## Anticoagulation for left ventricular thrombi secondary to COVID—Is 3 months too long?

Revised: 8 May 2022

Rimmy Garg<sup>1,2</sup> Amitoj Sachdeva<sup>1</sup> | Juan Del Cid Fratti<sup>1,2</sup> | Samuel Mortoti<sup>2</sup>

<sup>1</sup>University of Illinois College of Medicine Peoria, Peoria, Illinois, USA

<sup>2</sup>OSF Healthcare Cardiovascular Institute, Peoria, Illinois, USA

Correspondence

Rimmy Garg, 6710 N. Summershade Circle Apt H, Peoria, IL 61615, USA. Email: rimmy.p.garg@gmail.com

## Abstract

The length of anticoagulation for thrombotic events related to COVID-19 is unknown. We present a patient with COVID-19 complicated by a thrombotic anterior STEMI and multiple left ventricular (LV) thrombi that resolved after 8 weeks of anticoagulation. We suggest a shorter length of anticoagulation with COVID-19-related LV thrombus.

**KEYWORDS** anticoagulation, COVID, left ventricle thrombi, STEMI

#### **INTRODUCTION** 1

COVID-19 has been associated with venous and arterial thromboembolic disease likely due to potent local and systemic cytokine production with subsequent platelet activation, thrombin stimulation, and fibrin deposition.<sup>1</sup> Such thromboembolic disease is treated with anticoagulation, but there are no guidelines to direct the types and duration of oral anticoagulation (OAC). We present a case of COVID-associated acute coronary syndrome (ACS) and left ventricular (LV) thrombi with resolution of thrombi within 2 months of warfarin initiation.

#### 2 **CASE PRESENTATION**

A 62-year-old female patient with a past medical history of hypertension, hyperlipidemia, and tobacco use presented with left-sided chest pain with radiation to the left arm that started the night prior to admission. She was recently diagnosed with mild COVID-19 infection 2 weeks earlier and was treated conservatively as an outpatient. Neither she nor her family has any history of coronary artery

disease, heart failure, or any arrythmias. She was found to be tachycardic but with a regular rhythm and an otherwise normal physical examination.

Laboratory data were notable for an elevated troponin of 31.2 ng/ml (reference: <0.028 ng/ml) and elevated aspartate aminotransferase of 177 U/L. Electrolytes and an arterial blood gas values were within normal range.

ECG demonstrated ST elevations in leads II, III, aVF, and V3-V5 with ST depression in aVL (Figure 1). With a diagnosis of anterior and inferior myocardial infarction, patient was taken emergently for a left heart catheterization (LHC).

Left heart catheterization showed a thrombotic occlusion of the proximal subsection of the distal left anterior descending (LAD) coronary artery with evidence of organized thrombus, judged by the difficulty in passing a wire across (Figure 2A). Multiple balloon dilatations and multiple rounds of aspiration with a penumbra catheter were attempted, and intracoronary eptifibatide was administered with the restoration of TIMI 2 flow (Figure 2B, Video S2). Given the presence of organized clot, decision was to treat medically. The patient otherwise had nonobstructive disease of the other coronaries. A LHC was

This is an open access article under the terms of the Creative Commons Attribution License, which permits use, distribution and reproduction in any medium, provided the original work is properly cited.

© 2022 The Authors. Clinical Case Reports published by John Wiley & Sons Ltd.

 $\mathbf{v}$ \_Clinical Case Reports \_

repeated 2 days later to see whether the thrombus had resolved and stent could be placed. However, there was still residual thrombus in the distal LAD, unchanged from prior study (Figure 2C). As the patient was chest pain free and hemodynamically stable, no further intervention was attempted. A transthoracic echocardiogram (TTE) was performed, demonstrating apical akinesis with LV ejection fraction of 39% by Simpson's biplane method as well as multiple large, mobile LV thrombi with a maximum size of  $2 \text{ cm} \times 1.5 \text{ cm}$  (Figure 3A,B, Videos S3–S9). With systolic dysfunction in the setting of STEMI, the patient was maintained on aspirin, ticagrelor, atorvastatin, metoprolol succinate, spironolactone, and losartan. She was additionally started on a heparin drip with bridge to warfarin in the setting of multiple large LV thrombi. Given the size and number of thrombi and associated increased risk of stroke, cardiac surgery was consulted for potential surgical LV thrombus evacuation. Surgical intervention was not recommended due to high risk of complications in the setting of recent ACS, and plan was to continue medical management.



**FIGURE 1** EKG demonstrating ST elevations in leads II, III, avF, and V3–V5 with ST depression in aVL



**FIGURE 2** (A) Thrombotic occlusion of the proximal subsection of the distal LAD with evidence of organized thrombus. (B) Occluded LAD after percutaneous coronary intervention (PCI) with restoration of TIMI 2 flow. (C) Persistently occluded LAD 2 days after PCI. LAD, Left anterior descending



**FIGURE 3** (A) TTE without contrast at the off-axis apical 4 chamber view demonstrating multiple LV thrombi. (B) TTE with contrast at the off-axis apical 4 chamber view demonstrating multiple LV thrombi. TTE, Transthoracic echocardiogram; LV, left ventricular



**FIGURE 4** TTE without contrast at the apical 4 chamber view shows resolution of LV thrombi. TTE, Transthoracic echocardiogram; LV, left ventricular

It was thought that the LAD and LV thrombi were secondary to the patient's recent COVID infection. At the time of discharge, aspirin was discontinued. Ticagrelor was to be continued for a year and warfarin for at least 3 months based on the resolution of thrombi.

Our patient was followed up by a cardiologist and had a repeat TTE 2 months after hospital discharge. This TTE \_Clinical Case Reports

-WILEY

showed that there was a large apical aneurysm of the LV but no evidence of any thrombi in the apex (Figure 4, Videos S10– S13). The ejection fraction was still low around 35%. Warfarin was continued at this time due to the lack of contrast with the last TTE study (patient had refused contrast at that time), but discontinued 3.5 months later when a repeat TTE with contrast showed that the LV remained unchanged with no evidence of thrombi (Figure 5, Videos S14–S16). The patient was subsequently restarted on aspirin 81 mg daily for dual antiplatelet therapy with her ticagrelor, and she remained asymptomatic on the following clinic visits.

# 3 | DISCUSSION AND CONCLUSION

Cardiac manifestations of COVID-19 ACS are due to coronary thrombosis or acute plaque rupture from systemic inflammation and catecholamine surge.<sup>2,3</sup> For ACS due to plaque rupture, it is recommended that dual anti-platelet therapy and full-dose anticoagulation be administered in the acute setting per the American College of Cardiology (ACC)/ American Heart Association (AHA) and the European Society of Cardiology (ESC) guidelines.<sup>4</sup> There are currently no specific guidelines on the anticoagulant used or the duration of anticoagulation in the setting of coronary thrombosis and/or LV thrombosis related to COVID-19.

As LV thrombus is not an uncommon complication of an acute myocardial infarction, the 2013 ACC/AHA STEMI guidelines recommend OAC use in the setting of STEMI with anterior apical akinesis or dyskinesis to prevent the thrombus formation for 3 months, aiming for a lower international normalized ratio of 2.0–2.5.<sup>5</sup> The 2017 ESC STEMI guidelines recommend OAC for up to 6 months with final duration guided by a repeat TTE, risk of bleeding, and need for concomitant antiplatelet



**FIGURE 5** TTE with contrast at the apical 4 chamber view shows resolution of LV thrombi. TTE, Transthoracic echocardiogram; LV, left ventricular

-WILEY-Clinical Case Reports -

therapy.<sup>6</sup> The ACTION Study Group also concluded that anticoagulation for the LV thrombus for at least 3 months was associated with a lower risk of major cardiovascular events or all-cause mortality.<sup>7</sup>

There have not been any large prospective or direct comparison studies looking at direct OAC (DOAC) versus warfarin for the treatment of LV thrombus. One metanalysis did show that DOACs appear to be non-inferior to warfarin without any statistical difference in stroke or bleeding complications when treating for the LV thrombus.<sup>8</sup>

There have been cases that reported resolution of the LV thrombus in the setting of COVID-19 infection prior to the 3-month mark. One reported resolution on a 1-month follow-up TTE while on warfarin<sup>9</sup> while another reported resolution at 10 days on low molecular weight heparin.<sup>10</sup> Our patient had a large LV thrombus burden that resolved within 2 months with warfarin use.

Additionally, the risk of an LV thrombus is the highest in the first month after a myocardial infarction. In patients with a chronic LV aneurysm, the incidence of systemic emboli is extremely low, and thus, the use of long-term OAC is not justified.<sup>11</sup>

As such, a shorter duration of anticoagulation under close supervision should be considered in patients with COVID-19-related cardioembolic/thrombotic events, guided by echocardiographic imaging. Such imaging would require a thorough sweep of the left ventricle in on-axis and off-axis views so as not to miss any residual thrombi. Prospective data about proper OAC regimen and duration are still needed.

#### AUTHOR CONTRIBUTIONS

All authors contributed equally to the preparation of this manuscript, including literature review, writing, acquiring images, and final review. All authors have read and approved the final manuscript.

#### ACKNOWLEDGEMENT

None.

### **CONFLICT OF INTEREST**

The authors of this manuscript have no competing financial or non-financial competing interests.

#### ETHICAL APPROVAL

As this was a case report with de-identified patient information, an IRB approval was not needed.

#### CONSENT

The patient from this case graciously consented to the publication of this manuscript.

ORCID Rimmy Garg b https://orcid.org/0000-0003-1657-6250

#### REFERENCES

- Ortega-Paz L, Capodanno D, Montalescot G, Angiolillo DJ. Coronavirus disease 2019 – associated thrombosis and coagulopathy: review of the pathophysiological characteristics and implications for antithrombotic management. J Am Heart Assoc. 2021;10(3):e019650.
- 2. Xiong TY, Redwood S, Prendergast B, Chen M. Coronaviruses and the cardiovascular system: acute and long-term implications. *Eur Heart J.* 2020;41(19):1798-1800.
- 3. Dominguez-Erquicia P, Dobarro D, Raposeiras-Roubín S, Bastos-Fernandez G, Iñiguez-Romo A. Multivessel coronary thrombosis in a patient with COVID-19 pneumonia. *Eur Heart J*. 2020;41(22):2132.
- 4. Bikdeli B, Madhavan M, Jimenez D, et al. COVID-19 and thrombotic or thromboembolic disease: implications for prevention, antithrombotic therapy, and follow-up: JACC state-of-the-art review. *J Am Coll Cardiol*. 2020;75(23):2950-2973.
- O'Gara PT, Kushner FG, Ascheim DD, et al. 2013 ACCF/AHA guideline for the management of ST-elevation myocardial infarction: a report of the American College of Cardiology Foundation/American Heart Association task force on practice guidelines. *Circulation*. 2013;127(4):362-425.
- 6. Ibanez B, James S, Agewall S, et al. 2017 ESC guidelines for the management of acute myocardial infarction in patients presenting with ST-segment elevation: the task force for the management of acute myocardial infarction in patients presenting with ST-segment elevation of the European Society of Cardiology (ESC). *Eur Heart J.* 2018;39(2):119-177.
- Lattuca B, Bouziri N, Kerneis M, et al. Antithrombotic therapy for patients with left ventricular mural thrombus. *J Am Coll Cardiol.* 2020;75:1676-1685.
- Dalia T, Lahan S, Ranka S, et al. Warfarin versus direct oral anticoagulants for treating left ventricular thrombus: a systematic review and meta-analysis. *Thrombosis J.* 20211;19:7. doi:10.1186/s12959-021-00259-w
- Jeon HD, Patel J, Priester T, Moten M. Left ventricular thrombus caused by COVID-19: how long to treat? *J Am Coll Cardiol*. 2021;77(18\_Supplement\_1):2013.
- Sonaglioni A, Albini A, Nicolosi GL, Rigamonti E, Noonan DM, Lombardo M. Case report: an unusual case of biventricular thrombosis in a COVID-19 patient with ischemic dilated cardiomyopathy: assessment of mass mobility and embolic risk by tissue doppler imaging. *Front Cardiovasc Med.* 2021;29(8):694542.
- Lapeyre AC 3rd, Steele PM, Kazmier FJ, Chesebro JH, Vlietstra RE, Fuster V. Systemic embolism in chronic left ventricular aneurysm: incidence and the role of anticoagulation. *J Am Coll Cardiol.* 1985;6(3):534-538.

### SUPPORTING INFORMATION

Additional supporting information may be found in the online version of the article at the publisher's website.

How to cite this article: Garg R, Sachdeva A, Del Cid Fratti J, Mortoti S. Anticoagulation for left ventricular thrombi secondary to COVID—Is 3 months too long?. *Clin Case Rep.* 2022;10:e05950. doi: 10.1002/ccr3.5950

4 of 4