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## Case report

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# Takotsubo cardiomyopathy associated with opiate withdrawal

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## **Learning Point for Clinicians**

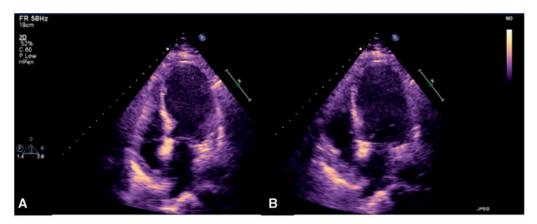
Classically, takotsubo cardiomyopathy (TTC) or 'broken-heart-syndrome', describes the association of a trigger factor such as 'death of a loved one' with induction of a transient cardiomyopathy. However, the trigger factor can be any one of a diverse group of physical or emotional stressors. It has recently been reported that drug withdrawl, particularly from opiates, can also initiate TTC. Thus, prompt recognition of withdrawal-associated TTC is essential for appropriate management and optimal clinical outcome.

## **Case report**

A 60-year-old woman with hepatitis C, major depression and chronic pain syndrome presented to the emergency department complaining of abdominal discomfort, chills, diaphoresis, nausea and emesis for 72 hr. Several days prior to presentation, she had run out of MS Contin 100 mg (qid), and because of her symptoms she had discontinued all her medications for 48 hr. Her cardiovascular risk factors were cigarette smoking and hypertension.

On admission, the patient was in distress, reiterating her need for morphine. Vital signs were blood pressure 141/89 mm Hg, pulse 117/min and

respiratory rate 24/min. Her chemistry panel and venous blood gases revealed a mixed acid base disorder and liver function tests were normal. Shortly after she complained of sudden onset of chest pain and her electrocardiogram showed ST-segment elevation in the precordial leads. Troponin I was elevated (21.12 ng/ml, ref <0.04). Echocardiography revealed left ventricular (LV) ejection fraction of 35% with apical akinesis and hypokinesis of the mid-anterior and mid-inferior segments (Figure 1). The patient received urgent cardiac catheterization, which revealed no significant coronary artery disease. Left ventriculography demonstrated LV apical akinesis, basal hypercontractility, and LV ejection fraction of 40%, and LV end diastolic pressure of 29 mm Hg. These findings were consistent with takotsubo cardiomyopathy (TTC) and the patient was transferred to the cardiac care unit where a nitroprusside infusion was initiated and morphine was administered. The patient's hospital course was complicated by pulmonary edema, which required intensive therapy including diuresis, vasodilators and brief ventilatory support. Over the next several days, therapy with an angiotensin converting enzyme inhibitor and beta-blocker was started. Morphine was progressively decreased and discontinued after <1 week. The patient's condition stabilized over the next 3 weeks. By hospital day 27, she 302 A. Sarcon et al.



**Figure 1.** Echocardiography reveals left ventricular ejection fraction of 35% with apical ballooning pattern (**A** in systole and **B** in diastole).

was fully ambulatory, her echocardiogram had normalized, and her electrocardiogram had only residual anterior T-wave inversion.

### **Discussion**

TTC is characterized by transient, regional systolic dysfunction of the left ventricle without obstructive epicardial coronary artery disease. It predominantly affects postmenopausal women and is frequently triggered by an emotional or a physical stressor. TTC related to medication withdrawal has been reported in a diverse group of patients. Interestingly, Americans account for approximately 80% of the global consumption of prescription analgesics. The rising consumption of analgesic medications worldwide suggests a potential increase in the frequency of withdrawal-associated TTC.

The pathophysiology of TTC is not well understood, but excessive catecholamine and adrenergic stimulation have been implicated.<sup>3</sup> Drug withdrawal can be associated with a hyperadrenergic state and central nervous system irritability,<sup>4</sup> which may provide a mechanism for withdrawal-associated TTC. We have identified seven additional cases of drug withdrawal-associated TTC in our International Takotsubo Registry at the leading hospital Zurich, Switzerland (www.takotsubo-registry.com). We suggest that withdrawal-associated TTC is under-diagnosed due to limited awareness. The majority of

these patients was women with ST-segment elevation and all demonstrated the LV apical ballooning pattern, with reduced ejection fraction of 36%. Inhospital complications occurred among our patients including one death. Our patient's complex presentation involving opiate withdrawal and TTC was manifested by acute pulmonary edema and metabolic alkalosis, which prolonged her hospitalization. Recognition of the potential for patients to develope TTC during drug withdrawal to develop TTC is essential for appropriate management and favorable outcome.

Conflict of interest: None declared.

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