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# **ABOUT COVER**

Editorial board member of World Journal of Cardiology, Dr. Dai is a Distinguished Cardiovascular Doctor at the Yan'an Affiliated Hospital of Kunming Medical University (China). After receiving his Bachelor's degree from Kunming Medical University in 2006, he continued his postgraduate training at the same institute, receiving a Master's degree in 2009 and a PhD in 2015. He quickly rose to Deputy Chief Physician in the Cardiology Division of Yan'an Affiliated Hospital, in 2016. His ongoing research interests involve the application of evidence-based medicine in cardiovascular diseases, particularly to study coronary heart disease, heart failure, congenital heart disease, and pulmonary artery hypertension. (L-Editor: Filipodia)

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ORIGINAL ARTICLE

# **Retrospective Study**

# Risk score for predicting abdominal complications after coronary artery bypass grafting

Dmitry Vladimirovich Belov, Dmitry Victorovich Garbuzenko, Ksenia Alekseevna Abramovskikh, Nikolay Olegovich Arefyev

### **ORCID number:** Dmitry

Vladimirovich Belov 0000-0003-4985-9716; Dmitry Victorovich Garbuzenko 0000-0001-9809-8015: Ksenia Alekseevna Abramovskikh 0000-0001-8462-3819; Nikolay Olegovich Arefyev 0000-0002-1770-064X.

Author contributions: Belov DV wrote the manuscript and performed data analysis; Garbuzenko DV contributed to the conception and design of the study, acquisition, analysis and interpretation of data, and wrote the manuscript; Abramovskikh KA contributed to writing the manuscript, drafting the conception and design of the study; Arefyev NO contributed to analysis of data, and wrote and revised the manuscript; all authors have read and approved the final manuscript.

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Dmitry Vladimirovich Belov, Department of Hospital Surgery, Federal Center of Cardiovascular Surgery of the Ministry of Health of Russia (the city of Chelyabinsk), Chelyabinsk 454003, Russia

Dmitry Victorovich Garbuzenko, Department of Faculty Surgery, South Ural State Medical University, Chelyabinsk 454092, Russia

Ksenia Alekseevna Abramovskikh, Department of Hospital Surgery, South Ural State Medical University, Chelyabinsk 454092, Russia

Nikolay Olegovich Arefyev, Department of Pathological Anatomy and Forensic Medicine, South Ural State Medical University, Chelyabinsk 454092, Russia

Corresponding author: Nikolay Olegovich Arefyev, MD, Academic Researcher, Surgeon, Department of Pathological Anatomy and Forensic Medicine, South Ural State Medical University, Vorovskogo, 64, Chelyabinsk 454092, Russia. nikolai.arefyev@gmail.com

# **Abstract**

# **BACKGROUND**

Although early abdominal complications after coronary artery bypass grafting (CABG) with cardiopulmonary bypass (CPB) are rare, the associated mortality remains high.

To develop a risk score for the prediction of early abdominal complications after CABG with CPB.

# **METHODS**

This retrospective study was performed in the Federal State Budgetary Establishment "Federal Center of Cardiovascular Surgery" of the Ministry of Health of Russia (the city of Chelyabinsk) and included data of 6586 patients who underwent CABG with CPB during 2011-2017. The risk factors taken for evaluation were compared between patients with early abdominal complications ( n = 73) and without them (n = 6513). We identified the most important risk factors and their influence on the development of early abdominal complications after CABG with CPB.

personal data" filled out and signed by the patient, where the patient gave permission to use his/her personal data for conducting scientific research.

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# RESULTS

Gender and the presence of postinfarction cardiosclerosis, chronic kidney disease, or diabetes in the anamnesis did not affect the occurrence of abdominal complications. The leading risk factors of the early abdominal complications after CABG with CPB were multifocal atherosclerosis, extracorporeal membrane oxygenation, intra-aortic balloon pump, atrial fibrillation, perioperative myocardial infarction, and the need for resternotomy in the postoperative period. The average value of the predicted probability was  $0.087 \pm 0.015$  in patients with early abdominal complications after CABG with CPB and 0.0094 ± 0.0003 in patients without these complications. The percentage of correct classification turned out to be 98.9%. After calculating a score for each of the leading risk factors, we counted a total score for each particular patient. The highest risk was noted in patients with a total score of 7 or more.

# **CONCLUSION**

The developed score predicts the risk of early abdominal complications after CABG with CPB and makes it possible to stratify patients by risk groups.

Key Words: Coronary artery bypass grafting; Cardiopulmonary bypass; Abdominal complications; Risk factors; Risk score

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Core Tip: Although early abdominal complications after coronary artery bypass grafting (CABG) with cardiopulmonary bypass (CPB) are rare, the associated mortality remains high. We developed a score for the prediction of early abdominal complications after CABG with CPB. The leading risk factors were multifocal atherosclerosis, extracorporeal membrane oxygenation, intra-aortic balloon pump, atrial fibrillation, perioperative myocardial infarction, and the need for resternotomy in the postoperative period. A risk score that involves these factors makes it possible to stratify patients, which is important for timely treatment and diagnosis and, ultimately, will help to reduce postoperative mortality.

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# INTRODUCTION

Abdominal complications in patients who underwent coronary artery bypass grafting (CABG) with cardiopulmonary bypass (CPB) are rare (1.1%) but associated with high mortality, which reaches 90%[1,2]. Predicting the risk of their development may be one of the ways to solve this problem.

One way to reduce mortality in this category of patients is to predict the risk of abdominal complications[3]. Our aim was to develop a score for predicting early abdominal complications after CABG with CPB.

# MATERIALS AND METHODS

# **Patients**

We retrospectively analyzed individual medical records of 6586 patients who underwent CABG with CPB in the Federal Center of Cardiovascular Surgery of the Ministry of Health of Russia (the city of Chelyabinsk) during 2011-2017. Cases of acute cholecystitis, acute pancreatitis, acute mesenteric ischemia, intestinal necrosis, and acute intestinal obstruction that developed prior to discharge of the patient were

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referred to as early abdominal complications.

Some patients had postinfarction cardiosclerosis, chronic kidney disease, or type 1 or type 2 diabetes in the anamnesis.

Diabetes was diagnosed using one of the following diagnostic criteria: (1) Classic symptoms of diabetes (polyuria, polydipsia, ketonuria, and rapid weight loss) and a random venous plasma glucose concentration of  $\geq 11.1 \text{ mmol/L}$ ; (2) Fasting glucose concentration of  $\geq 7.0$  mmol/L in venous plasma or  $\geq 6.1$  mmol/L in whole blood; and (3) 2-h plasma glucose concentration of ≥ 11.1 mmol/L during a 75-g oral glucose

In each case, measurement of glucose concentration was repeated on a second occasion to confirm the diagnosis[4].

Chronic kidney disease was diagnosed if either of the following was present for > 3 mo: Glomerular filtration rate of < 60 mL/min/1.73 m<sup>2</sup>; One or more markers of kidney damage: (1) Albuminuria (ACR  $\geq$  30 mg/g), (2) Urine sediment abnormalities, (3) Electrolyte and other abnormalities due to tubular disorders, (4) Abnormalities detected by histology, (5) Structural abnormalities detected by imaging, and (6) History of kidney transplantation<sup>[5]</sup>.

Each medical record contained a completed and signed informed voluntary consent of the patient to the processing and use of his/her personal data and written consent to treatment. Information that was taken into account to evaluate the risk factors for abdominal complications is presented in Table 1.

# Assessment of risk factors and statistical analysis

The anamnestic risk factors and the risk factors that occurred within 2 d from the date of surgery were taken for evaluation and compared between patients with early abdominal complications (n = 73) and without them (n = 6513), and the relative risks (RRs) were calculated. Statistical data processing was performed using Statistica 10.0 (Statsoft, Tulsa, OK, United States) and SPSS 23.0 (IBM Corp., Armonk, NY, United States). In order to describe the risks, contingency tables were used to calculate the RR and the odds ratio.

An integrated score for assessing the risk of early abdominal complications was created on the basis of the above-mentioned factors by using multivariate logistic regression analysis. The presence or absence of complications was a dependent variable, while the above-mentioned factors were independent variables. Then, the obtained risk score was evaluated by using a receiver operating characteristic (ROC) curve analysis. The presence or absence of early abdominal complications was an outcome variable. The ordinate axis (sensitivity) corresponded to the true positive rate; the abscissa axis (1-specificity) corresponded to the false positive rate. It is generally supposed that the area under the ROC curve that is in the range of 0.9-1.0 should be considered as an indicator of the highest informativeness of the diagnostic method. The range of 0.8-0.9 is good, 0.7-0.8 is acceptable, 0.6-0.7 is weak, and 0.5-0.6 is extremely weak.

# RESULTS

In order to determine the probability of early abdominal complications after CABG with CPB, the RRs of their development were analyzed according to the presence and absence of the factors included in the study. Gender and the presence of postinfarction cardiosclerosis, chronic kidney disease, or type 1 or type 2 diabetes in the anamnesis did not affect the occurrence of abdominal complications.

At the same time, abdominal complications most often occurred in patients with extracorporeal membrane oxygenation (RR = 38.7), intra-aortic balloon pump (RR = 29.3), and acute myocardial infarction diagnosed after surgery (RR = 21.9). In addition, the development of early abdominal complications was possible in patients who had the following risk factors: Resternotomy (RR = 14.1); ischemic and hemorrhagic brain stroke (RR = 9.4); atrial fibrillation (RR = 8.2); multifocal atherosclerosis, namely the presence of hemodynamically significant stenoses, not only in coronary arteries but also in the brachiocephalic arteries, arteries of the upper and lower extremities, and renal arteries (RR = 4.0); CPB with the duration of more than 125 min (RR = 3.19); a combination of CABG with valve correction (RR = 3.4); aortic clamping time of more than 70 min (RR = 2.5); and age over 70 years (RR = 2.3) (Table 2).

Based on the data obtained, a multivariate logistic regression model analysis was done in order to create an integrated score for assessing the risk of early abdominal complications after CABG with CPB. The presence or absence of complications was a

Table	1 Indicators	for the eva	duation of	the risk factors	for abdominal o	complications
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Factor	Indicator	Mean (95%CI) or <i>n</i> (%)
Male sex	1-Presence/0-Absence	5164 (78.4)
Age in years	Number	61 (56 to 66)
Weight, kg	Number	82 (73 to 93)
Height, cm	Number	169 (161 to 174)
Body mass index, kg/m <sup>2</sup>	Number	29.27 (26.35 to 32.56)
Left ventricular ejection fraction, %	Number	55.69 (47.74 to 61)
Post-infarction cardiosclerosis	1-Presence/0-Absence	4429 (67.2)
Cerebrovascular accident in the anamnesis	1-Presence/0-Absence	359 (5.5)
Chronic kidney disease	1-Presence/0-Absence	917 (13.9)
Multifocal atherosclerosis	1-Presence/0-Absence	673 (10.2)
Diabetes	1-Presence/0-Absence	1452 (22)
The duration of cardiopulmonary bypass in min	Number	77 (60 to 107)
Aortic clamping time in min	Number	41 (31 to 61)
Combination of coronary artery bypass grafting with valve correction	1-Presence/0-Absence	1049 (15.9)
Resternotomy	1-Presence/0-Absence	294 (4.5)
Postoperative bleeding requiring resternotomy	1-Presence/0-Absence	182 (2.8)
Other reasons for resternotomy	1-Presence/0-Absence	100 (1.52)
Atrial fibrillation in the postoperative period	1-Presence/0-Absence	695 (10.6)
Intra- and postoperative myocardial infarction	1-Presence/0-Absence	90 (1.37)
Intra- and postoperative cerebrovascular accident	1-Presence/0-Absence	74 (1.12)
Intra-aortic balloon pump	1-Presence/0-Absence	13 (0.19)
Extracorporeal membrane oxygenation	1-Presence/0-Absence	30 (0.45)

CI: Confidence interval.

dependent variable and the six most significant risk factors were independent variables (Table 3).

It was found that the most significant independent risk factors were extracorporeal membrane oxygenation, intra-aortic balloon pump, and resternotomy in the postoperative period, followed by atrial fibrillation, perioperative acute myocardial infarction, and multifocal atherosclerosis. The average value of the predicted probability was 0.087 ± 0.015 in patients with early abdominal complications after CABG with CPB and  $0.0094 \pm 0.0003$  in patients without these complications. The percentage of correct classification turned out to be 98.9%.

To facilitate the use of this model in practice, each factor was assigned a specific score reflecting its contribution to the total risk of early abdominal complications after CABG with CPB (Table 4).

To evaluate the probability of early abdominal complications for each patient, we calculated the exact predicted probability by using the created logistic regression model and the sum of scores. Next, both of them were compared by using a ROC analysis. The criterion "presence/absence" of complications was used as an outcome variable (Table 5 and Figure 1).

The data presented in Figure 1 indicates that the ROC-curve for the indicator "sum of scores" almost coincides with that for the predicted probability and has a good diagnostic value [area under the curve (referred to as the AUC) =  $0.805 \pm 0.033$ , P < 0.001].

After the ROC curve analysis, we found it reasonable to divide the curve into four segments that would reflect the low, moderate, high, and very high risk (Table 6).

We also prospectively evaluated our scale in a group of patients who underwent CABG with CPB in 2018. The identification of a high and very high risk group for

Table 2 Qualitative risk factors of abdominal complications and relative risk of abdominal complications

Factor -		Patients with the presence of a studied factor		Patients with the absence of a studied factor		P
		Risk of abdominal complications, <i>n</i> (%)	n	Risk of abdominal complications, n (%)	risk	value
Extracorporeal membrane oxygenation	30	11 (36.67)	6556	62 (0.95)	38.7	< 0.001
Intra-aortic balloon pump	13	4 (30.77)	6573	69 (1.05)	29.3	< 0.001
Perioperative myocardial infarction	90	17 (18.89)	6496	56 (0.86)	21.9	< 0.001
Resternotomy	294	29 (9.86)	6292	44 (0.7)	14.1	< 0.001
Cerebrovascular accident in the perioperative period	74	7 (9.46)	6512	65 (1.0)	9.4	< 0.001
Atrial fibrillation in the postoperative period	695	36 (5.18)	5891	37 (0.63)	8.2	< 0.001
Multifocal atherosclerosis	673	23 (3.42)	5913	50 (0.85)	4.0	< 0.001
Combination of coronary artery bypass grafting with valve correction	1049	29 (2.76)	5537	44 (0.79)	3.4	< 0.001
Cardiopulmonary bypass with the duration of more than 125 min	1238	31 (2.5)	5348	42 (0.79)	3.19	< 0.001
Aortic clamping time of more than 70 min	1374	29 (2.11)	5212	44 (0.84)	2.5	< 0.001
Age over 70 yr	830	18 (2.17)	5756	55 (0.96)	2.3	0.0018
Cerebrovascular accident in the anamnesis	359	8 (2.23)	6227	65 (1.04)	2.1	0.037
Chronic kidney disease	917	14 (1.53)	5669	59 (1.04)	1.4	0.19
Male sex		60 (1.16)	1422	13 (0.91)	1.3	0.43
Postinfarction cardiosclerosis		48 (1.08)	2157	25 (1.16)	0.94	0.78
Diabetes		10 (0.69)	5134	63 (1.23)	0.56	0.084

Table 3 Coefficients of the model of multivariate logistic regression (Nagelkerke R Square = 0.966, P < 0.0001)						
Risk fac	tor	B ± SE	Exp (B)	P value		
Multifoca	al atherosclerosis	0.74 ± 0.28	2.10	0.001		
Extracor	poreal membrane oxygenation	$2.323 \pm 0.546$	10.207	< 0.001		
Intra-aor	tic balloon pump	$1.762 \pm 0.887$	5.824	0.047		
Resterno	tomy	1.922 ± 0.316	6.833	< 0.001		
Atrial fib	rillation in the postoperative period	$1.62 \pm 0.262$	5.054	< 0.001		
Periopera	ative myocardial infarction	$1.384 \pm 0.415$	3.991	0.001		

 $5.508 \pm 0.193$ 

B: Coefficient of regression; SE: Standard error.

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abdominal complications contributed to early diagnosis in 14 patients. The number of lethal cases was 20 (27.4%) for 2011-2017 and decreased to 2 (14.3%) cases in 2018 (P <0.05). Hence, the hospital mortality decreased to 52.2% in the group of patients with early abdominal complications.

0.004

The division of patients into risk groups according to the development of abdominal complications after CABG with CPB suggests a differentiated approach to their management in the postoperative period. Patients with a score of more than 7 according to our score require particular attention.

< 0.001

Table 4 Distribution of scores assigned to the risk factors for abdominal complications					
Risk factor	Score				
Multifocal atherosclerosis	3				
Extracorporeal membrane oxygenation	10				
Intra-aortic balloon pump	6				
Atrial fibrillation in the postoperative period	5				
Perioperative myocardial infarction	4				
The need to perform postoperative resternotomy (excluding patients with extracorporeal membrane oxygenation)	7				

Table 5 Results of receiver operating characteristic curve analysis				
Factor	AUC	SE (AUC)	95%CI	P value
The exact predicted probability	0.805	0.033	0.739-0.869	< 0.001
Total score	0.805	0.033	0.740-0.871	< 0.001

AUC: Area under the curve; CI: Confidence interval; SE: Standard error.

Table 6 Distribution of patients according to the risk groups of abdominal complications after coronary artery bypass grafting with cardiopulmonary bypass

Risk group	Score	Risk of abdominal complications, %	n	The percentage of the risk group in the general population, %
Low risk of abdominal complications	0	0.4	5072	77.0
Moderate risk of abdominal complications	1-6	1.2	1047	15.9
High risk of abdominal complications	7-12	9.4	421	6.4
Very high risk of abdominal complications	≥13	44	46	0.7

# DISCUSSION

Abdominal complications after CABG with CPB are found in 0.2%-5.5% of operated patients<sup>[6]</sup> and are accompanied by a mortality rate from 11% to 74%<sup>[7]</sup>. Therefore, it is relevant to identify risk factors for predicting the development of abdominal complications.

Long-term CPB is accompanied by a wide range of pathophysiological disorders, which include non-pulsating blood flow, activation of hormonal immunity, anticoagulation, hypothermia, decreased organ perfusion, redistribution of blood flow, threat of embolism, and hyperkalemia<sup>[6]</sup>. It is known that subphysiological blood flow, the release of endogenous vasoconstrictors, including angiotensin II, and the subsequent increase in systemic vascular resistance during CPB may lead to abdominal ischemia, especially if atherosclerotic lesions are present<sup>[8,9]</sup>.

An increase in aortic clamping time may lead to low cardiac output and/or release of inflammatory mediators[10]. In addition, if CABG is combined with valve surgery, the risk of mesenteric embolism increases[11].

In patients with postoperative bleeding that requires a resternotomy and in the case when a ventilation time is more than 24 h, there is a decrease in cardiac output and in mean arterial pressure and an increase in vascular resistance of internal organs, which induce temporary hypoperfusion[12]. These disorders of splanchnic hemodynamics are exacerbated by the activation of the renin-angiotensin-aldosterone system and increased levels of catecholamines.

Severe acute perioperative myocardial infarction is accompanied by a decrease in cardiac output, systemic hypotension with the centralization of blood circulation, and a decrease in mesenteric perfusion<sup>[13,14]</sup>. They are also facilitated by the following factors: (1) Vasopressors that are used to treat myocardial infarction[15]; (2)

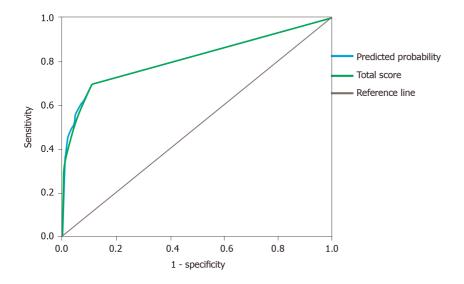


Figure 1 Receiver operating characteristic curve for the predicted probability and the sum of scores.

Prolongation of CPB and mechanical ventilation[16]; (3) Resternotomy for reshunting<sup>[6,8,17]</sup>; (4) Extracorporeal membrane oxygenation<sup>[16,18]</sup>; and (5) Intra-aortic balloon counterpulsation, when there is thromboembolism of visceral arteries or their obstruction due to incorrect positioning[17,19].

Atrial fibrillation in the postoperative period contributes to the embolism of the mesenteric vasculature<sup>[12]</sup>, which is also possible during manipulations on the aorta in multifocal atherosclerosis  $^{\text{[10]}}.$  In the latter, hemodynamically insignificant stenoses in the superior mesenteric artery exacerbate the impairment of mesenteric perfusion caused by hypovolemia and hypotension. In addition, multifocal atherosclerosis may act as an independent risk factor of early abdominal complications in patients older than 70 years<sup>[18]</sup>.

Comparing the literature data with the results of our own research, we identified the most significant risk factors and used them to develop a prognostic score, which had a good diagnostic value (AUC =  $0.805 \pm 0.033$ , P < 0.001). This score allows us to stratify patients with the risk for early abdominal complications after CABG with CPB, which is important for timely treatment and diagnosis and, ultimately, will help to reduce postoperative mortality<sup>[20]</sup>.

# CONCLUSION

In conclusion, the leading risk factors for early abdominal complications after CABG with CPB in our study were multifocal atherosclerosis, extracorporeal membrane oxygenation, intra-aortic balloon pump, atrial fibrillation, acute perioperative myocardial infarction, and the need for performing postoperative resternotomy. According to the developed score, patients with a score of 7 or more had the highest risk of abdominal complications. The score makes it possible to predict the risk of early abdominal complications after CABG with CPB and to stratify patients with the risk for their development, which is important for timely treatment and diagnosis and, ultimately, will help to reduce postoperative mortality.

# ARTICLE HIGHLIGHTS

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# Research background

Abdominal complications in patients who underwent cardiac surgery are rare but the associated mortality varies from 11.0% to 74.0%, which makes the problem relevant. One reason for high mortality rates is late diagnosis. Certain difficulties in diagnosing these complications are associated with the peculiarities of postoperative management and unclear clinical picture due to sedation, analgesia, prolonged mechanical ventilation, and the use of extracorporeal membrane oxygenation. Determining the risk factors for abdominal complications and identifying high-risk groups is an urgent task that may help diagnose abdominal complications and reduce related mortality.

# Research motivation

The topic of this study is the identification of leading risk factors for abdominal complications in patients after coronary artery bypass grafting (CABG) with cardiopulmonary bypass (CPB) and the development of a prediction score based on perioperative predictors to reveal patients at a high risk of abdominal complications.

# Research objectives

The objectives of this study were to identify risk factors for abdominal complications according to clinical, laboratory, and imaging findings, to determine their rank influence on the development of early abdominal complications after CABG with CPB, and to develop a risk prediction score. The risk score makes it possible to perform early screening and to develop an algorithm of therapeutic and diagnostic measures for the prevention, early diagnosis, and treatment of this pathology.

# Research methods

The factors taken for evaluation were compared between patients with early abdominal complications (n = 73) and without them (n = 6513), and the relative risks were calculated. Statistical data processing was performed using Statistica 10.0 (Statsoft, Tulsa, OK, United States) and SPSS 23.0 (IBM Corp., Armonk, NY, United States). In order to describe the risks, contingency tables were used to calculate the relative risk and the odds ratio. An integrated score for assessing the risk of early abdominal complications was created on the basis of the above-mentioned factors by using multivariate logistic regression analysis. The presence or absence of complications was a dependent variable, while the above-mentioned factors were independent variables. Then, the obtained risk score was evaluated by using a receiver operating characteristic curve analysis. The presence or absence of early abdominal complications was an outcome variable. The ordinate axis (sensitivity) corresponded to the true positive rate; the abscissa axis (1-specificity) corresponded to the false positive rate. It is generally supposed that the area under the receiver operating characteristic curve that is in the range of 0.9-1.0 should be considered as an indicator of the highest informativeness of the diagnostic method. The range of 0.8-0.9 is good, 0.7-0.8 is acceptable, 0.6-0.7 is weak, and 0.5-0.6 is extremely weak.

# Research results

The leading risk factors of the early abdominal complications after CABG with CPB were multifocal atherosclerosis, extracorporeal membrane oxygenation, intra-aortic balloon pump, atrial fibrillation, perioperative myocardial infarction, and need for resternotomy in the postoperative period. After calculating a score for each of the leading risk factors, we counted a total score for each particular patient. The highest risk was noted in patients with a total score of 7 or more. Further research may be devoted to a prospective assessment of the proposed scale for the prediction of the risk of abdominal complications and the development of an algorithm for the management of high-risk patients.

# Research conclusions

The developed score predicts the risk of early abdominal complications after CABG with CPB and makes it possible to stratify patients by risk groups. It is important for timely treatment and diagnosis and, ultimately, will help to reduce postoperative mortality in this group of patients.

# Research perspectives

Future studies should prospectively assess the effectiveness of the proposed method, as well as to create a protocol for the prevention, early diagnosis, and treatment of abdominal complications in high-risk patients.

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