Archives of the Balkan Medical Union Copyright © 2018 Balkan Medical Union vol. 53, no. 3, pp. 401-407 September 2018



MARKERS OF INCREASED RISK OF DEATH IN POLYTRAUMA PATIENTS WITH ASSOCIATED CHEST TRAUMA

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Received 13 July 2018 , Accepted 22 Aug 2018 https://doi.org/10.31688/ABMU.2018.53.3.14

ABSTRACT

Introduction. Polytrauma patients represent one of the most difficult pathology. Therefore, developing methods of assessing the risk of death in these patients is a continuous challenge for physicians.

The aim of the study was to evaluate the levels of creatine kinase isoenzyme MB (CK-MB) and troponin, as markers of increased risk of death in the first 24 hours after admission, in polytrauma patients with associated chest trauma.

Material and methods. The study included 33 polytrauma patients treated in the Emergency Department of the County Clinical Emergency Hospital Constanta, Romania, between 2014-2017. In all these patients, clinical evaluation and paraclinical investigations have been performed, including measurements of CK-MB levels.

The results of the study suggest that CK-MB does not offer additional information for the risk of death in these patients, while in the case of troponin, higher values were observed in patients that didn't survive after 24 hours.

Résumé

Paramètres enzymatiques à l'admission et risque de décès chez les patients polytraumatisés présentant un traumatisme thoracique associé

Introduction. Les patients polytraumatisés représentent l'une des conditions médicales les plus difficiles. Développer des méthodes pour évaluer le risque de décès chez ces patients est un effort continu pour le domaine médical.

Le but de l'étude est d'évaluer la créatine kinase isoenzyme MB (CK-MB) et la troponine en tant que marqueurs du risque accru de décès dans les 24 heures suivant l'admission chez des patients polytraumatisés présentant un traumatisme thoracique associé.

Méthodes. L'étude a été menée sur 33 patients traités dans le service des urgences de l'Hôpital Clinique d'Urgence du Département de Constanta, entre 2014 et 2017

Les résultats de l'étude suggèrent que CK-MB n'offre pas d'informations supplémentaires liées au risque de décès chez ces patients, tandis que dans le cas de la troponine, des valeurs plus élevées ont été observées

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County Clinical Emergency Hospital of Constanta, Romania Address: B-dul Mamaia 542, Bl. 2, Et. 1, Ap. 3, Constanta, Romania Email: drcristinaghinguleac@gmail.com; Phone number: +40799.990.221 **Conclusions.** The results of our study suggest that troponin values may be useful in identifying the increased risk of death in polytrauma patients.

Keywords: creatine kinase-MB, troponin, polytrauma, thoracic trauma.

INTRODUCTION

One of the most challenging emergency is polytrauma, that is an important cause of mortality and morbidity¹. Objective methods that allow a fast evaluation of the patients' outcome are crucial in the first hours following the traumatic event.

For these patients, different scores are used in order to determine the risk, like Glasgow Coma Score², Injury Severity Score³, Revised Trauma Score⁴ or the New Trauma Score⁵, the acute physiology and chronic health evaluation II⁶ or the sequential organ failure assessment⁷. While these scores offer great support for the doctors, in order to standardise the therapeutic approach, the addition of new methods of assessing the risk in patients with multiple traumas, and in particular in those with associated thoracic trauma, can offer valuable information^{8,9}.

THE OBJECTIVE OF THE STUDY was to analyze the potential of creatine kinase MB isoenzyme and troponin to be used as paraclinical tests for the initial evaluation of polytrauma patients, with thoracic trauma, in determining the risk of death within the first 24 hours after admission.

MATERIAL AND METHODS

We studied 33 patients with severe thoracic polytrauma, admitted to the emergency room in the County Clinical Emergency Hospital Constanta, Romania, between 2014-2017.

The inclusion criteria were: age over 16 years, diagnosis of major polytrauma with chest trauma. Patients with unknown mechanisms of trauma have been excluded from the study.

We compared the values of the creatine kinase isoenzyme MB(CK-MB) and troponin, on admission, in patients who survived 24 hours after admission and in patients who died during the same period of time.

In order to determine the statistical tests to run (parametric or nonparametric), we assessed the normality of the data by observing the histogram and running the Shapiro-Wilk test. Depending on the results, we ran T test or Mann-Whitney U test to compare the values among the two groups of patients. **Conclusions.** Les valeurs de la troponine semblent être utiles pour identifier les patients présentant un risque accru de décès.

Mots-clés: créatine-kinase MB, troponine, polytrauma, traumatisme thoracique.

After comparing the values, we decided to run a second type of test, in which we transformed the data according to the existing literature, considering the CK-MB cutoff value24 UI/L. For troponin, values less between 0.021 ng/mL have been considered negative, values between 0.021-0.06 ng/mL – positive +, values between 0.061-0.22 ng/mL – positive ++, and values above 0.22 ng/mL – positive +++. After the transformation, we ran Chi-square test for testing the association between the patients' status after 24 hours and enzymes values.

For all statistical tests, we considered the result as being statistically significant for p values less or equal to 0.05.

RESULTS

From the study group of 33 patients, 11 died during the first 24 hours, and 22 patients survived. The average age of the survivors group was 48.73 years, with a standard deviation of 14.01 and median value of 50. For the group of patients who died within 24 hours after admission, the average age was 50.64 years, with a standard deviation of 12.37 and median value of 57. The difference was not statistically significant (p=0.704). For both groups, males represented most of the patients, 16 patients (72.7%) being males in the survivors group and 9 patients (81.8%) being males in the group of deceased patients. There is no statistically significant difference, according to gender distribution, between the groups (p=0.566).

Creatine kinase MB (CK-MB)

Analysing the CK-MB values, we observed that average values in the case of deceased patients was more than double when compared to patients who survived (248 U/L compared to 106.64 U/L), while the median was similar, 60 UI/L (Table 1).

The Shapiro-Wilk test had statistically significant results, with p values less than 0.001 for both groups (Table 2). In Figure 1, the distribution of the patients according to CK-MB values is represented. We noted that most of the patients had values less



CK-MB Histogram

Figure 1. Distribution of CK-MB values on admission depending on evolution.

	Died within 24 hours of admission			
	No	Yes	Total	
Ν	22	11	33	
Mean	106.64	248.00	153.76	
Standard deviation	106.844	262.204	183.182	
Median	60.00	60.00	60.00	
Variance	11415.576	68751.000	33555.752	
Skewness	1.392	.699	1.640	
Kurtosis	.850	-1.317	1.919	
Minimum	19	20	19	
Maximum	350	700	700	

Table 1. Descriptive statistical analysis of CK-MB between the two groups.

Table 2. Shapiro-Wilk test for CK-MB.

	Died within 24 hours of _	Kolmogorov-Smirnov ^a			Shapiro-Wilk		
CK-MB at admission	admission	Statistic	df	Sig.	Statistic	df	Sig.
	No	.271	22	.000	.775	22	.000
	Yes	.309	11	.004	.803	11	.010

than 50 UI/L, summing up to 36% of the deceased patients and 41% of the patients in the survivors group. Based on the results of the test and the observation of the histogram, we concluded that the data do not follow a normal distribution, therefore, for comparison, we used nonparametric tests.

Ranks test (Table 3) indicates that the mean range is significantly higher for deceased patients

group, with a value of 19.64, while for the survivors is 15.98. This indicates, once more, that patients who died had higher values of CK-MB compared to the survivors.

The result of the test was not significant (z=-1/101, p=0.268) (Table 4)), suggesting that there is no statistically significant difference between the two groups, when comparing the CK-MB values.

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Table 3. CK-MB Ranks.				
	Died within 24 hours of admission	Ν	Mean Rank	Sum of Ranks
	No	22	15.68	345.00
CK-MB on admission	Yes	11	19.64	216.00
	Total	33		

	CK-MB on admission		
Mann-Whitney U	92.000		
Wilcoxon W	345.000		
Z	-1.109		
Asymp. Sig. (2-tailed)	.268		
Exact Sig. [2*(1-tailed Sig.)]	.281 ^b		





Figure 2. Distribution according to troponin.

Association between CK-MB level on admission and the risk of death

From the total of 11 patients who died during the first 24 hours after admission, one patient had normal values of CK-MB, while 10 of them had elevated values. At the same time, among the 22 patients that survived after the first 24 hours, 3 had normal values and 19 elevated values. Thus, the percentages of patients with normal values were 13.6% in the survivors group and 9.1% in the deceased group (Table 5).

By running the Chi-Square test, the result is not statistically sgnificant (p=0.7), indicating the lack of association between CK-MB category and risk of death.

Troponin

For troponin, the mean values in deceased patients were 1.24 ng/mL, with a standard deviation of 1.66 ng/mL. In patients who survived, the mean values for troponin were 0.039 ng/mL, with a standard deviation of 0.063 ng/mL. Median values are significantly different, with 0.012 ng/mL in surviving patients and 0.73 ng/mL in deceased patients (Table 6).

According to the result of Shapiro-Wilk test (Table 7) and the histogram (Figure 2), the distribution of troponin values differs, statistically significant (p<0.001), and graphical way from a normal distribution. Therefore, for comparing the values, we used the nonparametric test Mann-Whitney U.

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		Died with	in 24 hours of	admission
		No	Yes	Total
NT 1	Counts	3	1	4
Normal	% din Died within 24 hours of admission	13.6%	9.1%	12.1%
F1 1	Counts	19	10	29
Elevated	% din Died within 24 hours of admission	86.4%	90.9%	87.9%
	Counts	22	11	33
	% din Died within 24 hours of admission	100.0%	100.0%	100.0%
	Normal Elevated	Normal Counts % din Died within 24 hours of admission Elevated % din Died within 24 hours of admission Counts % din Died within 24 hours of admission % din Died within 24 hours of admission % din Died within 24 hours of admission	Died withi Died withi No Counts 3 % din Died within 24 hours of admission 13.6% Elevated Counts 19 % din Died within 24 hours of admission 86.4% Counts 22 % din Died within 24 hours of admission 100.0%	Died within 24 hours of NormalNormalCounts31% din Died within 24 hours of admission13.6%9.1%ElevatedCounts1910% din Died within 24 hours of admission86.4%90.9%Counts2211% din Died within 24 hours of admission100.0%

Table 5. Patients' evolution and CK-MB category crosstabulation

Table 6. Descriptive statistics for troponin.

	Died within 24 hours of admission			
	Yes	Total		
Ν	22	11	33	
Mean	.03955	1.24055	.43988	
Standard Deviation	.062872	1.666787	1.096050	
Median	.01200	.07300	.03000	
Variance	.004	2.778	1.201	
Skewness	2.109	.722	2.482	
Kurtosis	3.750	-1.738	4.583	
Minimum	.000	.020	.000	
Maximum	.220	3.800	3.800	

Table 7. Shapiro-Wilk test for troponin.

	Died within 24 hours	Kolmogo	rov-Smirn	ov^a	Shap	iro-Wilk	
	of admission		df	Sig.	Statistic	df	Sig.
Troponin on admission	No	.315	22	.000	.658	22	.000
	Yes	.395	11	.000	.685	11	.000

Table 8. Troponin ranks					
	Died within 24 hours of admission	Ν	Mean Rank	Sum of ranks	
	No	22	13.30	292.50	
Troponin on admission	Yes	11	24.41	268.50	
	Total	33			

Graphically, we noticed that in the case of deceased patients, none of them have troponin values less than 0.02 ng/mL, while in the case of the patients who survived the first 24 hours, almost 60% of them have values less than 0.02 ng/mL. Also, in the case of the deceased patients, 36.37% had values higher than 3 ng/mL.

Comparing the ranks, the average rank in the deceased patients group was 24,41, significantly higher when compared to the survivors group, for which the average rank was 13.3 (Table 8).

The observed difference was statistically significant (z=-3.126, p=0.002) (Table 9). This sustains, with

a high degree of confidence, that patients who survived the first 24 hours after admission had significantly lower values of troponin.

Risk of death related to troponin values

For estimating the association between troponin values and the risk of death, we crosstabulated the data. The results (Table 10) show that in the group of patients who died within 24 hours from admission, all of them were in the category positive ++ (4 patients) or positive +++ (7 patients). In the group of survivors, from the total of 22 patients, 6 were considered to be negative, 7 positive +, 5 positive ++ and 4 positive +++.

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	Troponin on admission
Mann-Whitney U	39.500
Wilcoxon W	292.500
Z	-3.126
Asymp. Sig. (2-tailed)	.002
Exact Sig. [2*(1-tailed Sig.)]	.001 ^b

Table 9. Mann Whitney-U test for troponin depending on patients' evolution.

			Died wi	thin 24 hours o	f admission
			No	Yes	Total
		Counts	6	0	6
	Negativ	% from Died within 24 hours of admission	27.3%	0.0%	18.2%
Positive + Troponin category Positive ++	Counts	7	0	7	
	Positive +	% from Died within 24 hours of admission	31.8%	0.0%	21.2%
	D	Counts	5	4	9
	Positive ++	% from Died within 24 hours of admission	22.7%	36.4%	27.3%
Positive +++		Counts	4	7	11
		% from Died within 24 hours of admission	18.2%	63.6%	33.3%
T 1		Counts	22	11	33
	Iotai	% from Died within 24 hours of admission	100.0%	100.0%	100.0%

Table 10. Patients' evolution and troponin category crosstabulation

Table 11. Chi-square test for troponin.

	Value	df	Asymptotic Significance (2-sided)		
Pearson Chi-Square	11.545ª	3	.009		
Likelihood Ratio	15.224	3	.002		
N of Valid Cases	33				
a. ^a 6 cells (75.0%) have expected count less than 5. The minimum expected count is 2.00.					

The graphical representation of the proportion (Figure 3) offers a better insight on the distribution of patients, based on troponin category.

When applying the chi-square test, the percentage of cells with expected values less than 5 is higher than 20%, therefore, we considered the statistical significance obtained from the Likelihood Ratio, which is 0.002. This result indicates a significant association between troponin category and risk of death, thus, positioning a patient into positive ++ or positive +++ indicates a significantly higher risk of death in the first 24 hours after admission.

DISCUSSION

In the case of CK-MB, we observed that almost 90% of all the patients had increased values. Combined with the fact that the percentage of patients with elevated values was almost identical in both groups (86.4% vs 90.9%), we can conclude that CK-MB is not helpful in identifying the risk of death in patients with polytrauma and associated thoracic trauma. Other studies concluded that CK-MB is not useful in determining the cardiac injury, as it is not specific enough¹⁰⁻¹².

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In the case of troponin, it proved to increase more in patients who didn't survive in the first 24 hours after admission. There are a number of studies that proved the fact that trauma patients with increased troponin values are of greater risk of death^{13,14}.

As a limitation of the study, we didn't try to identify subsequent organ (cardiac) involvement, as we considered only the main outcome, death within 24 hours from admission being the only objective of this study.

CONCLUSIONS

Increased values of creatine kinase MB are not associated with higher risk of death, while troponin values may be useful during initial assessment of polytrauma patients with thoracic trauma, in order to identify the patients with increased risk of death.



Patients distribution according to troponin category and survival at 24 hours after admission

Figure 3. Patients distribution according to troponin category and survival at 24 hours after admission

Compliance with Ethics Requirements:

"The authors declare no conflict of interest regarding this article"

"The authors declare that all the procedures and experiments of this study respect the ethical standards in the Helsinki Declaration of 1975, as revised in 2008(5), as well as the national law. Informed consent was obtained from all the patients included in the study"

"No funding for this study"

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