One Patient, Three Variants, Four Episodes of Takotsubo Cardiomyopathy

Eitan Frankel, MD, Patrick Prior, MD, Hetal Mchta, MD, Priya Rajagopalan, MD, David H. Wiener, MD, FACC, FAHA, FASE, Andrew Rose, MD, FACC, and Gregory D. Marhefka, MD, FACC, FACP, Philadelphia, Pennsylvania

INTRODUCTION

The first case reports of takotsubo cardiomyopathy (TCM) were published in the early 1990s and described a pattern of apical ballooning.1,2 Since then, the literature on TCM has rapidly expanded as the diagnosis has become an increasingly recognized clinical entity. Until the mid-2000s, TCM was considered primarily a disease of the apex and midventricular segments of the left ventricle. However, it is now understood that there are generally four recognized variants, with relative incidences of 81.7% (apical), 14.6% (midventricular), 2.2% (basal), and 1.5% (focal).3-6 In many patients, cardiac function recovers within a few months, and mortality is similar to that among age- and sex-matched cohorts.7,8 However, a smaller subset of patients develop in-hospital and chronic complications including cardiogenic shock, persistent heart failure, recurrence of TCM, and death.3 We present the case of a patient who developed four separate episodes and three unique variants of TCM over the course of 1.5 years.

CASE PRESENTATION

The patient is a 51-year-old woman with a history of hypertension, who initially presented in July 2017 to an emergency department with abdominal pain, nausea, and emesis. Electrocardiography (ECG) was notable for T-wave inversions, and her troponin level was elevated. Transthoracic echocardiography (TTE) demonstrated mild left ventricular (LV) enlargement, apical ballooning with mid and apical akinesis, and an estimated LV ejection fraction (LVEF) of 35%. Left heart catheterization was performed, which demonstrated no evidence of obstructive coronary artery disease, and ventriculography showed normalization of the left ventricle with normal wall motion.

Nine months later, she returned with recurrent gastroparesis. ECG demonstrated diffuse T-wave inversions and a prolonged corrected QT interval. Troponins were again mildly elevated. TTE demonstrated a different segmental pattern of TCM compared with her two prior episodes: midventricular-variant akinesis with preservation of the basal segments. The patient was discharged after a complicated hospital course and then presented again 1 month later with similar symptoms of abdominal pain, nausea, and emesis. Given her recent history of TCM, TTE was performed. Per the report, this study demonstrated resolution of the previously seen regional wall motion abnormalities and an LVEF of 55%. Her presenting symptoms were attributed to gastroparesis, and the patient was again discharged.

Unfortunately, between July 2017 and February 2019, the patient was admitted to the hospital seven times and required repeat endoscopies for management of her abdominal symptoms. This case report focuses on the multiple variants of TCM with which the patient presented throughout her hospitalizations (Figure 1). All echocardiograms in boldface type are available and presented in this case report.

The patient first presented to our hospital in December 2017 with a chief symptom of abdominal pain. During her hospitalization, TTE not only showed normalization of LVEF, but it actually showed hyperdynamic LV systolic function (LVEF 75%) without segmental wall motion abnormalities (Figure 2, Videos 1 and 2).

About 1 week into her hospitalization, the patient developed acute onset of substernal, nonradiating chest pain. ECG demonstrated new, diffuse T-wave inversions and poor R-wave progression. Troponin was again mildly elevated. TTE again demonstrated recurrent apical ballooning with midventricular and apical akinesis with preservation of the basal segments and an LVEF of 33% (Figure 3, Videos 3 and 4). With the patient’s known history, the findings were attributed to a recurrence of TCM, and she was treated medically. In follow-up with her outpatient cardiologist, repeat echocardiography showed normalization of the left ventricle with normal wall motion.

Most recently, 18 months after her initial episode of TCM, the patient was again admitted for abdominal pain. ECG demonstrated sinus bradycardia without T-wave inversions, and troponins were mildly elevated. TTE at this time demonstrated her third variant of TCM, with basal akinesis (reverse pattern) and an LVEF of 45% (Figure 5, Videos 7 and 8). She recovered without incident and was discharged home after repeat endoscopy with onabotulinum toxin A injection. Notably, since the patient underwent left heart catheterization for her initial presentation, she has been managed conservatively without repeat catheterization for her recurrent cardiomyopathy.

DISCUSSION

TCM is now a well-recognized entity in the differential diagnosis of acute onset chest pain, non-ST-segment elevation myocardial infarction, and ST-segment elevation myocardial infarction. Episodes are often incited by significant emotional or physical stressors, but there are cases in which no stressor can be identified.9 Female gender and the presence of neurologic and psychiatric disease have repeatedly been demonstrated to be associated with TCM. These groups of patients have higher levels of sympathetic activation, and there is an
incompletely understood relationship between neurohormonal function and the development of TCM.\(^\text{10}\)

However, although TCM is an important clinical entity, there is still a lack of consensus regarding diagnostic criteria. Perhaps the most well-known criteria are the Mayo criteria, which were proposed in 2004 and modified in 2008. These criteria define TCM as (1) transient hypokinesis, akinesis, or dyskinesis of a LV segment with regional wall motion abnormalities that extend beyond distribution of a single vascular distribution; (2) absence of coronary artery disease; (3) new electrocardiographic abnormalities; and (4) the absence of pheochromocytoma or myocarditis. However, does this mean that every patient presenting with possible TCM needs catheterization, biochem-

**VIDEO HIGHLIGHTS**

**Video 1:** Normal echocardiographic findings, apical two-chamber view. TTE demonstrating hyperdynamic LV function with no segmental wall motion abnormalities.

**Video 2:** Normal echocardiographic findings, apical four-chamber view. TTE demonstrating hyperdynamic LV function with no segmental wall motion abnormalities.

**Video 3:** Apical-variant TCM, apical two-chamber view. TTE demonstrating apical ballooning with midventricular and apical akinesis with preservation of the basal segments.

**Video 4:** Apical-variant TCM, apical four-chamber view. TTE demonstrating apical ballooning with midventricular and apical akinesis with preservation of the basal segments.

**Video 5:** Midventricular-variant TCM, apical two-chamber view. TTE demonstrating mid-ventricular variant akinesis with preservation of the apical and basal segments.

**Video 6:** Midventricular-variant TCM, apical four-chamber view. TTE demonstrating midventricular variant akinesis with preservation of the apical and basal segments.

**Video 7:** Basal-variant TCM, apical two-chamber view. TTE demonstrating basal akinesis (reverse pattern).

**Video 8:** Basal-variant TCM, apical four-chamber view. TTE demonstrating basal akinesis (reverse pattern).

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**Figure 1** Serial echocardiograms. This is a chronological reporting of the available echocardiograms between July 2017 and May 2019. Given that the patient presented to multiple hospitals and providers with different electronic medical records, not all the images were available to be reviewed. Locations at which reviewed echocardiograms were performed included Thomas Jefferson University Hospital (TJUH), local outside hospitals (OSH), and a private cardiologist’s office. For each echocardiogram, when available, both the images and reports were reviewed. This figure highlights the multiple variants of TCM that are explored in this report.
Figure 2 Normal echocardiographic findings. TTE demonstrating hyperdynamic LV function with no segmental wall motion abnormalities. Apical two-chamber view (A) and apical four-chamber view (B).

Figure 3 Apical-variant TCM. TTE demonstrating apical ballooning with midventricular and apical akinesis with preservation of the basal segments. Apical two-chamber view (A) and apical four-chamber view (B).

Figure 4 Midventricular-variant TCM. TTE demonstrating midventricular-variant akinesis with preservation of the apical and basal segments. Apical two-chamber view (A) and apical four-chamber view (B).
comorbidities of TCM. In addition, increased awareness of this phenomenon may help physicians more effectively and efficiently manage patients with recurrent episodes of TCM.

CONCLUSION

Since TCM was first described, four LV variants have been recognized: apical, midventricular, basal, and focal. We describe a unique case of one patient, with four discrete episodes of TCM, demonstrating three distinct variants.

SUPPLEMENTARY DATA

Supplementary data related to this article can be found at https://doi.org/10.1016/j.case.2020.01.009.

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