Brief Report

The Impact of the COVID-19 Pandemic on Hospitalization **Rates due to Prosthetic Valve Thrombosis**

Mana Jameie, MD^{1,2} Mohammad Safarian Nematabad, MD³, Pejman Mansouri, MD¹, Arash Jalali, PhD^{1,2}, Faezeh Aghajani, MD⁴, Masoumeh Lotfi-Tokaldany, MD^{1,2}, Hassan Aghajani, MD, FSCAI^{1,2*}

⁴Research Development Center, Arash Women's Hospital, Tehran University of Medical Sciences, Tehran, Iran.

Received 06 December 2022; Accepted 18 March 2023

Abstract

Background: Studies have shown a decline in the admission rates of various diseases during the COVID-19 pandemic. Prosthetic valve thrombosis (PVT) is a rare condition followed by surgical or transcatheter valvular interventions. Considering the lack of data on hospitalization rates due to PVT during the pandemic, this study evaluated the implications of the COVID-19 pandemic on PVT admissions and characteristics in a tertiary referral center.

Methods: Data from all the consecutive patients hospitalized due to PVT between February 2020 and February 2021 (the first year of the pandemic) were collected from medical records and compared clinically with the corresponding time before the pandemic (February 2019 through February 2020). Variables of interest included the number of hospitalization, patient and valve characteristics, diagnostic and management strategies, and in-hospital events.

Results: Forty patients (32.5% male, age: 54.0 [46.5-62.0 y] comprised the study population. We observed a considerable decline in hospitalization rates during the pandemic, from 31 to 9 patients. Admitted patients were 8 years younger, had a higher proportion of the New York Heart Association functional class III or IV symptoms (44.4% vs 22.6%), were more often treated with fibrinolysis (33.3% vs 22.6%) or surgical approaches (33.3% vs 22.6%), and were discharged 6 days sooner.

Conclusion: We described a reduction in PVT hospitalization. Patients presented with a higher proportion of severe dyspnea and had increased treatment with fibrinolysis/surgical approaches. These observations highlight the necessity of the active surveillance of patients with prosthetic valves by caregivers for timely diagnosis and appropriate management during the pandemic.

J Teh Univ Heart Ctr 2023;18(2):136-141

This paper should be cited as: Jameie M, Safarian Nematabad M, Mansouri P, Jalali A, Aghajani F, Lotfi-Tokaldany M, et al. The Impact of the COVID-19 Pandemic on Hospitalization Rates due to Prosthetic Valve Thrombosis. J Teh Univ Heart Ctr 2023;18(2):136-141.

Keywords: Heart valve prosthesis; Heart valve diseases; COVID-19; Hospitalization

*Corresponding Author: Hassan Aghajani, Associate Professor of Interventional Cardiology, Department of Interventional Cardiology, Tehran University of Medical Sciences, Tehran Heart Center, North Kargar Street, Tehran, Iran. 1411713138. Tel: +98 21 88029256. Fax: +98 21 88029256. E-mail: aghajanih@sina.tums.ac.ir & aghajanihas@yahoo.com.

Copyright © 2023 Tehran University of Medical Sciences. Published by Tehran University of Medical Sciences.

136 -

10 (D) This work is licensed under a Creative Commons Attribution-NonCommercial 4.0 International license (https://creativecommons.org/licenses/by-nc/4.0/). Noncommercial uses of the work are permitted, provided the original work is properly cited.

¹Tehran Heart Center, Cardiovascular Disease Research Institute, Tehran University of Medical Sciences, Tehran Iran

²Cardiac Primary Prevention Research Center, Cardiovascular Disease Research Institute, Tehran University of Medical Sciences, Tehran, Iran.

³Ali IbnAbitaleb Educational and Treatment Hospital, Rafsanjan University of Medical Sciences, Rafsanjan, Iran.

Introduction

Prosthetic valve thrombosis (PVT) is a rare, albeit serious, complication of surgical or transcatheter valve replacement. It most frequently affects mechanical and, less often, bioprosthetic valves and is associated with higher mortality and morbidity.¹ Pathologically, valve thrombosis is characterized by thrombus formation on valve-related structures, with the potential ability to result in subsequent thromboembolism. Consequently, failure to diagnose and treat PVT promptly can lead to limb ischemia,² pulmonary thromboembolism, and disabling/fatal cerebrovascular accidents.³

The announcement of the COVID-19 outbreak⁴ adversely affected the health of millions of people. The pandemic has significantly decreased visits to care centers for a wide range of conditions,^{5, 6} including cardiovascular disease.⁷⁻⁹ Studies have suggested the fear of contamination as a culprit for this phenomenon.¹⁰ Cardiovascular comorbidities are particularly significant since they are known risk factors for the severe clinical presentation of COVID-19,11 causing a greater dread and delay for patients with these comorbidities to present to hospitals. Studies have described a decline in admission due to acute coronary syndromes,^{8,9} while the reported rate of out-of-hospital cardiac arrests has risen.¹² This simultaneity hints at a possible unmet burden of cardiovascular disease during the pandemic, concealed because of the COVID-19 restrictions and fear of contracting the disease at hospitals and clinics.

Given a lack of data on the impact of the COVID-19 pandemic on PVT-induced hospitalization, the present study evaluated hospitalization rates, patient and valve characteristics, management strategies, and in-hospital events among individuals with PVT before and during the pandemic.

Methods

This case series was conducted at Tehran Heart Center, a cardiovascular disease-dedicated hospital affiliated with the Tehran University of Medical Sciences.¹³ All consecutive patients hospitalized due to PVT in the first year of the COVID-19 outbreak (from February 21, 2020, through February 18, 2021) were included in the During-COVID group and were compared with all consecutive patients hospitalized due to PVT (the Pre-COVID group) during the corresponding period before the pandemic (from February 20, 2019, through February 19, 2020). There were no exclusion criteria. Patients either presented to the emergency department or were referred from the outpatient valve or other clinics. The study was approved by the local ethics committee and conformed to the Helsinki Declaration (IR.TUMS.THC.REC.1400.044). Due to the retrospective design and anonymization of the data, the need for informed consent was waived. Data, including hospitalization rates, patient and valve characteristics, management strategies, and in-hospital events, were collected from electronic and paper medical records.

Categorical variables were presented as numbers and frequencies and compared using the χ^2 or Fisher exact test as appropriate. Continuous variables were presented as medians and interquartile ranges and compared using the nonparametric Mann-Whitney *U* Test. IBM SPSS Statistics for Windows, version 21.0, was used for data analysis. Two-sided probability values were reported. Nevertheless, due to the small sample size (ie, lack of power), we relied on the clinical significance of our findings for data interpretation and not the statistical significance.

Results

Forty patients (32.5% male, age: 54.0 [46.5-62.0 y] comprised the study population. We identified 9 PVT patients during the pandemic compared with 31 patients before it. Table 1 compares the characteristics of the During-COVID and Pre-COVID groups. (Only clinically relevant differences are discussed hereafter.) Sex distribution did not differ between the 2 periods, with PVT affecting women almost twice as much as men. The During-COVID group patients were 8 years younger than the Pre-COVID group patients (50.0 [77.0-51.0 y] vs 58.0 [48.5-63.0 y]). The proportion of patients presenting with the New York Heart Association function class (NYHA FC) III/IV increased considerably (44.4% vs 22.6%). There was also an increase in patients with an international normalized ratio (INR) of less than 2.5 (77.8% vs 58.1%). The isolated mitral valve and simultaneous mitral valve and aortic valve involvement decreased (48.4% vs 22.2% and 25.8% vs 11.1%, respectively), while that of isolated aortic valve increased (44.4% vs 22.6%). The use of transesophageal echocardiography (TEE) as the first-line diagnostic imaging technique dropped from 45.2% to 11.1%. In contrast, the proportion of patients who underwent fluoroscopy and subsequent TEE (due to abnormal fluoroscopic findings) rose from 51.6% to 66.7%. In the During-COVID group, treatment with anticoagulants was lower by 21.5%, while fibrinolysis and surgical technique were each higher by 10.7%. Further, the hospital stay was shortened from 14.0 [9.5-17.0 d] to 8.0 [6.0-19.0 d].

Discussion

Figure 1 illustrates the principal findings of the current study. We found a considerable reduction in the number of patients with PVT admitted to our center during the

```
http://jthc.tums.ac.ir
```

Table 1. Patient characteristics*

	The Pre-COVID Group (n=31, 77.5%)	The During-COVID Group (n=9, 22.5%)	Р
Baseline Characteristics			
Demographics			
Age	58.0 [48.5-63.0]	50.0 [44.0-51.0]	0.024
Male	10 (32.3%)	3 (33.3%)	>0.999
Clinical Presentations			
NYHA FC III / IV	7 (22.6%)	4 (44.4%)	0.227
Unstable hemodynamic	2 (6.5%)	0	>0.999
AF rhythm at admission	9 (29.0%)	2 (22.2%)	>0.999
Paraclinical Evaluations			
LVEF	50.0 [42.5-50.0]	50 [45.0-50.0]	0.323
INR	2.2 [1.5-2.9]	2.1 [1.3-2.4]	0.759
INR Range			0.337
<2.5	18 (58.1%)	7 (77.8%)	
2.5-3.5	7 (22.6%)	2 (22.2%)	
>3.5	6 (19.4%)	0	
VTH Characteristics			
Valve Type			0.689
Mechanical	27 (87.1%)	8 (88.9%)	
Bioprosthetic	4 (12.9%)	1 (11.1%)	
Valve Site			0.120
Isolated MV	15 (48.4%)	2 (22.2%)	
Isolated AV	7 (22.6%)	4 (44.4%)	
MV-AV	8 (25.8%)	1 (11.1%)	
Isolated TV	1 (3.2%)	1 (11.1%)	
Isolated PV	0	1 (11.1%)	
Diagnostic Method			0.080
First-line TTE #	1 (3.2%)	1 (11.1%)	
First-line TEE #	14 (45.2%)	1 (11.1%)	
Fluoroscopy-TEE	16 (51.6%)	6 (66.7%)	
Fluoroscopy-TTE	0	1 (11.1%)	
Type of Treatment			0.525
Anticoagulant	17 (54.8%)	3 (33.3%)	
Fibrinolytic	7 (22.6%)	3 (33.3%)	
Surgery	7 (22.6%)	3 (33.3%)	
Type of Surgery (n=10)			0.500
Valve replacement	2 (28.6%)	2 (66.7%)	
Surgical thrombectomy	5 (71.4%)	1 (33.3%)	
In-hospital Events			
LOS	14.0 [9.5-17.0]	8.0 [6.0-19.0]	0.211
mortality	2 (6.5%)	0	>0.999
CVA	1 (3.2%)	0	>0.999

*Data are presented as frequencies (percentages) or medians (Q1-Q3).

+ First-line refers to performing the mentioned diagnostic imaging method as the first step and without preceding fluoroscopy.

NYHA, New York Heart Association; FC, Functional class; AF, Atrial fibrillation; LVEF, Left-ventricular ejection fraction; INR, International normalized ratio; VHT, Valvular heart thrombosis; MV, Mitral valve; AV, Aortic valve; TV, Tricuspid valve; PV, Pulmonary valve; TTE, Transthoracic echocardiography; TEE, Transesophageal echocardiography; LOS, Length of stay; CVA, Cerebrovascular accident



Figure 1. The image depicts the impact of COVID-19 on hospitalization rates due to prosthetic valve thrombosis. NHYA, New York Heart Association; TEE, Transesophageal echocardiography; LOS, Length of stay

COVID-19 pandemic. The During-COVID group patients were 8 years younger, had more frequent presentations with severe dyspnea and treatment with fibrinolytic/surgical therapies, and had shorter hospital stays than the Pre-COVID group patients.

The impact of the COVID-19 pandemic on various cardiovascular conditions has been previously reported.^{7.9, 14, 15} However, information on admissions due to PVT is missing. Chiming with our observation, a recent systematic review reported a 40-50% reduction in acute coronary syndromes and a 12-40% reduction in stroke admissions during the pandemic.¹⁶ Likewise, a large multisite study reported a significant reduction in admission rates due to various cardiovascular conditions, one of which was "other selected acute cardiovascular conditions" including prosthetic valve-related diseases.¹⁷

Multiple explanations can be proposed for the reduced hospitalization, the most worrisome of which is that patients, particularly those with milder symptoms, were more reluctant to seek medical care due to the fear of contracting the disease,¹¹ which is also suggested by others.¹⁰ Nonetheless, it is difficult to evaluate the pure role of viral fear without considering its intricate interaction with COVID-19 restriction rules and media effects, which might have heightened this effect.¹⁸ Moreover, dyspnea as a common symptom of PVT might be mistaken for COVID-19 symptoms by both patients and physicians, thereby causing delays in further evaluations for PVT. Additionally, patients might have presented to COVID-19 dedicated centers due to dyspnea; thus, their PVT diagnosis might have been made elsewhere rather than at our center. Still, we continued to encourage our patients with valvular interventions to come to our center if they had any symptoms.

Patients were considerably younger during the pandemic, possibly reflecting that older people are more frightened to present to the hospital. Some studies have reported the younger age of admitted patients with acute coronary syndromes, as well,⁸ while others have not.¹⁹ The behavior of medical care avoidance might have more severe implications among older patients, hence the importance of encouraging this group not to ignore their symptoms during the ongoing pandemic.

As also noted with other cardiovascular conditions,^{7, 8, 19, 20, 21} the more frequent presentation with severe symptoms (eg, dyspnea in our study) during the pandemic might imply that patients with milder symptoms were more inclined to stay home and present to hospitals when their symptoms had become critical. The increase in the use of fibrinolytic and surgical treatments, which are preferred for patients with more severe symptoms and higher clot burdens, can support this claim.

During the pandemic, the proportion of patients undergoing first-line TEE dramatically decreased, possibly implying an enhanced tendency toward the use of fluoroscopy over TEE as the initial diagnostic technique. This is in line with the latest guidelines on cardiac imaging during the ongoing pandemic. Due to the high risk of staff contamination, TEE should be avoided as much as possible and be limited to occasions when there are crucially suspicious findings or when it might change the treatment approach.²² The decline in hospitalization length has been a worldwide issue since the start of the pandemic,^{14, 16, 17, 23} representing physicians' and patients' inclination toward an early discharge, as well as hospitals' endeavors to ensure bed availability.

INR changes could indicate that patients went to outpatient prothrombin time clinics or drugstores less frequently and more irregularly, as proposed by other researchers, as well.²⁴ Many people taking anticoagulants are older and have comorbidities; therefore, they may be more afraid of contracting the disease and more likely to abide by the country's "stay home" instructions. These behavioral changes can make patients liable to over-or

```
http://jthc.tums.ac.ir
```

under-anticoagulation.25

The observational and single-center nature of the current study precluded the possibility of finding a causal relationship between the pandemic and PVT admissions. Our findings are, accordingly, only hypothesis-generating and should be confirmed by other studies. Most importantly, as a consequence of the low PVT incidence and our single-center design, there was a lack of power to detect statistically significant differences. Consequently, we were able to discuss only clinically-relevant findings. It should be noted that many variables for which we reported a clinical difference had significant or borderline P values. Therefore, larger multicenter studies could further draw statistically significant differences in this regard. Our study is the first to report the effects of the COVID-19 pandemic on PVT admissions so far, which we hope will serve as a trigger for further multicenter research in the field.

Conclusion

Hospitalization due to PVT dropped considerably during the COVID-19 pandemic. Admitted patients were 8 years younger, with a higher proportion of severe dyspnea and treatment with fibrinolysis/surgical approaches. These changes highlight the significance of the active surveillance of patients with prosthetic valves to ensure that they are symptom-free, taking their anticoagulant medications properly, and having scheduled follow-up visits promptly during the pandemic.

References

- Dangas GD, Weitz JI, Giustino G, Makkar R, Mehran R. Prosthetic Heart Valve Thrombosis. J Am Coll Cardiol 2016;68:2670-2689.
- Barandon L, Clerc P, Chauvel C, Plagnol P. Native aortic valve thrombosis: a rare cause of acute ischemia of the lower limb. Interact Cardiovasc Thorac Surg 2004;3:675-677.
- Schoen FJ, Gotlieb AI. Heart valve health, disease, replacement, and repair: a 25-year cardiovascular pathology perspective. Cardiovasc Pathol 2016;25:341-352.
- WHO COVID-19 Dashboard. Geneva: World Health Organization. 2020 [Available from: https://covid19.who.int/.)Access date:1 March 2023)
- McAndrew J, O'Leary J, Cotter D, Cannon M, MacHale S, Murphy KC, Barry H. Impact of initial COVID-19 restrictions on psychiatry presentations to the emergency department of a large academic teaching hospital. Ir J Psychol Med 2021;38:108-115.
- Ferrero F, Ossorio MF, Torres FA, Debaisi G. Impact of the COVID-19 pandemic in the paediatric emergency department attendances in Argentina. Arch Dis Child 2021;106:e5.
- Bromage DI, Cannatà A, Rind IA, Gregorio C, Piper S, Shah AM, McDonagh TA. The impact of COVID-19 on heart failure hospitalization and management: report from a Heart Failure Unit in London during the peak of the pandemic. Eur J Heart Fail 2020;22:978-984.
- 8. Gluckman TJ, Wilson MA, Chiu ST, Penny BW, Chepuri VB, Waggoner JW, Spinelli KJ. Case Rates, Treatment Approaches, and

Outcomes in Acute Myocardial Infarction During the Coronavirus Disease 2019 Pandemic. JAMA Cardiol 2020;5:1419-1424.

- Solomon MD, McNulty EJ, Rana JS, Leong TK, Lee C, Sung SH, Ambrosy AP, Sidney S, Go AS. The Covid-19 Pandemic and the Incidence of Acute Myocardial Infarction. N Engl J Med 2020;383:691-693.
- Barten DG, Latten GHP, van Osch FHM. Reduced Emergency Department Utilization During the Early Phase of the COVID-19 Pandemic: Viral Fear or Lockdown Effect? Disaster Med Public Health Prep 2022;16:36-39.
- Garrafa E, Levaggi R, Miniaci R, Paolillo C. When fear backfires: Emergency department accesses during the Covid-19 pandemic. Health Policy 2020;124:1333-1339.
- Chen T, Wu D, Chen H, Yan W, Yang D, Chen G, Ma K, Xu D, Yu H, Wang H, Wang T, Guo W, Chen J, Ding C, Zhang X, Huang J, Han M, Li S, Luo X, Zhao J, Ning Q. Clinical characteristics of 113 deceased patients with coronavirus disease 2019: retrospective study. BMJ. 2020 Mar 26;368:m1091. doi: 10.1136/bmj.m1091. Erratum in: BMJ 2020;368:m1295.
- Poorhosseini H, Abbasi SH. The Tehran heart center, Eur Heart J 2018;39:2695–2696
- Page EM, Ariëns RAS. Mechanisms of thrombosis and cardiovascular complications in COVID-19. Thromb Res 2021;200:1-8.
- 15. Roudaut R, Serri K, Lafitte S. Thrombosis of prosthetic heart valves: diagnosis and therapeutic considerations. Heart 2007;93:137-142.
- 16. Cannatà A, Bromage DI, Rind IA, Gregorio C, Bannister C, Albarjas M, Piper S, Shah AM, McDonagh TA. Temporal trends in decompensated heart failure and outcomes during COVID-19: a multisite report from heart failure referral centres in London. Eur J Heart Fail 2020;22:2219-2224.
- Birkmeyer JD, Barnato A, Birkmeyer N, Bessler R, Skinner J. The Impact Of The COVID-19 Pandemic On Hospital Admissions In The United States. Health Aff (Millwood) 2020;39:2010-2017.
- Kiss P, Carcel C, Hockham C, Peters SAE. The impact of the COVID-19 pandemic on the care and management of patients with acute cardiovascular disease: a systematic review. Eur Heart J Qual Care Clin Outcomes 2021;7:18-27.
- Ghoreishi A, Arsang-Jang S, Sabaa-Ayoun Z, Yassi N, Sylaja PN, Akbari Y, Divani AA, Biller J, Phan T, Steinwender S, Silver B, Zand R, Basri HB, Iqbal OM, Ranta A, Ruland S, Macri E, Ma H, Nguyen TN, Abootalebi S, Gupta A, Alet M, Lattanzi S, Desai M, Gagliardi RJ, Girotra T, Inoue M, Yoshimoto T, Isaac CF, Mayer SA, Morovatdar N, Nilanont Y, Nobleza COS, Saber H, Kamenova S, Kondybayeva A, Krupinski J, Siegler JE, Stranges S, Torbey MT, Yorio D, Zurrú MC, Rubinos CA, Shahripour RB, Borhani-Haghighi A, Napoli MD, Azarpazhooh MR. Stroke Care Trends During COVID-19 Pandemic in Zanjan Province, Iran. From the CASCADE Initiative: Statistical Analysis Plan and Preliminary Results. J Stroke Cerebrovasc Dis 2020;29:105321.
- Scognamiglio G, Fusco F, Merola A, Palma M, Correra A, Sarubbi B. Caring for adults with CHD in the era of coronavirus disease 2019 pandemic: early experience in an Italian tertiary centre. Cardiol Young 2020;30:1405-1408.
- 21. Firouzi A, Hosseini Z, Norouzi Z, Hosseini Z, Amirpour A, Talakoob H, Amin A, Soleimani A, Moradifar N, Karbalai S, Mozafarybazargani M, Hekmat H, Maleki M, Sadeghipour P, Mirbod SM, Ghorbanpoor Kohnaki M, Bakhshandeh H, KalaeiNia M, Habibizade FS, Iraninejad S, Baay M, Khalilipur E. The Pragmatic Role of COVID-19 on the Thrombus Grade of Patients with Contemporary ST-Segment-Elevation Myocardial Infarction. J Tehran Heart Cent 2022;17:103-111.
- 22. De Rosa S, Spaccarotella C, Basso C, Calabrò MP, Curcio A, Filardi PP, Mancone M, Mercuro G, Muscoli S, Nodari S, Pedrinelli R, Sinagra G, Indolfi C; Società Italiana di Cardiologia and the CCU Academy investigators group. Reduction of hospitalizations for myocardial infarction in Italy in the COVID-19 era. Eur Heart J 2020;41:2083-2088.

- 23. Mohsenizadeh SA, Alidoosti M, Jalali A, Tofighi S, Salarifar M, Poorhosseini H, Jenab Y, Ahmadian T. Comparison of Angiographic and Clinical Outcomes After Primary Percutaneous Coronary Intervention for ST-elevation Myocardial Infarction Between Patients With and Without Concomitant COVID-19 Infection. Crit Pathw Cardiol 2022;21:141-146
- Omidi N, Forouzannia SK, Poorhosseini H, Tafti SHA, Salehbeigi S, Lotfi-Tokaldany M. Prosthetic heart valves and the COVID-19 pandemic era: What should we be concerned about? J Card Surg 2020;35:2500-2505.
- Clerkin KJ, Fried JA, Raikhelkar J, Sayer G, Griffin JM, Masoumi A, Jain SS, Burkhoff D, Kumaraiah D, Rabbani L, Schwartz A, Uriel N. COVID-19 and Cardiovascular Disease. Circulation 2020;141:1648-1655.

The Journal of Tehran University Heart Center 141