CASE PRESENTATION

A challenging case of Takotsubo cardiomyopathy complicated by acute mitral regurgitation

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Abstract: Introduction – Takotsubo cardiomyopathy (TCM) is a cardiac condition that mimics the clinical presentation of acute coronary syndrome, but without any evidence of obstructive atherosclerotic coronary artery disease. The prognosis is usually good, with improvement of the left ventricular (LV) dysfunction within a few weeks. There are some possible complications that may occur in the acute phase. Case presentation – We report the case of a 77 years old female patient, who presented with intense chest pain, occurred in conditions of a major emotional stress, associated with pulmonary oedema and cardiogenic shock. Echocardiographic exam showed the presence of a major mitral regurgitation (MR) in addition to the severely impaired LV systolic function (estimated ejection fraction of 20-25%). The patient underwent emergent coronary angiography, revealing no significant coronary arteries lesions, but instead an image of LV “apical ballooning” at ventriculography, suggestive for TCM. The patient evolution was slightly favourable, under supportive medical treatment, with full recovery of LV function at 4 weeks follow-up, and mild MR. Conclusion – This case illustrates an unexpected evolution of a critical patient, with severe LV dysfunction and acute ischemic MR, that had instead a full recovery, typical for TCM, which raised major difficulties in terms of diagnosis and management, but with excellent outcome. Keywords: Takotsubo cardiomyopathy, acute mitral regurgitation, pulmonary oedema, cardiogenic shock, echocardiography.


INTRODUCTION

Takotsubo cardiomyopathy (TCM), also called transient left ventricular (LV) apical ballooning or “broken heart syndrome,” is a cardiac condition that mimics the clinical presentation of acute coronary syndrome, but without any evidence of obstructive atherosclerotic coronary artery disease (CAD)¹¹. The incidence of TCM is postulated in most studies between 1% to 2.2% of ACS and is higher in postmenopausal women³⁷⁻⁸.

Emotional or physical stress at onset is common, although a triggering event is not always present. Proposed etiopathogenic mechanisms include: multi-vessel coronary artery spasm, impaired cardiac microvascular function, endogenous catecholamine induced myocardial stunning and microinfarction or possible direct catecholamine-mediated myocyte injury.²⁻⁶⁻⁹

Although it is considered to be a benign condition, with generally favourable outcome, there are possible

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complications that may occur in the acute phase (up to 50% of patients develop complications)^10. Heart failure is the most common among it, mainly due to the impaired LV systolic function. There are also mentioned life-threatening arrhythmias, sudden cardiac death, right ventricular involvement, thrombus formation, resulting in stroke or arterial embolism or rarely, ventricular rupture^4,10-13. Up to 20–25% of patients with TCM may develop acute mitral regurgitation (MR) and/or left ventricular outflow tract (LVOT) obstruction, severe complications that can significantly worsen the clinical patient status, leading to cardiogenic shock and/or pulmonary oedema, with major impact on vital prognosis^7.

CASE PRESENTATION
We present the case of a 77 years old female patient that was admitted in our emergency care unit at two hours after the onset of a persistent, intense, constriction chest pain, associated with shortness of breath, occurred in conditions of a major emotional stress (unexpected loss of a son). She had never experienced a similar chest pain before. Her medical background included 20 years history of hypertension, treated intermittently with calcium channel blocker and type 2 diabetes mellitus, diagnosed 5 years ago, with inadequate glycemic control under treatment with Metformin. She had neither history of smoking nor family history of CAD.

On presentation, the physical exam revealed a hemodynamic unstable patient, blood pressure (BP) of 80/50 mmHg, pulse of 100 beats/min, 90% oxygen saturation under the use of a face mask at 6 L/min, respiratory frequency of 30 breaths/min, chest pain present 6/10. Cardiac exam showed tachyrhythmic heart sounds, intense systolic murmur 3/6 at the apex, normal symmetrical peripheral pulse, inspiratory crackles over posterior bases in both lungs.

The electrocardiogram (ECG) on admission showed sinus rhythm, ST segment depression followed by deep T-wave inversions in both anterior and inferior leads (Figure 1).

The transthoracic echocardiography showed severe impaired LV systolic function (estimated ejection fraction- EF of 20-25%), with severe hypokinesia of the apex and the mid-apical segments of the LV walls and also severe mitral regurgitation (MR), with eccentrically directed posterior jet, normal-sized atrial cavities, no pericardial effusion (Figure 2).

Laboratory exams showed elevated cardiac Tropinin I (4 ng/ml, normal 0.0-0.3 ng/ml) with slightly elevated creatine kinase on admission (30 U/L, then normal values), normal lactate dehydrogenase and aspartate transaminase values, both on admission and in evolution, elevated pro-brain natriuretic peptide (1500 pg/ml, normal 125 pg/ml), mild leucocytosis (12.000/mm³), elevated C-reactive protein (15 mg/L, normal 5 mg/L), normal creatinine levels, persistently elevated blood glucose levels (160 mg%-130 mg%) and dyslipidemia (total cholesterol 235 mg%, HDLc 40 mg%, LDLc 160 mg%, triglyceride 175 mg%).

The patient underwent a coronary angiography, which showed no significant coronary artery lesions (Figure 3). Left ventriculography revealed severe hypo-

Figure 1. ECG on admission: sinus rhythm, 100 bpm, ST oblique descendent segment depression of 1-2 mm in V3- V6 and deep T-wave inversions in DI, aVL, DII, aVF, V3- V6.
kinesia of the apex, outlining an image of LV “apical ballooning”, suggestive for TCM (Figure 4).

Under supportive medical treatment, with intravenous dobutamine and noradrenaline, the hemodynamic patient’s status improved. After reaching normal BP values, low doses of nitrates was used in order to relief dyspnoea. Furthermore, the evolution was slightly favourable, with symptoms relief and without arrhythmic events until discharge. Regarding the presence of acute significant MR, which was thought to be in the context of ischemia, according to the initial cardiovascular surgical evaluation, the mitral valve repair procedure was delayed, taking into account the severe LV systolic dysfunction.

Treatment at discharged included antiplatelet therapy (acetyl salicylic acid), statine, beta-blocker and angiotensin-enzyme converting inhibitor in low doses (limitation for BP).

At 4 weeks follow-up, at transthoracic echocardiographic examination it was noticed a full recovery of LV function and mild MR (Figure 5), aspects that further sustained the diagnosis of TCM. The patient was asymptomatic, without residual chest pain and the ECG showed normal morphology of QRS complex, without ST-T segment changes (Figure 6).

DISCUSSIONS
This case illustrates a particular evolution of TCM, complicated by acute MR, pulmonary oedema and cardiogenic shock, which raised major issues in terms of management, considering the paucity of the literature data in this condition.
There are several hypotheses regarding the mechanisms of acute MR associated with TCM.

De Backer et al. showed in a two-year, two-center experience study, that up to 20% of TCM patients may develop left ventricular outflow tract (LVOT) obstruction, being in accord with some previous studies\textsuperscript{14-15}. Factors associated with this complication were advanced age, septal bulging, systolic anterior motion (SAM) - induced MR, and hemodynamic instability\textsuperscript{7}. In this study, all patients with LVOT obstruction developed SAM of the anterior mitral valve leaflet and high-grade MR, and nearly half of them presented cardiogenic shock\textsuperscript{7}.

On the other hand, Izumo et al. found in their study another mechanism involved: mitral valve leaflet tethering secondary to papillary muscle dysfunction or dis-
placement associated with regional or global LV systolic dysfunction and LV enlargement. This is previously known as an important cause of ischemic MR. The transient ischemia, with classic LV apical “ballooning” and consecutively impaired LV systolic function is a common finding in TCM, those it is reasonable to be aware of the occurrence of acute MR in these conditions.

In the case presented, echocardiography revealed besides a severely impaired LV systolic function, a major MR, but no signs of SAM or obstruction in LVOT. In the absence of SAM or LVOT obstruction, the low EF of 20-25%, together with the mid-apical ischemia and the LV apical ballooning, consists in arguments for the hypothesis of transient ischemic MR.
Furthermore, the identification of the specific mechanism of acute MR in this condition is important for proper management of the patients. For instance, in cases with LVOT obstruction associated, the use of inotropic agents or vasodilators, otherwise often used in the treatment of ACS, can worsen the grade of obstruction. Instead, there are recommended beta-blockers, fluid intake or intra-aortic balloon pump (IABP), for refractory cardiogenic shock, as a bridge to clinical recovery.\textsuperscript{7,20-21}.

In cases with MR without SAM, because of severe LV dysfunction associated, the inotropic therapy is often necessary, with the recommendation for careful titration of these in order to avoid triggering LVOT obstruction.\textsuperscript{22-23} We used double association of inotropic agents, both dobutamine and noradrenaline, in order to obtain hemodynamic stability of our patient.

Another important and disputed question was what is best to do regarding the indication of mitral valve replacement in these cases? The current European and American guidelines recommend mitral valve surgery in symptomatic patients with severe acute MR.\textsuperscript{24,25} On the other hand, the fact that both LV dysfunction and MR associated with TCM are reversible, the idea of supportive medical treatment until complete recovery is feasible.\textsuperscript{21,26}

In the case presented, we asked for cardiovascular surgical evaluation, that suggested us to postpone the mitral valve replacement intervention, considering the unacceptable high risk associated with the severe LV systolic dysfunction. The fact that we adopted a conservative strategy was the best thing to do in this case, which was confirmed later; when the patient returned after 4 weeks, with complete recovery of LV function and mild MR, that eliminated the previous indication of surgery.

CONCLUSIONS

Despite the mechanism involved, the association of significant acute MR in a patient with TCM, increases the risk of adverse outcomes. This underline the importance of an accurate diagnosis, systematic and repeated echocardiographic evaluations of a patient diagnosed with TCM, in order to unmask as earlier as possible such complication.\textsuperscript{27}

This case exposed the difficulties of diagnosis and treatment of a critical patient, presented with severe LV dysfunction and acute MR, leading to pulmonary oedema and cardiogenic shock that had an excellent recovery under conservative treatment, typical for TCM.

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References


