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GJMR-F Classification : NLMC Code: WI 140, WS 205, WD 300

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The Importance of Pediatric Scoring Systems of Multiorgan Failure in Intensive Care Unit

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Results: The mean value of PRISM score in the group of patients with favorable outcome of treatment was 8 and with lethal outcome 18. The mean value of PELOD score in the group of patients with favorable outcome of treatment was 7.7 and with lethal outcome 17.7. The area under the ROC curve for the PRISM score was 0.8306, for the PELOD score it was 0.7967. Calibration values expressed in HL-GOF for PRISM score were 2.913, while the PELOD score they were 0.60971).

Conclusion: Initial assessment, daily monitoring and reliable prediction of the final outcome of the application of pediatric scoring systems allow rising efficiency and rationalization of work in an intensive care unit.

Keywords: pediatric scoring systems, multiple organ dysfunction syndrome- MODS, intensive care unit.

I. INTRODUCTION

ysfunction and organ failure in the sepsis is very common and serious complication of the most serious ill patients. Research on various factors in explanation of sepsis occurrence, imposed a concept that was accepted at the International Conference on definitions of sepsis from 2001. The diagnosis of sepsis is based on defined criteria Association Consensus Conference Chest Physicians and Intensivists (ACCP / SCCM) [1,2] Multiorgan dysfunction syndrome (Multiple Organ Dysfunction Syndrome-MODS) is the most common cause of death in the pediatric intensive care units, with frequency range of 26-50% [3,4]. In adult patients, mortality due to MODS is of a similar value [5], according to Bilevicius and associates data the level of mortality reaches 85% [6]. It is believed that the high mortality is a direct result of the progression of organ system failure [7,8] because a direct link has been proved between mortality and the number of affected organ systems, as well as between mortality and the severity of their dysfunction [9,10]. Wilkinson, Proulx and associates also point out in their studies that the mortality caused by MODS directly depends on the number of affected organ systems [11].

Different scoring systems for the estimate of severity of illness in intensive care units emerged from the gaining of clinical experiences, primarily as a response to a question on the efficiency and quality of a treatment. In the last decade scoring system has developed on the basis of results of multicenter studies. A separate category for the estimate of the severity of illness are so-called score table which make it possible to statistical calculate the probability of survival (PS-Probability of Survival) and the probability of dying (PM-Probability of Mortality) [12]. The ideal scoring system does not exist, but a good score definitely has to meet several basic requirements: first of all, the system must be simple, mathematically consistent, research results should have high sensitivity and specificity, and environmental factors should not affect the tests which make point system [13].

The largest number of scores that are now used in intensive care units are primarily related to the adult population. Threfore, the pediatric scores that are used in clinical practice, usually resulting from the pre-existing scores for adult patients. However, the specific physiology of pediatric patients has imposed a need to establish scores that just respect these facts. These are primarily characteristic pulse and systolic pressure caused by age, specific dynamics of water and electrolyte metabolism with an increased tendency towards metabolic acidosis, a relatively small amount of circulating volume, the immaturity of immune system and the difficulty in the maintaining of body temperature.

a) Pediatric Risk Score of Mortality (PRISM)

PRISM score is used in age from the newborn to adolescence and shows the seriousness of the illness on the basis of disorders of the observed physiological 2014

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and clinical parameters, with the additional verification of pathological findings in special laboratory tests. This score, however, did not appear to be useful enough with premature babies [14]. There are several versions of this scoring system which was first applied in clinical practice under the name the physiological stability index (Physiologic Stability Index-PSI). Originally, 24 physiological parameters were followed by this score [15,16]. It was published in the literature 1986, as a dynamic assessment of the patients condition [17]. Pollack and his collaborators published a new version of the score in 1988, giving it the final name Pediatric Risk of Mortality (PRISM). This point system was by then named PRISM II score by some intensivists, which definitely separated it from the initial PSI scoring system, which was also defined as PRISM I score. PRISM II scoring system included 14 parameters, and its dynamic estimation according to the daily analysis type was first shown in 1991's [18]. Score values in the range of 0-76. Table 1 shows the parameters that make the PRISM II score.

b) Paediatric Logistic Organ Dysfunction score (PELOD)

The scoring system which is applied to the dysfunction of various organs in the pediatric age group was described in detail by Leteurtre in 1999. The score included the evaluation of the condition of the six organ systems (cardiovascular, respiratory, hepatic, renal, hematological and central nervous), and the possible existence of the dysfunction some of the selected organ systems. This score is also part of the dynamic scores, because it means a daily record of 12 observed parameters of organ dysfunction [3]. In addition, physiological parameters that are dependent on the age of patients (neonatal, infant, toddler, school age and adolescents) are continuously recorded [19,20]. Table 2 gives the parameters of PELOD score.

II. Aims

Analyzing the clinical value of pediatric scoring systems, Pediatric logistics organ dysfunction score (PELOD) and Pediatric Risk score of Mortality (PRISM), in the presence of sepsis accompanied by multiple organ dysfunction syndrome (MODS) in the surgical intensive care unit of tertiary pediatric hospital.

III. Methods

The study was conducted at the Intensive Therapy of Pediatric Surgery Clinic in Novi Sad in the period of 36 months. The method of random selection was applied, and it included 90 patients who were previously treated surgically or primarily located in the intensive care unit. On the basis of the results of analyzed scoring systems and their completed correlation with the expected and actual (real) mortality, checking out of the prognostic reliability of the examined systems carried out. The study data are numerical and they were analyzed by standard procedures of descriptive and comparative statistics. ROC (Receiver Operating Characteristics) analysis was also used in the evaluation of score value for predicting of hospital mortality. Lemeshov Hosmer goodness of fit test was also used in the evaluation of the expected and actual (real) mortality and it present the measure of the degree of calibration. [21,22,23].

IV. Results

The study included 90 patients aged 0-18 years treated at the Pediatric Surgery Clinic in Novi Sad, in the Intensive Care Unit. The average annual number of patients hospitalized in the intensive care unit of the Pediatric Surgery Clinic was 195, with a reported mortality of 8.8%. Out of the total number of patients, in 10% of patients MODS have developed, with a mortality rate of 62%. In the group of newborn babies there were total of 39 examined patients (43.3%). The group of infants had a total of 10 examined patients (11.1%). In the group of patients over 12 months there were 41 patients (45.5%). The gender distribution in the study was equal: 42 female subjects (46.6%) and 48 males (53.3%). The average length of stay patients in the ICU for the total number of patients was 10.3 days. With patients with lethal results 18.6 days, and in cured patients 8.2 days. By the analysis of outcome, 72 patients (80%) survived, and death outcome was noted in 18 (20%) patients.

a) PRISM score

For the total number of examined patients, the mean PRISM score was 10.0. The mean value of the PRISM score in the group of patients with favorable outcome was 8, and with patients with lethal results 18.

b) PELOD score

For the total number of examined patients mean PELOD score value were 9.5 The mean PELOD score value in the group with favorable outcome was 7.7. In the group with lethal outcome it was 17.7.

c) The results of ROC analysis

The showing of PRISM score values using ROC curve (Figure 1). In area under the curve in our results for the PRISM score was 0.8306, which indicates a very good prediction of PRISM score in relation to the final outcome.

d) Application of ROC analysis for PELOD score

The showing of PELOD score values using ROC curves (Figure 2). The area under the curve was 0.7967,

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which also proves a good prediction of PELOD score in relation to the final outcome.

The prediction of outcome based on the PRISM score value using logistic regression

Based on the PRISM score value, using the Hosmer-Lemeshov goodness-of-fit test, the predicted mortality was compared to the current. (Table 3).

Predicting outcomes based on the PELOD score value using logistic regression

The table 4 shows the probability of lethal outcome based on the of PELOD score values and using logistic regression. (Table 4).

Table 5 gives a collective survey of the examined scores and their comparisons. Hosmer-Lemeshov GOF, ROC curve and standard error tests were valued. Based on the comparison of the examined scores better calibration results were obtained for PELOD score, while the results of discrimination with the ROC curve indicates greater reliability PRISM score, using the statistical method.(Table 5).

V. Discussion

In recent years, the complexity of intensive treatment need for more objective assessment of weight status of patients and their ultimate prognosis. Using of scoring systems, it is possible to not only predict the final outcome, but also to compare groups of patients within one health facility or among multiple institutions.

In our study, the average number of patients hospitalized in the intensive care unit of the Pediatric Surgery Clinic at the annual level was 195, with recorded mortality of 8.8%. MODS developed in 10% with a mortality rate of 62%. In our study, despite the relatively low incidence of MODS, mortality rate is high. Data from the literature for the pediatric population are very often different, mostly because in some pediatric intensive units both pediatric and surgical patients are treated together. The data of Tantalean and associates suggest that the frequency of MODS in the intensive care unit is 25% and mortality from 26-50% [4]. Proux states that the frequency of MODS was recorded from 11-27% [9], and Wilkinson gives data on mortality of 54% [11]. The analysis of the study results considering patients age, showed that nearly half of examined patients belong to the group of infants (43.3%). According to published data from the USA, age is a significant factor in the epidemiology of sepsis. It is thought that children up to one year, especially newborns, are the patients with the significantly greatest risk of sepsis development, with even up to 10 times more bigger risk compared to the other categories of pediatric patients [8]. The total number of participants, males and females was almost equal: 46.6% female and 53.3% males. Watson and associates, in their epidemiological study of sepsis in the pediatric age, indicate that boys under 10 years occur more frequently than girls, but there is not a

significant difference in their recording mortality [10]. When it comes to the final outcome of treatment, no gender predisposing is noticed in the analyzed literature.

For the total number of observed patients, the mean PRISM score value was 10.0. The mean PRISM score value in the group of observed patients with favorable outcome was 8, and with lethal outcome it was 18. In our study, PRISM score values over 10 indicated a possible unfavorable outcome. Different from this, Tantalean and associates, in their study, with patients with lethal outcome, got results which showed the average values of PRISM score of 22.07, but also relatively high mean values of PRISM score even with patients who survived [4]. For the total number of the observed patients, the results of mean values of PELOD score were 9.5. In our survey, the mean value of PELOD score in the group with favorable outcome of treatment was 7.7, whereas in the group with lethal outcome it was 17.7. In Leteurtre and associates study, the mean value of PELOD score in patients with favorable outcome was 9.4, whereas in the group with lethal outcome mean scores were 31.0 PELOD (3). For both point systems, according to the literature data, values are significantly higher for the observed patients with lethal outcome. Considering the difference of our results compared to the literature data, the obtained results suggest the need for analysis on a greater number of the observed patients.

By using the ROC curve the values of the area under the curve were obtained for the Pediatric risk of mortality score (PRISM) which were 0.8306, whereas for the Pediatric logistic organ dysfunction score (PELOD) the values of area under the curve were 0.7967. These results show that by the use of PRISM and PELOD numerical system it is possible to achieve high reliability and safety is satisfying certainty in prediction of the final outcome of treatment. Both scores meet the ROC criteria for a good prediction, which means that the value of the area under the curve is higher than 0.70. A similar survey, with the help of ROC curve was carried out by A. Thurkal, where the validity of PRISM score was verified by the values of ROC curve 0.80 [24]. Singhal et associates by the determining of mortality prediction using the PRISM score received a value of discrimination with the ROC curve 0.72 which indicates a satisfactory value of prediction [25]. Moreno and in their multicenter study, associates got а discrimination value of PELOD score of 0.91 [26]. Research H. Iskandar survey gave the ROC analysis value for PELOD score of 0.954, and for the PRISM score 0.868. Both score in this study gave high values of the prediction of the final outcome of treatment [27]. Pedro Garcia and associates analysis of PELOD score in two pediatric intensive units, with its statistic results indicated a very good discrimination value for PELOD

score (ROC 0.93) [28]. Regardless of the different values of the results obtained by different researchers, it is clear that both point systems show strong reliability in the assessment of the final outcome prediction

A calibration degree was established by Hosmer-Lemeshov goodness-of-fit test. When Hosmer-Lemeshov goodnes-of-fit test (HL GOF) was applied in the prediction of a treatment outcome, based on the results of PRISM score, the values were 2.913 with the risk factor p = 0.405. Assessment of treatment outcomes based on the results PELOD score, applying this test the values of 0.609 with a risk factor p = 0.434. Leteurtre and associates in their multicenter study showed the values of PELOD score calibration using HL-HL GOF $\chi 2 = 4.03$ with the risk factor p = 0.54 [3]. V.F. Martha survey similarly suggests the use of PRISM score, because the calibration values PRISM score are χ 2-HL = 9.23 with the value of p = 0.10, [29]. The analysis of PELOD score, Pedro Garcia and associates in two pediatric intensive unit, according to its statistical results indicated very poor calibration values (χ 2-HL = 72.3, with risk values p lower than 0.001) [28]. Considering the difference in our results compared to literature data, where the calibration values, using HL-GOF test indicated good reliability in the prediction assessment of the final outcome of treatment with both point systems, but with slightly better results for PELOD score, the obtained results indicate the need for analysis to be carried out on a larger number of subjects. Costa and associates in their retrospective cohort study, got results in a period of one year, at a general tertiary pediatric intensive care unit. The pediatric risk of mortality score (PRISM) showed adequate discriminatory capacity and thus constitutes a useful tool for the assessment of prognosis for pediatric patients admitted to a tertiary pediatric intensive care units [29].

VI. Conclusion

The average number of hospitalized patients in Intensive Care Unit of the Pediatric Surgery Clinic, annually was 195, with an average mortality of 8.8%. The reported incidence of multiple organ dysfunction syndrome (MODS) in the intensive care unit in the study was 10%, with a mortality of 62%. Most at-risk for the development of severe sepsis were neonates.

For the total number of respondents, the median PRISM score was 10.0. The mean value of PRISM score in the group of patients with favorable outcome was 8, and with lethal results 18. The results were obtained for the mean values of PELOD score of 9.5 for the total number of respondents. In our study, the median PELOD score in the group with favorable outcome of treatment was 7.7, whereas in the group with lethal results mean was 17.7. Based on these results, it was confirmed that the higher values of the

PRISM score and PELOD directly related to an unfavorable outcome. In addition, this study confirms that both scores reliably reported and described the clinical condition of patients analyzed.

Using ROC curves the values of area under the curve were obtained for the pediatric risk of mortality score (PRISM) of 0.8306, while for the pediatric logistic organ dysfunction score (PELOD) values of area under the curve 0.7967. These results indicate that the application of the PRISM and PELOD numerical system achieves high reliability and satisfactory safety in predicting the outcome of treatment.

Using Hosmer-Lemeshov goodnes-of-fit test in the assessment the outcome, based on the results of PRISM score, the values of risk factors p = .405. Assessment of treatment outcomes based on the results PELOD score, applying this test the values of the likelihood ratio p = 0.434were obtained. Results of the application Hosmer-Lemeshov goodnes-of-fit test indicate that it is possible to predict the outcome in the regression model.

The study results show that both scores, based on statistical methods of discrimination (ROC curve) and calibration (HL-GOF test) reliably show and describe the clinical condition of patients analyzed.

Daily use and use of numeric system is needed to achieve full effectiveness of the therapy. Initial assessment, daily monitoring and reliable prediction of the final outcome of the application of pediatric scoring systems allow to raise efficiency and rationalization of work in an intensive care unit.

Previous clinical trials need to be upgraded through a multi-center study, with the aim of finding, as more reliable parameters, as new scoring systems to predict the outcome of treatment for multiple organ dysfunction in the pediatric age.

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VII. Abbreviations

PELOD-Pediatric Logistics Organ Dysfunction Scoring System

PRISM- Pediatric Risk Score of Mortality MODS- Multiple Organ Dysfunction Syndrome ICU -Intensive Care Unit

ROC-Receiver Operating Characteristics Curve HL-GOF- Hosmer-Lemeshow goodness of fit test

ACCP / SCCM -Association Consensus Conference Chest Physicians and Intensivists

SIRS- Systemic inflammatory response syndrome PIRO classification (P predisposition, I for infection, R for response of organism and O for organ dysfunction)

PS- Probability of Survival

PM- Probability of Mortality

PSI - Physiologic Stability Index

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Competing interests: none to declare.

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PARAMETER	AGE	VALUE	POINT
Systolic	Infant	130-160	2
pressure(mmHg)			
		55-65	2
		> 160	6
		40-54 < 40	6 7
	Child	150-200	2
		65-75	2
		> 200	6
		50-64	6
Diastolic pressure	All ages	< 50	7
(mmHg)		>110 > 160	6 4
Heart rate/min	Infant	< 90	4
	Child	> 150	4
Breathing frequency/min	Infant	< 80	4
		61-90	1
		> 90	5 5
	Child	apnea 51-70	5 1
		> 70	5
$Pa O_2/Fi O_2$	All ages	apnea	5
	5	200-300	2
Pa CO ₂ (mmHg)	All ages	< 200 51-65	3 1
GCS		> 65	5
Pupil reaction	All ages All ages	< 8	6
	All ages	Unequal or dilated	4
PT/PTT	All ages	Fixed and dilated	10
Total bilirubin (µmol/l)	> 1 month	1.5 puta	2 6
Potassium (µmol/l)	All ages	> 3.5 3.0-3.5	о 1
		6.5-7.5	1
		< 3.0	5
Calcium (µmol/l)	All ages	> 7.5	5
. ,	-	7.0-8.0	2
		12.0-15.0 < 7.0	2 6
Glucosa (mg/dl)	Allacoo	> 15.0	6
Giucosa (mg/ui)	All ages	40-60	4
		250-400	4
		< 40	8
Bicarbonates (µmol/l)	All ages	> 400 < 16	8 3
		> 32	3
			Ŭ

Points	0	1	10	20
ORGAN DYSFUNCTION NEUROLOGICAL GCS and Pupil reaction CARDIOVASCULAR DYSFUNCTION	12-15 or reactive	7-11 Not assessed	4-6 Both fixed	3
Heart rate/min < 12 yr = 12 yr	= 195 = 150		>195 > 150	
Sistolic TA (mmHg) < 1 month 1 month-1 yr 1-12 yr = 12 yr	> 65 > 75 > 85 > 95		35-65 35-75 45-85 55-95	< 35 < 35
RENAL DYSFUNCTION Creatinine (ųmol/l)				< 45 < 55
< 7 days 7 days- 1 yr 1-12 yr = 12 yr RESPIRATORY DYSFUNCTION	< 140 < 55 < 100 < 140		= 140 = 55 = 100 =140	
PaO ₂ (kPa/FiO ₂) Pa CO ₂ (kPa) Mehanic ventilation	> 9.3 and = 11.7 and Without MV	Ventilation	= 9.3 ili > 11.7	
HEMATOLOGY SYSTEM DYSFUNCTION				
Leukocyties (x 10 ⁹ /l) Platelets (x 10 ⁹ /l) HEPATIČ DYSFUNCTION	= 4.5 and = 35	1.5-4.4 or < 35	< 1.5	
Aspartate transaminase (IU/I) Protrombin time (or INR)	< 950 and > 60 (< 1.40)	= 950 or = 60 (= 1.40)		

Table 2 : Paediatric logistic organ	dysfunction score	(PELOD score)
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Table 3 : The probability of lethal	outcome based on PRISM score using logistic regression
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PRISM score	Expected outcome of the LR (number of patients)	Actual result (number of patients)
0-2 (18)	0,84	0
3-5 (20)	1,47	1
6-10 (18)	2,12	2
11-19 (16)	3,7	6
>20 (18)	. 9,78	9

Table 4: The probability of lethal outcome based on PELOD score using logistic regression

PELOD score	 Expected outcome of the LR (number of patients) 	Actual result (number of patients)
0-9 (32)	1,61	2
10 (33)	5,28	4
>11 (25)	11,09	12

Table 5 : Comparasion of tested values (PRISM and PELOD scores)

The values of tests	PRISM	PELOD
H-L GOF test x ² (p)	2,913 (p=0,405)	0,609 (p=0,434)
Area of ROC curve-AUC (CI	0,8306	0,7967
95%)		
Standard error AUC	0,062	0,066

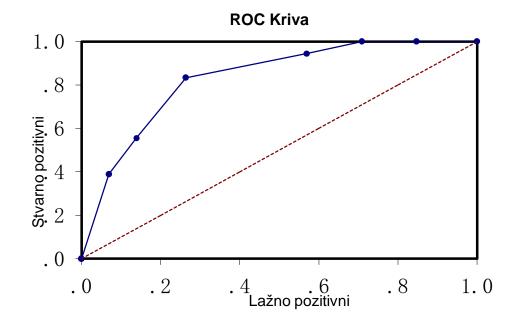


Figure 2 : Pelod score values using ROC curve