# COVID-19 as an independent predictor of aspiration thrombectomy in STEMI. National data from the ORPKI register in the years 2020–2022

# Patrycja Zając¹, Karol Kaziród-Wolski², Janusz Sielski², Magdalena Wolska³, Krzysztof Piotr Malinowski⁴, Zbigniew Siudak²

<sup>1</sup>Rheumatology Department of the Province Hospital, Końskie, Poland

<sup>2</sup>Collegium Medicum, Jan Kochanowski University, Kielce, Poland

<sup>3</sup>Outpatient Treatment Facility "CenterMed", Kielce, Poland

<sup>4</sup>Department of Bioinformatics and Telemedicine, Jagiellonian University Medical College, Krakow, Poland

Adv Interv Cardiol 2023; 19, 2 (72): 119–126 DOI: https://doi.org/10.5114/aic.2023.127893

#### Abstract

**Introduction**: Coronavirus disease 2019 (COVID-19) exacerbates intravascular thrombosis that occurs in the coronary artery in ST-elevation myocardial infarction (STEMI).

Aim: To analyze the impact of COVID-19 on the application and effect of thrombectomy in STEMI patients.

**Material and methods:** 29915 STEMI patients were analyzed, of whom 3139 (10.5%) underwent thrombectomy. COVID-19 (+) was reported in 311 (10.8%). The clinical characteristics and management of STEMI in COVID-19 (+) and COVID-19 (–) patients were compared. A multivariable logistic regression analysis was performed in search of factors influencing thrombectomy.

**Results:** COVID-19 (+) patients had higher Killip class (IV class; n = 33 (12.31%) vs. n = 138 (5.84%); p < 0.0001) and cardiac arrest at baseline was more frequent in this group (n = 25 (8.04%) vs. n = 137 (4.84%); p = 0.016). Thrombolysis in myocardial infarction (TIMI) 3 after percutaneous coronary intervention was less frequent (n = 248 (80.52%) vs. n = 2388 (87.19%); p = 0.001) in the COVID-19 (–) group. Periprocedural mortality was similar in both groups (n = 28 (0.99%) vs. n = 4 (1.29%); p = 0.622). In multivariable regression analysis COVID-19 increased the risk of thrombectomy (OR = 1.23; 97.5% Cl: 1.05–1.43; p = 0.001).

**Conclusions:** STEMI patients undergoing aspiration thrombectomy who were COVID-19 (+) were more likely to be in a severe clinical condition (higher Killip class, more frequent cardiac arrest before the procedure) than COVID-19 (–) patients. Despite more intensive antiplatelet and anticoagulant treatment, PCI procedures were less likely to result in an optimal TIMI 3 effect. COVID-19 is an independent strong predictor of patient qualification for aspiration thrombectomy in STEMI.

Key words: thrombectomy, percutaneous coronary intervention, acute coronary syndrome, COVID-19, myocardial infarction, coronary thrombosis.

#### Summary

The authors found that coronary thrombosis in patients undergoing thrombectomy because of ST-elevation myocardial infarction with concomitant COVID-19 is more severe. In these patients percutaneous coronary intervention less frequently ends in an optimal result. COVID-19 is an independent and strong predictor of patient qualification for aspiration thrombectomy in ST-elevation myocardial infarction.

# Introduction

Initially, with SARS-CoV-2 coronavirus infections, the main focus was placed on the infectious symptoms and the pulmonary complications that could eventually lead to pro-

gression of respiratory failure and, in many cases, inevitably lead to the patient's death. It was quickly recognized that infection with the new coronavirus could generate symptoms from other systems, likely due to the prevalence of the receptor by which the virus enters the host cell. One of

#### Corresponding author:

Karol Kaziród-Wolski MD, PhD, Collegium Medicum, Jan Kochanowski University, Kielce, Poland, phone: +48 41 36 71 493, e-mail: karol.kazirod-wolski@ujk.edu.pl **Received:** 28.12.2022, **accepted:** 6.03.2023. the first observations made was the effect of infection on the coagulation system. It was reported that patients with coronavirus disease 2019 (COVID-19) had an increased incidence of complications such as ischemic stroke, pulmonary embolism and acute coronary syndromes [1–3]. It has not been fully clarified what leads to the procoagulant state in COVID-19. Most likely, there are complex mechanisms between the innate immune response, coagulation and fibrinolysis pathways and vascular endothelial damage, as well as individual variables including mutations of factor V Leiden and prothrombin G20201A [4, 5]. The pathology observed in the course of COVID-19 in a mouse model was similar to that observed in patients with severe COVID-19, with evidence of diffuse bronchoalveolar injury with edema, fibrin deposits, leukocyte infiltration and pneumocyte hypertrophy [4]. Also there are significant histopathological changes in the accumulation of T lymphocytes (CD4-CD8 located around blood vessels are visible in the post-mortem examination of COVID-19 patients) [6]. The impact of disruptions of the fibrinolysis system in the course of COVID-19 infection may play a significant role not only in terms of cardiac diseases. The importance of this system and its disorders in the course of infection is also confirmed by the observations of other specialists. A complication that can be observed in patients with severe COVID-19 is pulmonary fibrosis, in which the formation of fibrin in the intraalveolar compartment and proteases of the coagulation system and plasminogen activation (plasminergic) system play an important role, which respectively create and break down fibrin [7]. In a study of the effect of drugs on pulmonary fibrosis in mice, lower fibrinogen levels and higher levels of tPA, the main intravascular activator of fibrinolysis, were found in the treatment group, which correlated with inhibition of fibrosis progression in mice [8]. Experts suggest that therapy to reduce fibrotic lesions used in the early stages of SARS-CoV-2 infection may have significant benefits [9].

It seems that the procoagulant potential in the course of SARS-CoV-2 virus infection may be enhanced in the course of acute coronary syndrome (ACS), where coagulation also occurs. As a result, this additive effect may lead to a more severe course and be associated with a higher risk of death [10, 11].

# Aim

The aim of the study is to determine whether and how COVID-19 affects the clinical parameters of ST-elevation myocardial infarction (STEMI) patients undergoing aspiration thrombectomy during percutaneous coronary intervention (PCI) as well as the application and effect of thrombectomy in STEMI patients.

# Material and methods

In order to determine the most important factors that, due to the enhanced clotting potential of SARS-

CoV2 coronavirus infection, may have affected thrombus formation in the coronary vessel, as well as whether COVID-19 infection alone affected the frequency of aspiration thrombectomy, we analyzed patients with STEMI, based on the National Registry of Invasive Cardiology Procedures (ORPKI) [10].

The study group consisted of patients undergoing invasive treatment for STEMI from March 4, 2020, when the first COVID-19 infection was confirmed in Poland, until March 4, 2022.

The group of patients collected for analysis, namely those who qualified for treatment of STEMI and were registered in ORPKI, included 29915 patients, of whom 3139 underwent aspiration thrombectomy. This group consisted of 2828 COVID-19 negative patients and 311 patients who were diagnosed with COVID-19. The antigen tests were used to confirm positive cases, either in the ambulance or at the destination hospital. Due to the need to adhere to time standards, there was no waiting for PCR test results. Patients with suspected COVID-19 (as recommended for triage by the National Institute of Public Health and the Ministry of Health) were treated as potentially COVID-19 (+). A diagnosis of COVID-19 was always available before any interventional procedure (angiography or percutaneous coronary intervention) and was recorded in the ORPKI online database. Swabs for molecular RT-PCR were always collected before the procedure. The study included cases of patients who underwent primary PCI; rescue and facilitated PCI were excluded. The time from symptoms onset to PCI was less than 24 h.

We carried out a pooled analysis of factors predisposing to aspiration thrombectomy in STEMI patients undergoing invasive treatment. The mentioned patients received acetylsalicylic acid (ASA), P2Y12 or anticoagulation during PCI, which was included in the ORPKI registry; the remaining patients received the drugs earlier, which was not shown in the ORPKI registry. The ORPKI registry does not provide specific information about thrombolysis during PCI. Data on thrombus burden grades were not collected as part of the ORPKI registry. Chronic kidney disease was defined in compliance with the recommendations as the presence of an estimated glomerular filtration rate (eGFR) of less than 60 ml/min/1.73 m<sup>2</sup>. Two groups of patients were compared: patients with STEMI but without confirmed infection (COVID-19(-)) and patients with STEMI with confirmed infection (COVID-19(+)). We carried out multivariable regression analysis of clinical, prehospital and pharmacological factors to the endpoint of aspiration thrombectomy. The authors selected variables with a potential impact on the course of PCI; these were mainly clinical parameters. Patients qualified for invasive treatment signed informed consent forms in accordance with the recommendations of the 1964 Declaration of Helsinki. As we used anonymous data from the ORPKI database, the study did not require approval from the Bioethics Committee.

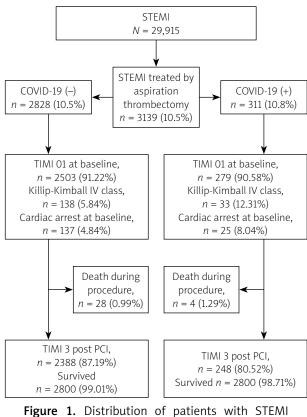
#### Statistical analysis

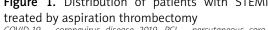
Nominal variables were presented as counts and percentages, whereas continuous variables were presented as means with standard deviation as well as median with the first and the third quartiles. Normality was assessed by Shapiro-Wilk test. Equality of variances was assessed using Levene's test. Differences between two groups were compared using Student's or Welch's t test depending on the equality of variances for normally distributed variables. The Mann-Whitney U test was used for non-normally distributed continuous variables. Comparison of nominal variables between analyzed groups was performed using the Pearson  $\chi^2$  test or Fisher's exact test. Multivariable logistic regression models were used to estimate odds ratios with 95% confidence intervals and p-values. A p-value of less than 0.05 was considered significant. No missing data imputation was performed. All available data were analyzed. The final model was run on all cases. The outcome was thrombectomy and exposure was COVID-19. Predictors were age, gender, weight, diabetes, previous myocardial infarction (MI), previous PCI, previous coronary artery bypass grafting (CABG), smoking status, psoriasis, arterial hypertension, kidney disease, chronic obstructive pulmonary disease (COPD), Killip class at admission and cardiac arrest at baseline. Effect modifiers were advanced forms of coronary artery disease (stenosis of left main coronary artery (LMCA), multivessel disease with or without stenosis of LMCA). Confounders were time from pain to first medical contact (FMC) and time from FMC to angiography. All statistical calculations were performed with R version 4.1.3 (R Core Team, Vienna, Austria, 2022).

# Results

Among 29915 patients with STEMI included in the study 3139 underwent aspiration thrombectomy. 2828 patients were COVID-19 (–) and 311 patients were COVID-19 (+).

Patients' distribution is presented in Figure 1. Baseline characteristics of clinical factors showed that patients with STEMI and concomitant COVID-19 who had undergone aspiration thrombectomy were in a higher Killip class (class IV; n = 33 (12.31%) vs. n = 138 (5.84%); p < 0.0001), more frequently had chronic obstructive pulmonary disease (COPD) (n = 66 (2.33%) vs. n = 16(5.14%); p = 0.003), chronic total occlusion (CTO) (n = 63)(2.23%) vs. *n* = 13 (4.18%); *p* = 0.034), had higher weight (*n* = 82 (73; 95) vs. *n* = 85 (77; 95); *p* = 0.026) and longer time from first contact to inflation or angiogram in minutes (75 (55; 120) vs. 90 (58.5; 142.5); *p* < 0.001) as well as being more likely to experience out-of-hospital cardiac arrest (n = 137 (4.84%) vs. n = 25 (8.04%); p = 0.016) (Table I). Patients with STEMI and concomitant COVID-19 who underwent aspiration thrombectomy during PCI more often received unfractionated heparin (UFH) (n = 2295 (81.15%) vs. n = 268 (86.17%); p = 0.03), acetylsalicylic acid (n = 908 (32.11%) vs. n = 124 (39.87%); p = 0.006), clopidogrel (n = 370 (13.08%) vs. n = 50(16.08%); p = 0.001) and ticagrelor (n = 843 (29.81%) vs.)n = 119 (38.26%); p = 0.001) as well as bivalirudin (n =20 (0.71%) vs. 6 (1.93%); *p* = 0.024). Total radiation dose was higher in COVID-19 patients (551 (315; 938) vs. 621 (376.5; 1130.5); *p* < 0.001) and these patients required thrombolysis during PCI more often (n = 24 (0.85%) vs. n = 9 (2.89%); p < 0.001). Post-PCI Thrombolysis In Myocardial Infarction (TIMI) 3 flow was obtained less frequently (n = 2388 (87.19%) vs. n = 248 (80.52%);p < 0.001) (Table II). Multivariable regression analysis showed that independent predictors of thrombectomy were Killip class IV (OR = 2.06 (1.66-2.55); p < 0.001), COVID-19 (OR = 1.23 (1.05-1.43); p = 0.001), active smoking status (OR = 1.18 (1.07 - 1.31); p = 0.001), male gender (OR = 1.13 (1.01–1.27); *p* = 0.032), age (OR = 0.99 (0.98–0.99); *p* < 0.001), weight (OR = 1.01 (1.002–1.008); p = 0.003), multivessel disease with left main coronary artery (OR = 0.81 (0.74–0.89); p < 0.001) or without LMCA stenosis (OR = 0.63 (0.51–0.77); *p* < 0.001) and time from first medical contact to angiography (OR = 0.99 (0.9987-(0.9996); p = 0.001) (Table III). Factors affecting qualification for thrombectomy are also shown in Figure 2.





COVID-19 – coronavirus disease 2019, PCI – percutaneous coronary intervention, STEMI – ST-elevation myocardial infarction, TIMI – Thrombolysis In Myocardial Infarction.

<b>Table I.</b> Baseline characteristics of clinical factors in STEMI patients ( $n = 29915$ ) undergoing aspiration throm-
bectomy during PCI

Aspiration thrombectomy	Total (n = 3139; 10.5%)	COVID-19 (–) (n = 2828; 10.5%)	COVID-19 (+) (n = 311; 10.8%)	P-value
Age [years]	64 (56; 71)	64 (56; 71)	64 (55; 71)	0.555
Gender, male	2311 (73.76%)	2068 (73.26%)	243 (78.39%)	0.051
Weight [kg]	82 (74; 95)	82 (73; 95)	85 (77; 95)	0.026
Diabetes, n (%)	511 (16.28)	452 (15.98)	59 (18.97)	0.195
Previous stroke, n (%)	82 (2.61)	75 (2.65)	7 (2.25)	0.674
Previous MI, n (%)	387 (12.33)	342 (12.09)	45 (14.47)	0.226
Previous PCI, n (%)	380 (12.11)	337 (11.92)	43 (13.83)	0.327
Previous CABG, n (%)	58 (1.85)	48 (1.70)	10 (3.22)	0.059
Smoking status, n (%)	1157 (36.86)	1054 (37.27)	103 (33.12)	0.15
Psoriasis, n (%)	13 (0.41)	12 (0.42)	1 (0.32)	0.79
Arterial hypertension, n (%)	1802 (57.41)	1618 (57.21)	184 (59.16)	0.509
COPD, n (%)	82 (2.61)	66 (2.33)	16 (5.14)	0.003
Killip class:				
	2107 (80.05%)	1923 (81.35%)	184 (68.66%)	< 0.0001
II	272 (10.33%)	241 (10.19%)	31 (11.57%)	•
	82 (3.12%)	62 (2.62%)	20 (7.46%)	-
IV	171 (6.50%)	138 (5.84%)	33 (12.31%)	-
Cardiac arrest at baseline, n (%)	162 (5.16)	137 (4.84)	25 (8.04)	0.016
Time from pain to first contact [min]	120 (60; 360)	120 (60; 360)	120 (60; 480)	0.733
Time from pain to inflation or angiogram [min]	220 (135; 530)	220 (135; 504.5)	250 (135; 805.5)	0.05
Time from first contact to inflation or angiogram [min]	75 (55; 120)	75 (55; 120)	90 (58.5; 142.5)	< 0.001
Angiography result:				
1-vessel	1559 (49.67%)	1403 (49.61%)	156 (50.16%)	0.979
Multivessel	1418 (45.17%)	1278 (45.19%)	140 (45.02%)	
Multivessel + LMCA	154 (4.91%)	140 (4.95%)	14 (4.50%)	
LMCA	8 (0.25%)	7 (0.25%)	1 (0.32%)	
СТО, п (%)	76 (2.42)	63 (2.23)	13 (4.18)	0.034

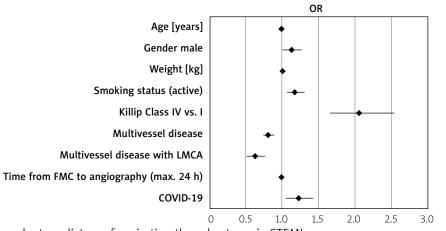
CABG – coronary artery bypass grafting, COPD – chronic obstructive pulmonary disease, COVID-19 – coronavirus disease 2019, CTO – chronic total occlusion, LMCA – left main coronary artery, MI – myocardial infarction, PCI – percutaneous coronary intervention, STEMI – ST-elevation myocardial infarction.

# Discussion

In the course of COVID-19 infection, an increased incidence of thromboembolic complications was very quickly noted, and disorders of the hemostasis system became distinguished from the previously known coagulopathy, which was referred to as COVID-19-associated coagulopathy (CAC). Equally quickly, a different course of STEMI in the course of SARS-CoV2 infection began to be observed. It was found that some of the infected patients undergoing invasive examination did not have an identifiable coronary lesion [12], and the occurrence of non-obstructive coronary artery disease was significantly more frequently observed in COVID-19 STEMI than in the control group (13% vs. 1%, p = 0.03) [13]. In addition, there have been reports that this problem occurred with an increased frequency in women (1/3 of women vs. 1/5 of men) [14]. Information is available in the literature on cases of patients who developed coronary artery embolism during COVID-19 infection. During angiography, thrombectomy was performed, and due to the absence of a coronary lesion, stent implantation was abandoned, and then anticoagulant therapy was continued with a satisfactory effect [15]. Similar clinical cases have also been reported by other authors [16]. The angiographic image in the form of extensive and multivessel thrombosis, regardless of the presence of atherosclerotic plaques, poses a new therapeutic challenge and may be associated with an increase in the incidence of stent thrombosis [17], which in in-hospital settings may increase up to five times [18]. In a large multicenter, retrospective study in patients with COVID-19 and STEMI disease, the percentage of stent thrombosis in patients undergoing PCI treatment was estimated to be 21% vs. 1% compared to the control group [19]. Interesting conclusions were provided by the study of Cornelissen et al., who assessed the inflammatory potential and thrombogenicity of different stents using the blood of healthy volunteers, enriched with high levels of IL-6 and TNF- $\alpha$  to simulate a cytokine storm, similar to that occurring in the course of COVID-19. The study indi 
 Table II. Descriptive characteristics of procedural factors in STEMI patients undergoing aspiration thrombectomy during PCI

Aspiration thrombectomy	Total (n = 3139; 10.5%)	COVID-19 (–) (n = 2828; 10.5%)	COVID-19 (+) (n = 311; 10.8%)	<i>P</i> -value
Infarct-related artery:				
LMCA	53 (1.69%)	48 (1.70%)	5 (1.61%)	0.907
LAD	1245 (39.66%)	1108 (39.18%)	137 (44.05%)	0.096
Cx	298 (9.49%)	268 (9.48%)	30 (9.65%)	0.923
RCA	1481 (47.18%)	1345 (47.56%)	136 (43.73%)	0.199
LIMA-LAD	3 (0.10%)	2 (0.07%)	1 (0.32%)	0.269
SVG	36 (1.15%)	31 (1.10%)	5 (1.61%)	0.421
Number of implanted stents:				
0	341 (10.86%)	306 (10.82%)	35 (11.25%)	0.94
1	2400 (76.46%)	2160 (76.38%)	240 (77.17%)	
2	342 (10.90%)	310 (10.96%)	32 (10.29%)	
3	54 (1.72%)	50 (1.77%)	4 (1.29%)	
4	2 (0.06%)	2 (0.07%)	0 (0.00%)	
TIMI 01 at baseline, n (%)	2782 (91.15%)	2503 (91.22%)	279 (90.58%)	0.711
Post-PCI TIMI 3, n (%)	2636 (86.51%)	2388 (87.19%)	248 (80.52%)	0.001
Total amount of contrast used during procedure [ccm]	150 (120; 200)	150 (120; 200)	150 (120; 200)	0.682
Total radiation dose during procedure [mGy]	558 (321; 959.25)	551 (315; 938)	621 (376.5; 1130.5)	< 0.001
Death during procedure, n (%)	32 (1.02%)	28 (0.99%)	4 (1.29%)	0.622
No reflow, n (%)	111 (3.54%)	95 (3.36%)	16 (5.14%)	0.107
Cardiac arrest during PCI, n (%)	48 (1.53%)	42 (1.49%)	6 (1.93%)	0.545
GPI IIb/IIIa, n (%)	1635 (52.09%)	1472 (52.05%)	163 (52.41%)	0.9
ASA during PCI, n (%)	1032 (32.88%)	908 (32.11%)	124 (39.87%)	0.006
UFH during PCI, n (%)	2563 (81.65%)	2295 (81.15%)	268 (86.17%)	0.03
LMWH during PCI, n (%)	9 (0.29%)	7 (0.25%)	2 (0.64%)	0.216
Thrombolysis during PCI, n (%)	33 (1.05%)	24 (0.85%)	9 (2.89%)	< 0.001
Bivalirudin during PCI, n (%)	26 (0.83%)	20 (0.71%)	6 (1.93%)	0.024
P2Y12 during PCI:				
None	1708 (54.41%)	1568 (55.45%)	140 (45.02%)	0.001
Clopidogrel	420 (13.38%)	370 (13.08%)	50 (16.08%)	
Prasugrel	49 (1.56%)	47 (1.66%)	2 (0.64%)	
Ticagrelor	962 (30.65%)	843 (29.81%)	119 (38.26%)	

ASA – acetylsalicylic acid, Cx – critical stenosis of circumflex artery, COVID-19 – coronavirus disease 2019, GPI IIb/IIIa – IIb/IIIa glycoprotein inhibitor, LAD – critical stenosis of left anterior descending artery, LIMA-LAD – critical stenosis of left internal mammary artery – left anterior descending bypass, LMCA – critical stenosis of left main coronary artery, LMWH – low molecular weight heparin, PCI – percutaneous coronary intervention, RCA – critical stenosis of right coronary artery, STEMI – ST-elevation myocardial infarction, SVG – critical stenosis of saphenous vein graft, TIMI – Thrombolysis In Myocardial Infarction, UFH – unfractionated heparin.



**Figure 2.** Independent predictors of aspiration thrombectomy in STEMI COVID-19 – coronavirus disease 2019, FMC – first medical contact, LMCA – stenosis of left main coronary artery, STEMI – ST-elevation myocardial infarction.

Variable		P-value
	OR (97.5% CI)	
Age [years]	0.99 (0.98–0.99)	< 0.001
Gender male	1.13 (1.01–1.27)	0.032
Weight [kg]	1.01 (1.002–1.008)	0.003
Diabetes	0.97 (0.85–1.1)	0.587
Previous stroke	1.00 (0.75–1.31)	0.993
Previous MI	0.98 (0.79–1.21)	0.862
Previous PCI	1.03 (0.83–1.27)	0.815
Previous CABG	1.42 (0.98–2.01)	0.053
Smoking status (active)	1.18 (1.07–1.31)	0.001
Psoriasis	0.84 (0.41–1.55)	0.603
Arterial hypertension	1.01 (0.92–1.11)	0.847
Kidney disease	1.22 (0.93–1.57)	0.139
COPD	1.12 (0.83–1.48)	0.438
Killip class II vs. I	1.12 (0.96–1.29)	0.156
Killip class III vs. I	1.16 (0.87–1.51)	0.300
Killip class IV vs. I	2.06 (1.66–2.55)	< 0.001
Multivessel disease	0.81 (0.74–0.89)	< 0.001
Multivessel disease with LMCA	0.63 (0.51–0.77)	< 0.001
LMCA	1.3 (0.51–2.86)	0.547
Cardiac arrest at baseline	1.03 (0.82–1.27)	0.819
СТО	0.78 (0.58–1.03	0.093
Time from pain to FMC (max 24 h) [min]	1.0 (0.99–1.0)	0.219
Time from FMC to angiography (max 24 h) [min]	0.99 (0.9987–0.9996)	0.001

Table III. Clinical facto	rs influencing the us	e of aspiration throm	bectomy in all patients
---------------------------	-----------------------	-----------------------	-------------------------

CABG – coronary artery bypass grafting, COPD – chronic obstructive pulmonary disease, COVID-19 – coronavirus disease 2019, CTO – chronic total occlusion, FMC – first medical contact, LMCA – critical stenosis of left main coronary artery, MI – myocardial infarction, PCI – percutaneous coronary intervention, STEMI – ST-elevation myocardial infarction.

1.23 (1.05-1.43)

cates that patients with SARS-COV2 infection should be treated with stents with the lowest thrombogenicity, using nanocoatings that prevent platelet adhesion [20]. The prevention of stent thrombosis is also influenced by other variables that have already been studied. Preference is given to stents made of materials that reduce adhesion and activation of platelets through albumin adsorption. Of great importance after stent implantation is the rate of endothelial coverage, which is influenced by factors such as thinner struts, type of polymer, type of drug and its quantity [21].

Our results indicate that COVID-19 infection is a strong independent predictor of a patient's eligibility for aspiration thrombectomy in the course of STEMI. Similar conclusions were described by Rodriguez-Leor *et al.*, in whose study mechanical thrombectomy was performed significantly more often in STEMI patients with confirmed SARS-COV2 virus infection compared to controls (44% vs. 33.5%, p = 0.046) [22]. In our analysis, STEMI and COVID+ patients were more likely to be both in a serious condition and to experience out-of-hospital sudden cardiac arrest. The study group of patients undergoing aspiration thrombectomy required the use of more intensive anticoagulant therapy. During PCI they were more often subjected to thrombolysis; moreover, a satisfactory effect (TIMI 3) was less often obtained than in the control group. In this work, among the predisposing factors for aspiration thrombectomy in STEMI patients enrolled in the study were COVID-19, Killip-Kimball class IV, male gender, and smoking.

0.001

The prothrombotic state described in the course of COVID-19 infection and the difference in its course compared to coagulation disorders previously known may worsen the course of ACS and require a different therapeutic approach. In the early stages of the pandemic, according to some authors, fibrinolysis may have been a reasonable alternative to STEMI treatment [23]. The authors of a large review concluded that intracoronary thrombolytic drugs significantly improved myocardial perfusion and significantly reduced the occurrence of serious adverse cardiovascular events without increasing the risk of bleeding compared to aspiration thrombectomy [24]. Polish research on the phenomenon of thrombectomy also brings important data. Siudak et al. in the NRDES study, which involved 13 hemodynamic labs, divided patients into two large groups: those who were treated with thrombectomy at the time of primary PCI and those not treated with this method. The study involved 2686 patients. As a result, thrombectomy in patients undergoing primary PCI in STEMI was not associ-

COVID-19

ated with an improvement in long-term, 1-year clinical outcome [25].

The results regarding the time from first medical contact to balloon inflation or angiogram in patients with STEMI and COVID-19 in the available literature are contradictory. Although most studies indicate that this time was longer compared to the controls [26, 27], some authors did not find significant differences in this respect [28]. In our analysis, the time from first contact to balloon inflation or angiogram was significantly longer for STEMI patients with co-infection with COVID-19 compared to controls, but the median ischemia time was similar in both groups. The reasons for this state of affairs could be e.g. the patient's fears of internal infection, operator covers, lack of barrier clothing available at the beginning of the pandemic, or longer time of patient transport to the interventional cardiology center. In the regression analysis, time from first contact to balloon inflation or angiogram had a significant but small effect on the occurrence of thrombectomy.

Summing up these reports, it seems that the hyperinflammatory response generated by COVID-19 infection strongly affects changes in the coagulation system, which can cause hypercoagulability and related complications. The fact that in patients with STEMI and COVID-19 non-obstructive coronary artery disease is more common, aspiration thrombectomy is more frequently performed, incidence of stent thrombosis is higher, and higher doses of anticoagulants are necessary raises the question of what the actual role of the coagulation and fibrinolytic systems is in the context of this disease and how this knowledge can be applied for potential risk-based benefits. It seems that in the case of patients with STEMI and COVID-19 who did not show a coronary vascular lesion of the nature of concomitant atherosclerotic plaque during PCI, thrombectomy, intravascular thrombolysis, without stent implantation, may be a promising direction of treatment, especially since stent thrombosis was more common among patients diagnosed with COVID-19. In the group with concomitant atherosclerotic lesions, the best solution seems to be the use of thinner stents, covered with a polymer with the lowest possible thrombogenicity. In addition, it remains to be determined which therapy affecting the coagulation system would be the most optimal, taking into account the fact that despite more intensive anticoagulant treatment in the group of COVID-19 STEMI patients (+), PCI treatments were less likely to result in an optimal clinical outcome (TIMI 3).

Summing up these data and current knowledge, in the course of COVID-19 and STEMI infection, in some cases a different pathophysiology of coronary vascular lesion formation should be taken into account, and the occurring coagulation disorders and shifting the balance in the procoagulation direction can be treated as a consequence of the inflammatory reaction generated in the course of SARS-CoV2 coronavirus infection. This may lead to a different therapeutic approach to this group of patients. Unlike many other studies on thrombectomy in STEMI with concomitant COVID-19, our study considered perioperative mortality (instead of in-hospital mortality), which was similar in both groups. The study presented here is the first to focus directly on examining the factors affecting eligibility for thrombectomy. When analyzing the factors influencing the eligibility for thrombectomy, we considered STEMI with a short time from the onset of pain to FMC and from FMC to angiography (< 24 h). COVID-19 independently increased the risk of thrombectomy.

### Study limitations

No techniques were used to adjust for baseline differences in patients of the 2 unequal groups such as propensity matching. Whether a patient was included in the STEMI COVID-19 (+) group was determined by the positive result of the antigen test performed in the ambulance or at the destination hospital. Swabs for molecular RT-PCR were always performed before the procedure; however, due to the urgency of the intervention, its results were not waited for. There is no database that presents these data, which is a limitation of this study.

# Conclusions

COVID-19 (+) STEMI patients undergoing aspiration thrombectomy were more often in a severe clinical condition (higher Killip-Kimball class, more often pre-procedure cardiac arrest) than COVID-19 (–) patients. Despite more intensive antiplatelet and antithrombotic treatment, PCI procedures less often resulted in an optimal TIMI 3 effect. COVID-19 is an independent strong predictor of patients' qualification for aspiration thrombectomy in STEMI.

# Conflict of interest

The authors declare no conflict of interest.

#### References

- Katsoularis I, Fonseca-Rodríguez O, Farrington P, et al. Risk of acute myocardial infarction and ischaemic stroke following COVID-19 in Sweden: a self-controlled case series and matched cohort study. Lancet 2021; 398: 599-607.
- 2. Candeloro M, Schulman S. Arterial thrombotic events in hospitalized COVID-19 patients: a short review and meta-analysis. Semin Thromb Hemost 2023; 49: 47-54.
- 3. Di Minno A, Ambrosino P, Calcaterra I, et al. COVID-19 and venous thromboembolism: a meta-analysis of literature studies. Semin Thromb Hemost 2020; 46: 763-71.
- 4. Conway EM, Mackman N, Warren RQ, et al. Understanding COVID-19-associated coagulopathy. Nat Rev Immunol 2022; 22: 639-49.
- 5. Ali EW, Ibrahim IK. Multi-factorial mechanism behind COVID-19 related thrombosis. Med Arch 2022; 76: 62-5.
- Zaremba M, Płusa A, Radowicz-Chil A. Cytotoxic T lymphocytes could contribute in COVID-19-induced passing anosmia. Medical Studies 2021; 37: 258-60.

- 7. Schuliga M, Grainge C, Westall G, Knight D. The fibrogenic actions of the coagulant and plasminogen activation systems in pulmonary fibrosis. Int J Biochem Cell Biol 2018; 97: 108-17.
- 8. Xu J, Li W, Xu S, et al. Effect of dermatan sulphate on a C57mouse model of pulmonary fibrosis. J Int Med Res 2019; 47: 2655-65.
- 9. George PM, Wells AU, Jenkins RG. Pulmonary fibrosis and COVID-19: the potential role for antifibrotic therapy. Lancet Respir Med 2020; 8: 807-15.
- 10. De Luca G, Silverio A, Verdoia M, et al. Angiographic and clinical outcome of SARS-CoV-2 positive patients with ST-segment elevation myocardial infarction undergoing primary angioplasty: a collaborative, individual patient data meta-analysis of six registry-based studies. Eur J Intern Med 2022; 105: 69-76.
- 11. De Luca G, Debel N, Cercek M, et al. Impact of SARS-CoV-2 positivity on clinical outcome among STEMI patients undergoing mechanical reperfusion: Insights from the ISACS STEMI COVID 19 registry. Atherosclerosis 2021; 332: 48-54.
- 12. Diaz-Arocutipa C, Torres-Valencia J, Saucedo-Chinchay J, Cuevas C. ST-segment elevation in patients with COVID-19: a systematic review. J Thromb Thrombolysis 2021; 52: 738-45.
- 13. Alvarez Villela MA, Alkhalil A, Weinreich MA, et al. Atypical ST-segment-elevation myocardial infarction presentation in patients with COVID-19 at a High-Volume Center in New York City. Tex Heart Inst J 2021; 48: e207446.
- 14. Quesada O, Van Hon L, Yildiz M, et al. Management strategies, and outcomes of STEMI with COVID-19: NACMI Registry. J Soc Cardiovasc Angiogr Interv 2022; 1: 100360.
- 15. Kumar S, Chawla S, Karimi H, et al. Cardiac thromboembolism in COVID-19: a case series. Cureus 2022; 14: e25193.
- 16. Green C, Nadir A, Lester W, Dosanjh D. Coronary artery thrombus resulting in ST-elevation myocardial infarction in a patient with COVID-19. BMJ Case Rep 2021; 14: e243811.
- 17. Kermani-Alghoraishi M. A review of coronary artery thrombosis: a new challenging finding in COVID-19 patients and ST-elevation myocardial infarction. Curr Probl Cardiol 2021; 46: 100744.
- Gitto M, Novelli L, Reimers B, et al. Specific characteristics of STEMI in COVID-19 patients and their practical implications. Kardiol Pol 2022; 80: 266-77.
- 19. Hamadeh A, Aldujeli A, Briedis K, et al. Characteristics and outcomes in patients presenting with COVID-19 and ST-segment elevation myocardial infarction. Am J Cardiol 2020; 131: 1-6.
- 20. Cornelissen A, Kutyna M, Cheng Q, et al. Effects of simulated COVID-19 cytokine storm on stent thrombogenicity. Cardiovasc Revasc Med 2022; 35:129-38.
- 21. Jinnouchi H, Mori H, Cheng Q, et al. Thromboresistance and functional healing in the COBRA PzF stent versus competitor DES: implications for dual antiplatelet therapy. EuroIntervention 2019; 15: e342-53.
- 22. Rodriguez-Leor O, Cid Alvarez AB, Pérez de Prado A, et al. In-hospital outcomes of COVID-19 ST-elevation myocardial infarction patients. EuroIntervention 2021; 16: 1426-33.
- 23. Wang N, Zhang M, Su H, et al. Fibrinolysis is a reasonable alternative for STEMI care during the COVID-19 pandemic. J Int Med Res 2020; 48: 300060520966151.
- 24. Kaddoura R, Mohamed Ibrahim MI, Al-Badriyeh D, et al. Intracoronary pharmacological therapy versus aspiration thrombectomy in STEMI (IPAT-STEMI): a systematic review and meta-analysis of randomized trials. PLoS One 2022; 17: e0263270.
- 25. Siudak Z, Mielecki W, Dziewierz A, et al. No long-term clinical benefit from manual aspiration thrombectomy in ST-elevation

myocardial infarction patients. Data from NRDES registry. Catheter Cardiovasc Interv 2015; 85: E16-22.

- Rodríguez-Leor O, Cid-Álvarez B, Pérez de Prado A, et al. Impact of COVID-19 on ST-segment elevation myocardial infarction care. The Spanish experience. Rev Esp Cardiol 2020; 73: 994-1002.
- 27. Chew NW, Sia CH, Wee HL, et al. Impact of the COVID-19 pandemic on door-to-balloon time for primary percutaneous coronary intervention – results from the Singapore Western STEMI Network. Circ J 2021; 85: 139-49.
- 28. Matsushita K, Hess S, Marchandot B, et al. Clinical features of patients with acute coronary syndrome during the COVID-19 pandemic. J Thromb Thrombolysis 2021; 52: 95-104.