagnostic challenges are tried to be overcome with both imaging methods and laboratory findings. Unfortunately, to date, no clear laboratory parameters have been described in diagnosing or predicting mortality of mesenteric ischemia. In our study, we aimed to evaluate the relationship between biochemical markers and mortality in AMI.
Materials and Methods	The patients diagnosed with acute arterial mesenteric ischemia between the dates 2010-2019 were enrolled in the study. Demographic characteristics, diagnostic biochemical parameters (lymphocyte, neutrophil, albumin, C-reactive protein, lactate, lactate dehydrogenase, Di-Dimer, troponin, white blood count, platelet, fibrinogen, creatine kinase-MB) levels, as well as comorbidities of the patients, were evaluated.
Results	A total of 149 patients was enrolled in the study, with a mean age of 66.11. In the non-survivor group, neutrophil, RDW, lactate, DDM, Fibrinogen, and CK-MB levels, furthermore albumin levels were defined as significantly elevated in the survivor group (p<0.05). Hypertension was the major comorbidity by far in both groups.
Conclusion	High levels of fibrinogen, lactate, DDM, CK-MB levels and low levels of albumin levels may be predictors of mortality in mesenteric ischemia.
Keywords	Mesenteric ischemia; Mortality; Predictive factors
Abstract	
Amaç	Akut mezenterik iskemi (AMI), sıklıkla acil cerrahi müdahale gerektiren hayatı tehdit eden bir patolojidir. Yaşlı hastalarda sıklıkla karşılaşılmaktadır. AMI tanısındaki gelişmelere rağmen morbidite ve mortalite oranları hala yüksektir. Komorbiditelerin varlığı, tanısal zorluklar nedeniyle cerrahi müdahaleyi geciktirir ve genellikle kardiyak problemleri olan yaşlı hastalar, daha mortal seyredebilirler. Gerek görüntüleme yöntemleri gerekse laboratuvar bulguları ile tanısal zorluklar aşılmaya çalışılmaktadır. Günümüzde halen mezenterik iskeminin mortalitesini teşhis etmek veya tahmin etmek için net bir laboratuvar parametresi tanımlanmamıştır.Çalışmamızda AMI'de biyokimyasal belirteçlerin mortalite ile ilişkisini değerlendirmeyi amaçladık.
Gereç ve Yöntemle	2010-2019 tarihleri arasında akut arteriyel mezenterik iskemi tanısı alan hastalar çalışmaya dahil edildi. Demografik özellikler, tanısal biyokimyasal parametreler (lenfosit, nötrofil, albümin, C-reaktif protein, laktat, laktat dehidrojenaz, D-Dimer, troponin, beyaz kan sayımı, trombosit, fibrinojen, kreatin kinaz-MB) seviyeleri ve ayrıca komorbiditeler değerlendirildi.
Bulgular	Çalışmaya yaş ortalaması 66.11 olan toplam 149 hasta alındı. Mortal seyirli grupta nötrofil, RDW, laktat, DDM, Fibrinojen ve CK-MB seviyeleri, ayrıca albümin seviyeleri hayatta kalan grupta anlamlı olarak yüksek olarak tanımlandı (p < 0.05). Her iki grupta da açık ara en önemli komorbidite hipertansiyondu.
Sonuç	Yüksek fibrinojen, laktat, DDM, CK-MB seviyeleri ve düşük albümin seviyeleri mezenterik iskemide mortaliteyi saptamada prediktif faktörler olabilirler.
Anahtar Kelimeler	Mezenterik iskemi; Mortalite; Prediktif faktörler

INTRODUCTION

Acute mesenteric ischemia (AMI) is a life-threatening disease that progresses with the occlusion of the mesenteric vessels that feed the intestines, which is often misdiagnosed and requires urgent surgical intervention.¹ The incidence of AMI increases with age, and while the mean age of onset was 67 years, there was no difference in incidence between men and women.^{2,3} The most common causes of impairment of blood flow to the intestine are mesenteric vascular insufficiency due to underlying causes such as atherosclerosis, mesenteric artery embolism, vasospasm, and mesenteric vein thrombosis.4,5 The duration of ischemia, degree of mesenteric artery occlusion, and rate of collateral flow are factors that determine bowel injury after acute arterial occlusion.⁶ Four main etiological factors can be mentioned in AMI, and 75% of these are arterial occlusions (50% arterial embolism, 25% arterial thrombosis).¹⁻³ The most leading reason for the delay in diagnosis of AMI is the rise of acute abdominal examination findings in the last period of the disease. Despite advanced diagnostic methods, the critical diagnostic step is to suspect that the patient may have AMI. Although the American Society of Gastroenterology practice guidelines set angiography as the gold standard in mesenteric ischemia, availability to this intervention may not be appropriate in every hospital.⁷ The most vital advantage of catheter angiography is that it can favorably affect mortality and morbidity, as well as the surgical requirement of the patient, by recanalization of the occluded artery in patients to whom can be applied and detected at an early stage of the disease. Nevertheless, in recent years, computed tomography angiography (CTA) has been replaced the role of catheter angiography, especially in terms of faster application and as a less invasive method in the diagnose of AMI.^{1,8}

Surgical intervention is the crucial part of the treatment of AMI. The results of surgical intervention, which can be performed without the signs of peritonitis, can be promising. In the surgical approach, while trying to ensure the continuity of the gastrointestinal system by resection of the ischemic bowel and observing the continuity of the blood flow, this patient group should be followed up in intensive care in the post-operative period.⁹

In addition to the radiological diagnosis of AMI, the contribution of laboratory data in this process may be valuable in determining the prognosis and severity of the patient. Nevertheless, to date, the specificity of any laboratory parameter or group cannot be mentioned in the prognosis or diagnosis of AMI.^{3,10} Biomarkers such as L-lactate, D-dimer, leukocyte, C reactive protein (CRP), and NLR levels are the frequently utilized laboratory parameters that can be taken into account.

This retrospective study aims to evaluate the effectiveness of preoperative biochemical parameters in predicting mortality and survival in patients diagnosed with AMI.

MATERIAL and METHODS

Our study was carried out in accordance with the Declaration of Helsinki with the approval of Sakarya University Faculty of Medicine Non-Invasive Clinical Research Ethics Committee dated 29.01.2021 and numbered E-71522473-050.01.04.32179-296. The files of patients who were diagnosed with AMI (arterial occlusion) between January 2010 and June 2019 and underwent surgical intervention were retrospectively analyzed. The diagnosis of AMI was based on clinical laboratory and CTA findings. All patients having the signs of peritonitis underwent surgery. Patients with venous ischemia defined in imaging methods and cases with complete root occlusion of the superior mesenteric artery were excluded from the study. Demographic characteristics, diagnostic biochemical parameters (lymphocyte, neutrophil, albumin, CRP, lactate, lactate dehydrogenase (LDH), Di-Dimer, troponin, white blood count (WBC), platelet, fibrinogen, creatine kinase-MB) levels, as well as comorbidities of the patients, were evaluated.

Statistical Analysis

SPSS (Statistics Package for Social Sciences, SPSS Inc., Chi-

cago, IL, USA) Windows 11 program was used to statistically analyze the results of the study. Kolmogorov-Smirnov test was used for the compatibility of the normal distribution of the data. P <0.05 values are considered significant for all variables.

RESULTS

A total of 149 patients was enrolled in the study, with a mean age of $66.11 \pm$ in the survivor group (SG) and $66.79 \pm$ in the non-survivor group (NSG). One hundred sixteen patients were survived (77.9%). There was no statistically significant difference in terms of gender between the groups. (Table 1).

Table 1. Frequency distribution of the gender and age of the patients according to their life status.						
		vivor ; 77,9%)	Non-survivor (n=33; 22,1%)			
	f	%	f	%		
Male	57	57,8	16	48,5		
Female	49	42,2	17	51,5		
Sum	116	100	33	100		
Mean age	66	,11	66,79			
Highest age	97		89			
Lowest age	1	9	26			

When patients' data on comorbid diseases are evaluated, it is figured out that hypertension (HT) ranks first in both the SG (31.0% and the NSG (21.2%) in terms of the most common comorbidities. Although the SG generally has more comorbidities, diabetes mellitus (DM) is the second most common disease with 7.8% after HT. On the other hand, when the data of the SG are examined, DM comes second with 9.1%, and COPD comes third with 6.1% respectively (Table 2).

Table 2. Frequency distribution according to their life status.	of comor	bid disea	ses of pa	tients
	Survivor (n=116)		Non-survi- vor(n=33)	
	f	%	f	%
Alzheimer	5	4,3	-	-
Arhythmia	1	,9	-	-
Atrial fibrillation	3	2,6	1	2,6
Benign prostatic hypertrophy	1	,9	1	3,0
Buerger	1	,9	-	-
Crohn	1	,9	-	-
Diabetes mellitus	9	7,8	3	9,1
Deep vein thrombosis	1	,9	-	-
Epilepsy	1	,9	-	-
Hepatitis B	-	-	-	-
Hepatitis C	-	-	-	-
Hypothyroidism	1	,9	-	-
Hypertension	36	31,0	7	21,2
Coronary artery disease	4	3,4	1	3,0
Chronic renal failure	3	2,6	2	2,6
Chronic obstructive pulmo- nary disease	7	6,0	2	6,1
Colon cancer	2	1,7	-	-
Heart failure	4	3,4	-	-
Parkinson's disease	2	1,7	-	-
Kidney failure	1	,9	-	-
Cerebrovascular disease	2	1,7	1	3,0

In the non-survivor group, neutrophil, RDW, lactate, DDM, Fibrinogen, and CK-MB levels, furthermore albumin levels were defined as significantly elevated in the survivor group (p<0.05) (Table 3).

The age and gender of the patients were compared according to the survival status, and it ascertained that there was no statistically significant difference (Table 4). Considering the surviving patients in the context of the age category, it was noticed that 39.7% of the survivor patients were under 65 years, and 60.3% were 65 years aged and over. In addition, it has been noted that 39.4% of the non-survivor patients are below 65 years, and 60.6% are 65 years aged and higher. However, there was no statistically significant difference between groups (Table 5).

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	son results of Lymphocy ts According to Their Life				, i Li, i loi inogen an	
	Status	n	Mean value Rank Average	Rank Sum	U	р
Tanan ali a anta	Survivor	116	76,89	8919,50	1604 50	,320
Lymphocyte	Non-Survivor	33	68,35	2255,50	- 1694,50	
NT (11	Survivor	116	70,83	8216,00	1420.00	,030
Neutrophil	Non-Survivor	33	89,67	2959,00	- 1430,00	
D I	Survivor	116	72,27	8383,00	1505.00	,150
Rdw	Non-Survivor	33	84,61	2792,00	1597,00	
Albumin	Survivor	116	80,34	9319,00	1205.00	,010
Albumin	Non-Survivor	33	56,24	1856,00	1295,00	
C	Survivor	116	71,81	8330,50	1544,50	,090
Crp	Non-Survivor	33	86,20	2844,50		
T	Survivor	116	65,88	7642,00	856,00	,000
Lactate	Non-Survivor	33	107,06	3533,00		
D-dm	Survivor	68	38,47	2616,00	270,00	,000
D-dm	Non-Survivor	32	76,06	2434,00		
T	Survivor	69	42,20	2912,00	497,00	,000
Troponin	Non-Survivor	31	68,97	2138,00		
1471	Survivor	116	76,89	8919,50	1694,50	,320
Wbc	Non-Survivor	33	68,35	2255,50		
Fihainagan	Survivor	72	76,22	8841,50	485,00	,000
Fibrinogen	Non-Survivor	33	70,71	2333,50		
Ch Mh	Survivor	74	43,24	3113,00	014.00	,010
Ck_Mb	Non-Survivor	32	74,30	2452,00	814,00	

Table 4. Statistical results of the gender of the patients according to their life status							
			Gender		¥2		
			Male	Female	X^2	р	
Life Status	Survivor	n	67 (57,8%)	49 (42,2%)	,89	,340	
	Non-Survivor	n	16 (48,5%)	17 (51,5%)			

Tablo 5. Statistical results of the patients' ages according to their life status							
			Age		X^2		
		Age under 65	Age over 65	Λ-	Р		
I for Chattan	Survivor	n	46 (%39,7%)	70 (60,3%)	,00	,970	
Life Status	Non-Survivor	n	13 (39,4%)	20 (60,6%)			

DISCUSSION

AMI was first described in detail in the 15th century.11 Risk factors affecting mortality in acute mesenteric ischemia have not been studied much, and published studies are incompatible with each other.^{12,13} AMI is a low volume emergency, with an incidence of 0.63 per 100,000 in Europe and 1 in 1000 hospitalized patients in Europe and the United States.^{14,15} Even there is no gender difference in AMI a handful of studies indicate that 70% proportion of the patients are female.^{2,15} The majority of cases in AMI arise in patients of advanced age and especially in patients with additional cardiovascular diseases. Hypertension and diabetes mellitus, which predispose cardiovascular comorbidities such as atrial fibrillation, ischemic heart disease, and coronary artery disease, can be mentioned as the leading reasons for AMI.^{2,14,16} When we examine our patient groups, we detected that the average age of AMI experience was close to the literature average (Table 1). When the previous studies are evaluated, particularly cardiovascular diseases come to the fore however, hypertension and diabetes mellitus, which are the main reasons for the emergence of cardiovascular disorders, were ascertained to be significant comorbidity for AMI in both the survivor and the non-survivor group (Table 2). Interestingly, while comorbidities were detected in several diverse groups in the survivor group, the non-survivor group consisted of a limited group of diseases. Thereby, beyond the variety and multiplicity of comorbidity in AMI, the presence of cardiovascular pathologies can be mentioned as chief etiologic factors on mortality (Table 2).

Although laboratory results are inconclusive, the existence of nonidentical values may assist clinical suspicion. A specific biomarker for the early diagnosis of AMI has not yet been defined.^{14,17} Acidosis, amylase, urea, creatinine, and increased leukocyte count have been associated with mortality in various studies.^{12,15,16,18–20} In addition, the levels of D-dimer, which is the enzymatic product of intravascular coagulation, and lactate can be found to be high even if not specific for AMI.²¹ In a large meta-analysis in which

biomarkers have been evaluated to date has been emphasized that amylase, leukocyte count, LDH are not fitting tests for the diagnosis of AMI due to the low predictive value of intestinal ischemia.^{1,3,22} In this current study, a key role may be referred to the biomarkers we checked due to selecting the same type of AMI group (arterial occlusion). Thus, the determined parameters can be more decisive. In our study, we found that the percentage of neutrophils, RDW, Lactate, D-dimer, troponin, fibrinogen, and CK-MB levels were significantly higher in the non-survivor group, and similar results are available in the literature (Table 3, Figure 1,2,3). Albumin levels were statistically elevated in the survivor group (Figure 1). Although not surprising, average values of albumin levels contribute significantly to the recovery of many surgical diseases but have never been described for AMI.

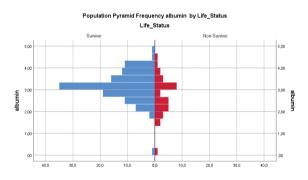


Figure 1 : Population pyramid of frequency between life status and albumin

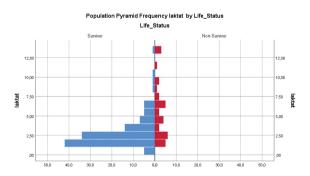


Figure 2 : Population pyramid frequency between life status and lactate

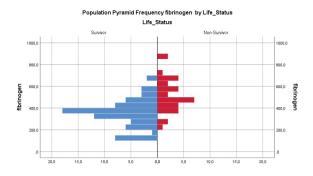


Figure 3 : Population pyramid frequency between life status and fibrinogen

The weakness of our study is that although it is not possible to design a prospective study on this subject, it is a retrospective review as well as a single-center experience. The fact that the number of patients included in the research is relatively satisfactory, so the results of the study are explicit and may contribute to the literature.

AMI is a disease frequently discerned in the elderly, and despite advances in diagnosis and treatment, morbidity moreover mortality rates are still high. In our series of 149 patients, we observed that high levels of lactate, fibrinogen, DDM, CK-MB and low albumin levels at the time of admission had an increasing effect on mortality. Based on these findings, we speculate that lactate, fibrinogen, and albumin levels at presentation may be prognostic factors in acute mesenteric ischemia.

Our study was carried out in accordance with the Declaration of Helsinki with the approval of Sakarya University Faculty of Medicine Non-Invasive Clinical Research Ethics Committee dated 29.01.2021 and numbered E-71522473-050.01.04.32179-296. The files of patients who were diagnosed with AMI (arterial occlusion) between January 2010 and June 2019 and underwent surgical intervention were retrospectively analyzed.

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