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## 7 **Para-cardiac Inflammatory Mass Compressing the Heart**

### 8 *A possible association with COVID-19*

9 **Ahmad Mirdamadi,<sup>1</sup> Amir M.M. Sadeghi,<sup>2</sup> Mohsen M.M. Sadeghi,<sup>3</sup> Majid**  
10 **Rabani,<sup>4</sup> \*Shakiba Hassanzadeh<sup>5</sup>**

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12 *<sup>1</sup>Department of Cardiology, Medical School, Islamic Azad University, Najafabad Branch,*  
13 *Isfahan, Iran; <sup>2</sup>Department of Cardiovascular Surgery, Isfahan University of Medical Sciences,*  
14 *Isfahan, Iran; <sup>3</sup>Department of Cardiac Surgery, Chamran Heart Center, Isfahan University of*  
15 *Medical Sciences, Isfahan, Iran; <sup>4</sup>Cardiac Ward, Khanévadeh Hospital, Isfahan, Iran; <sup>5</sup>Division*  
16 *of Pathology, Nickan Research Institute, Isfahan, Iran*

17 *\*Corresponding Author's e-mail: [shak.hassanzadeh@gmail.com](mailto:shak.hassanzadeh@gmail.com)*

#### 19 **Abstract**

20 Infection with the SARS-CoV-2 virus causes coronavirus disease 2019 (COVID-19). COVID-19  
21 usually affects the lungs but may also involve other organs such as the heart. We report a case of  
22 a para-cardiac mass in a previously healthy 45-year-old man who developed persistent dyspnea  
23 following SARS-CoV-2 infection. The patient underwent cardiac surgery since the mass was  
24 attached to the pericardium and was causing constrictive pericarditis. The pathology report  
25 indicated an inflammatory pattern for the mass. Based on our knowledge there has been no  
26 previous report of developing a para-cardiac inflammatory mass after SARS-CoV-2 infection. In  
27 conclusion, we would like to increase awareness regarding the possibility of developing a para-  
28 cardiac inflammatory mass following COVID-19.

29 **Keywords:** SARS-CoV-2; Pericarditis; Constrictive pericarditis; COVID-19; Cardiac tumor;  
30 Mediastinal tumor

31

## 32 **Introduction**

33 Severe acute respiratory syndrome coronavirus 2 (SARS-CoV-2) is caused by the SARS-CoV-2  
34 virus. The most common clinical manifestations of SARS-CoV-2 infection are respiratory  
35 symptoms; however, this infection may also involve other organs such as the heart and kidneys.<sup>1</sup>  
36 Some of the most common cardiac complications following coronavirus disease 2019 (COVID-  
37 19) include myocarditis, pericardial effusion and tamponade, myocardial infarction (MI),  
38 arrhythmias, acute heart failure (HF) with cardiogenic shock, and pericarditis. Most of these  
39 patients have shortness of breath and chest pain.<sup>1</sup> On the other hand, primary cardiac tumors are  
40 very rare and have an incidence of 0.001%–0.03% on autopsy findings and about 90% of them  
41 are benign.<sup>2</sup> In addition, 6 to 10% of primary cardiac tumors are primary pericardial masses  
42 which are also usually benign. The clinical signs and symptoms of pericardial masses include  
43 dyspnea, edema, pleural effusion, orthopnea, pericardial effusion, or murmur.<sup>3</sup> We report a case  
44 of a previously healthy 45-year-old male who developed persistent dyspnea and edema following  
45 his recovery from SARS-CoV-2 infection. Cardiac evaluation revealed a para-cardiac mass  
46 (attached to the pericardium) and constrictive pericarditis for which he underwent cardiac  
47 surgery. The mass was removed and pathological evaluation reported an inflammatory mass.  
48 This study follows the principles of the Declaration of Helsinki. Ethics approval from an  
49 International Review Board was not applicable. Informed consent was obtained from the patient  
50 for the publication of the information and images.

51

## 52 **Case Report**

53 A 45-year-old male came to our clinic complaining of persistent exertional dyspnea and edema  
54 in 2021. The patient had a history of hospital admission been due to COVID-19 two months  
55 earlier and his symptoms had developed following his SARS-CoV-2 infection. Based on the  
56 medical records of his previous admission, the patient was a healthy and active individual with  
57 no limiting conditions and there was no report of any thoracic lesions. Upon evaluation, a large  
58 hypoechoic mass with a size of 9×3.1cm was observed on echocardiography, (Figure 1). The  
59 report was suggestive of a possible extra pericardial inflammatory mass or a large hematoma  
60 with central liquefaction over the right atrium (RA) and right ventricle (RV) that had  
61 compressive effects. In addition, there was bilateral pleural effusion (PE) which was turbid with

62 possible exudative effusion. There was also a mild to moderate RV dysfunction as well as a  
63 moderate left ventricular (LV) systolic dysfunction with normal LV size. LV ejection fraction  
64 (LVEF) by Simpson's mode was 45%. Based on the echocardiography result, although there was  
65 no thickening or calcification of the pericardium, physiological constrictive pericarditis was  
66 assumed to have occurred due to a mass. Therefore, the patient was referred to a cardiac surgeon  
67 with the impression of a localized exudative para-cardiac mass or hematoma to remove the  
68 mass, evaluate the pericardium, and drain the pleural effusion. A multi-slice high-resolution  
69 computed tomography (HRCT) scan of the chest revealed bilateral mild pleural effusion. In  
70 addition, there was patchy and ground-glass opacity in the right lower lobe (RLL) which was  
71 suggestive of a sequel of his previously SARS-CoV-2 infection. Linear atelectasis was detected  
72 in both the right and left upper lobes as well as the left lower lobe (LLL). The report also noted  
73 minimal atelectasis with pericardial effusion (Figure 2). Evaluation with cardiac magnetic  
74 resonance (CMR) confirmed the findings detected on echocardiography and CT scan. CT scan  
75 and MRI images are shown in Figure 2. Consequently, based on the results of echocardiography,  
76 CT scan, and CMR, a diagnosis of para-cardiac mass with compressive effects on the heart and  
77 physiological constrictive pericarditis was established and the patient was scheduled for surgery.  
78 Treatment with anti-inflammatory drugs (such as corticosteroids or interleukin-receptor  
79 antagonist) to reduce the para-cardiac mass was not initiated before cardiac surgery. On  
80 admission, his condition was good with unremarkable signs and normal physical examination.  
81 The patient had no significant medical or family histories and did not use any medication.  
82 Laboratory results and electrocardiography (ECG) were also normal. A summary of the patient's  
83 laboratory workup is shown in Table 1. A day after admission, the patient underwent open-heart  
84 surgery with general anesthesia for two hours. Upon surgery, a median sternotomy was  
85 performed and a large mass with the size of 10x4x3cm with severe adhesion to the RA and RV  
86 was detected (Figure 3). Cardiopulmonary bypass (CPB) with an arterial cannula in the  
87 ascending aorta was established and the venous return was achieved through the femoral vein.  
88 With beating heart bypass surgery, the mass was completely excised by sharp and blunt  
89 dissections. Subsequently, the patient was weaned from CPB without any difficulty. Chest  
90 closure was performed with the placement of two drains (mediastinal and right pleural). There  
91 were no signs of any complications during surgery. The patient was discharged from the hospital  
92 four days later with stable vital signs and no evidence of post-surgical complications. The

93 macroscopic pathological evaluation of the mass revealed a 10×6×3.5cm mass with elastic  
94 brown tissue and a soft cystic change at the center of the tissue. On microscopic evaluation, some  
95 lesions had necrosis, neutrophilic infiltration, and foamy macrophages. In addition, some of the  
96 fibrotic areas contained lymphocytic infiltration and lympho-plasma cells. However, there was  
97 no evidence of malignancy or granuloma. The patient did not have any medical complaints on  
98 his cardiology and cardiosurgery follow-up visits.

99  
100 Informed consent was obtained from the patient for publication of this case report and any  
101 accompanying images.

### 103 **Discussion**

104 There have been reports of cardiovascular complications and death or deterioration of pre-  
105 existing cardiac disorders following influenza infection.<sup>1</sup> SARS-CoV-2 infection may also lead  
106 to cardiovascular complications. Post-COVID-19 cardiac complications include myocarditis  
107 (42.1%), pericardial effusion (15.8%), acute MI (15.8%), cardiac arrhythmias (10.5%), RV  
108 mural thrombus with pulmonary embolism (5.3%), acute HF with cardiogenic shock (5.3%),  
109 cardiac tamponade, Takotsubo cardiomyopathy, pericarditis, and myopericarditis.<sup>1</sup> In addition,  
110 the majority of symptoms of post-COVID-19 cardiac complications include shortness of breath  
111 (52.6%), chest pain (36.8%), fever (26.3%), cough (26.3%), fatigue (10.5%), abnormal troponin  
112 levels (68.4%), abnormal B-type natriuretic peptide (BNP) levels (42.1%), and ST-segment  
113 elevation (52.6%).<sup>1</sup> Different factors are involved in myocardial injury and the development of  
114 cardiovascular complications of COVID-19 and include an increased hypercoagulable status,  
115 systemic inflammatory response to a viral infection which may lead to increased metabolic  
116 activity, angiotensin-converting enzyme 2 (ACE2) expression on cardiac cells (which act as a  
117 receptor for the SARS-CoV2 virus leading to the invasion of the cardiac cells by the virus), and  
118 direct viral injury of cardiac cells.<sup>1</sup>

119  
120 On the other hand, around 90% of primary cardiac tumors are benign including myxoma (50%),  
121 fibroelastoma (26%), fibroma (6%), lipoma (4%), inflammatory myofibroblastic tumors (IMTs)  
122 (< than 5%), and other benign tumors.<sup>2</sup> Primary pericardial masses are also usually benign and  
123 constitute about 6% to 10% of primary cardiac tumors. These benign primary pericardial masses

124 include lipomas, pericardial cysts, and paragangliomas, and hemangiomas. In addition,  
125 pericardial tumors are associated with some disorders such as Erdheim-Chester disease and  
126 IgG4-related disease. The clinical signs and symptoms of pericardial masses include dyspnea,  
127 edema, pleural effusion, orthopnea, pericardial effusion, or murmur.<sup>3</sup> Inflammatory  
128 pseudotumors of the pericardium are among the primary pericardial tumors that are benign.  
129 Although the exact cause of inflammatory pseudotumors is unknown, these masses have been  
130 reported to occur following surgery, trauma, IgG4-related sclerosing disease, or infection (with  
131 bacteria such as *Mycoplasma* and *Nocardia*). In addition, some have argued that inflammatory  
132 pseudotumors develop through a low-grade neoplastic process.<sup>4</sup> Cardiac and pericardial tumors  
133 are rare and to our knowledge, there have been no reports of SARS-CoV-2 infection inducing the  
134 development of a cardiac or para-cardiac mass (that may present as a cardiac mass such as in our  
135 case). We report a previously healthy 45-year-old male that developed persistent dyspnea and  
136 edema following SARS-CoV-2 infection. He underwent cardiac surgery due to the presence of a  
137 compressive mass with physiological constrictive pericarditis. The pericardium was intact and  
138 the mass was removed. The pathology report indicated an inflammatory nature for the mass.  
139 Confirmation of an association between the development of a para-cardiac inflammatory mass  
140 and SARS-CoV-2 infection requires further investigation.

141

## 142 **Conclusion**

143 We report the first reported case of a para-cardiac mass development following SARS-CoV-2  
144 infection in a previously healthy 45-year-old man.

145

## 146 **Author Contributions**

147 AM, AMS, and MMMR, MR clinically managed the patient. The primary draft was conducted by  
148 SH. AM, AMS, MMMR, and MR conducted the first edit. All authors have read and approved  
149 the final manuscript.

150

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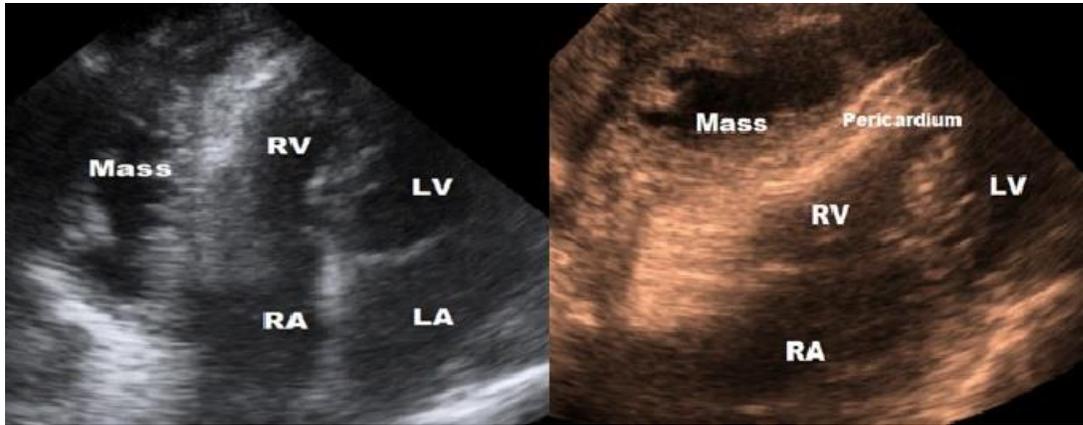
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 164 **Table 1.** A summary of the patient’s laboratory workup before surgery.

Laboratory test	Result	Normal range
WBC 10 <sup>9</sup> /L	8.77	Adult: 5-10
RBC 10 <sup>6</sup> /uL	4.27 (Low)	Male: 4.7-6.1
Hb g/dl	11.5 (Low)	Male: 13.2-16.2
HCT %	34.8 (Low)	Male: 39-52
Neutrophil %	63.5 (High)	40-60
Lymphocyte %	28.8	20-40
Platelets 10 <sup>3</sup> /ml	469 (High)	150-450
PTT seconds	64	30-45
PT seconds	18.1 (High)	12-16.5
INR (ratio)	1.36	Ratio: 1.34
Blood group	O positive	-
ESR (1hr) mm/hr	67 (High)	<10
CRP mg/l	32.5 (High)	Up to 6
Magnesium mg/dl	2.2	1.8-2.6
Hbs Ag (Index)	Negative	<1: Negative
HCV Ab (Index)	Non-reactive	Non-reactive: <0.9
HIV (IFA) (Index)	Non-reactive	Negative: <0.25

165 **Abbreviations:** White blood cells (WBC), Red blood cells (RBC), Hemoglobin (Hb),  
 166 Hematocrit (HCT), PTT (Partial thromboplastin time), PT (Prothrombin time), INR  
 167 (international normalized ratio), ESR (erythrocyte sedimentation rate), CRP (C-reactive protein),

168 Hbs Ag (Hepatitis B virus surface antigen), HCV Ab (Hepatitis C virus antibody), HIV (human  
169 immunodeficiency virus), and Indirect fluorescent antibody (IFA).

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171

172 **Figure 1.** Echocardiography: A large hypoechoic mass (size: 9x3.1cm in size) suggestive of an  
173 extra pericardial inflammatory mass or a large hematoma with central liquefaction over the right  
174 atrium (RA) and right ventricle (RV) causing compressive effects.

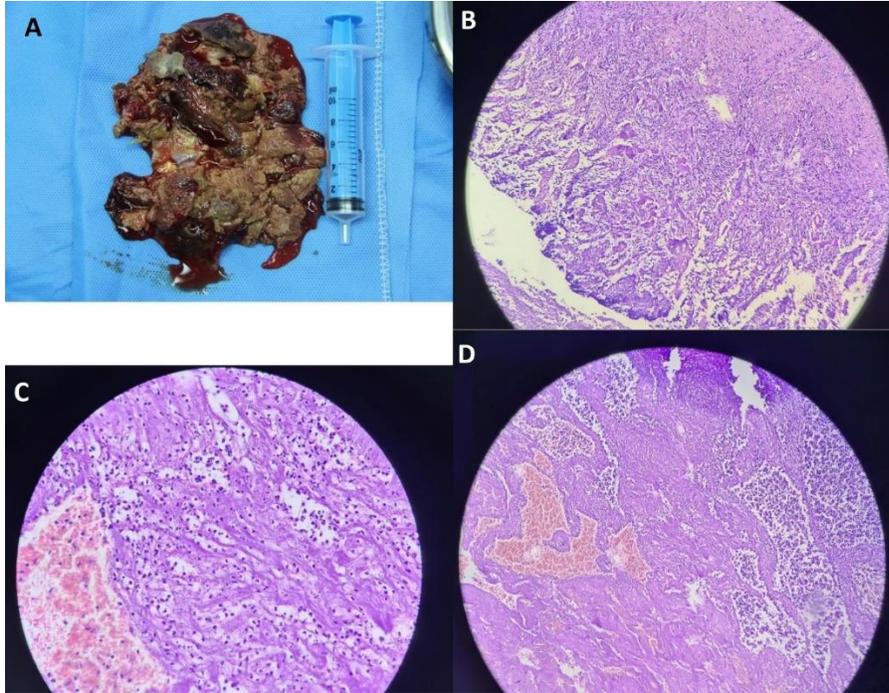
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177 **Figure 2. A.** Multi-slice high-resolution computed tomography (HRCT) scan of the chest  
178 revealing bilateral mild pleural effusion; Patchy and ground-glass opacity in the right lower lobe  
179 (RLL) [suggestive of a sequel of the patient's previously COVID-19 infection]; Linear  
180 atelectasis in both the right and left upper lobes as well as the left lower lobe (LLL); Minimal  
181 atelectasis with pericardial effusion; Arrow showing the mass. **B and C.** MRI of the chest;  
182 Arrows showing the mass (RV: Right Ventricle; LV: Left ventricle; PE; pleural effusion).

183



184

185 **Figure 3. A.** Image of the large mass (size of 10x4x3cm) that was excised upon surgery. **B, C,**  
186 **and D.** Microscopic evaluation of the mass (some lesions had necrosis, neutrophilic infiltration,  
187 and foamy macrophages; some of the fibrotic areas contained lymphocytic infiltration and  
188 lympho-plasma cells. No evidence of malignancy or granuloma).