Case report

Unusual cause of chest pain, Bornholm disease, a forgotten entity; case report and review of literature

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ARTICLE INFO

Keywords:
Coxsackievirus
Chest pain
Chest X ray
Pleurodynia

ABSTRACT

Chest pain is a common symptom culminating in hospital admissions and specialist referrals. Although cardiac work up is pursued in most of the cases, cardiac etiology is found to be the culprit in minority of the cases. Acute chest pain is a clinical syndrome that may be caused by almost any condition affecting the thorax, abdomen, or internal organs. On occasions this extensive and expensive diagnostic work up can be avoided with awareness of commoner and non-lethal reasons. We report a case of a woman with Bornholm disease secondary to Coxsackievirus B5 (CBS) infection and supplementary review of literature till date.

1. Introduction

Chest pain is a common symptom culminating in hospital admissions and specialist referrals. Although cardiac work up is pursued in most of the cases, cardiac etiology is found to be the culprit in minority of the cases [1–5]. Data from epidemiologic studies have indicated that the overall proportion of non-cardiac chest pain (NCCP) among patients with chest pain is reported between 20 and 40%. This fraction is robust and internationally similar in different countries, such as Germany, Europe, USA, China, or Australia [6]. Acute chest pain is a clinical syndrome that may be caused by almost any condition affecting the thorax, abdomen, or internal organs. When a patient presents with chest pain in the emergency room, the first priority is to consider life-threatening causes such as acute coronary syndromes, aortic dissection, pulmonary embolism (PE), ruptured aortic aneurysm, and tension pneumothorax [7]. The diagnostic work up for these etiologies bear a rightful demand on time of the emergency room physicians and other resources. On occasions this extensive and expensive diagnostic work up can be avoided with awareness of commoner and non-lethal reasons. Absence of knowledge of this entity often confuses clinicians to pursue the alternative diagnoses such as acute abdomen and acute chest syndromes including pleurisy and coronary syndromes. We report a case of a woman with Bornholm disease secondary to Coxsackievirus B5 (CBS) infection and supplementary review of literature till date.

2. Case

A 68-year-old female was admitted to the hospital with acute onset and progressively worsening sharp epigastric and left sided “gripping” chest pain of severe intensity without any radiation or change in character with position, two days prior to the presentation. The onset was without any trauma. Her past medical history was remarkable for hypertension, hyperlipidemia and hypothyroidism. Her medications included aspirin, amlodipine, lovastatin and levothyroxine. Patient described her pain as pleuritic in nature with exacerbation on taking a deep breath and no position relieved the pain. Patient denied any recent travel. There were no family members with similar symptomatology. On examination, patient was in significant discomfort and clutching pain in the left side of chest and epigastrium. She had normal blood pressure but was mildly tachycardic with a heart rate of 105 beats per minute. She had a low grade temperature of 99.9 degree Fahrenheit. Her oxygen saturation was 97% without any supplemental oxygen. The pain was not reproducible on palpation and there was no visible rash. Upon auscultation of the chest there was appreciable pleural rub on the left, and the right lung examination was unremarkable. Best of her systemic examination was unremarkable. Patient was given appropriate
analgesia.

Laboratory data showed normal white blood cell count of 9.4 × 1000/μL (3.4–11.0 × 1000/μL) with 54% neutrophils, along with normal hemoglobin and hematocrit. Serum chemistry was normal with creatinine of 0.75 mg/dL (0.5–1.5 mg/dL) and normal liver function tests. Due to concerns of chest pain and acute coronary syndrome her cardiac biomarkers were checked and troponin T was found to be normal (less than 0.030 ng/mL) on three occasions, each 6 hours apart. Creatine kinase (CK) was normal 122 U/L (24–173 U/L). MB component of CK was found to be normal as well at 3.7 ng/mL (0.0–5.3 ng/mL). EKG done for the patient to rule out any ongoing cardiac ischemia was normal (Fig. 1) without any ST wave changes.

Chest x-ray done showed small bilateral effusions (Fig. 2).

Due to unclear diagnosis of pleuritic chest pain at this time and sinus tachycardia, patient underwent computed tomography (CT) of the chest to rule out pulmonary embolism and was found to be negative for any embolic event or pulmonary infarction, aortic abnormality and gastro-esophageal abnormality. In addition there was no muscular edema noted on the scan. After ruling out the most ominous causes of chest pain, patient was still symptomatic. Now suspicion was high for rare causes of chest pain. Wider net was now casted for vasculitic disorders and rare infections causing similar symptomatology. Erythrocyte sedimentation rate and C-reactive protein were elevated to 65 mm/hr (2–15 mm/hr) and 17.2 mm/hr (0.0–4.9 mg/L) respectively. Her complement levels for C3 were 161 mg/dL (82–167 mg/dL) and for C4 were 16 mg/dL (14–44 mg/dL). Her C-ANCA and p-ANCA panel was negative. Antinuclear antibodies were negative. Further serological testing was done and she was found to have negative serology for coxsackievirus A types 7, 9, 16 and 24. Finally serology returned positive for coxsackievirus B type 5 (CB5) by complement fixation reaction (CFR, Virion/Serion, Clindia Benelux b.v., Almere, the Netherlands) and negative for coxsackievirus B types 1, 2, 3, 4 and 6. Patient was now continued to be treated as epidemic pleurodynia or Bornholm disease (caused by CB5) with symptom control using non-steroidal anti-inflammatory drugs and intravenous fluids. Patient’s pain improved significantly over the next 48 hours with resolution of fever and tachycardia. Patient was discharged home on the fourth day with complete resolution of symptoms. At 2 weeks follow up, patient was completely chest pain free. Repeated chest x-ray at this time showed normal lung fields with clearance of bilateral small pleural effusions (Fig. 3).

3. Discussion

First described in Daae and Homann in Norway, disease has been known with many forenames such as Bornholm (City in Denmark) disease, Devil’s grippe, epidemic myalgia and epidemic pleurodynia [8]. However in the early years there was no identifiable pathogen associated with the disease. Since then there have been glaring advances in microbiological and virological sciences. It was not until 1949
when the Coxsackie virus was thought to be associated with epidemic pleurodynia [8,10]. Ever since multiple studies have described the association of Bornholm disease with Coxsackievirus B (CBV) and less frequently echovirus types 1, 6, 8, 9 and 19 and further less frequently with Coxsackievirus A (CVA) types 4, 6, 9 and 10 [11,12]. Typical presentation with excruciating pleuritic chest pain or epispiacastic pain, fevers and complete recovery with supportive care is the norm [8,11,13–15]. Characteristic positive history with negative objective findings makes this malady easily distinguishable. Over the years the disease has been reported in conjunction with various strains of CBV. Data on association of epidemic pleurodynia with coxsackievirus B5 (CB5) is scanty. Bain et al., in 1961, for the first time described epidemic pleurodynia, benign pericarditis and aseptic meningitis which was attributable to CB5 [16]. In general, respiratory complaints are more common with type B virus infections as compared to type A viral strains. Also coxsackievirus type B strains result is more number of hospitalizations [14]. Extensive review of literature is provided in the following table with significant studies and reports (in English) till date (Table 1).

| Title                                                                 | Author                                      | Year |
|----------------------------------------------------------------------|---------------------------------------------|------|
| Pleurodynia caused by an echovirus 1 brought back from the tropics   | Leendertse et al.                          | 2013 |
| Bornholm disease—a pediatric clinical entity that can alert a thoracic surgeon | Tagarakis et al.                          | 2011 |
| Epidemic pleurodynia caused by coxsackievirus B3 at a medical center in northern Taiwan | Huang et al.                               | 2010 |
| Epidemic pleurodynia (Bornholm disease) outbreak in Singapore. A clinical and virological study | Chong et al.                               | 1975 |
| Association of Group B Coxsackieviruses with Cases of Pericarditis, Myocarditis, or Pleurodynia by Demonstration of Immunoglobulin M Antibody | Schmidt et al.                             | 1973 |
| Epidemic pleurodynia in Aden associated with infection by echovirus type 1 | McCracken et al.                          | 1969 |
| The occurrence of Bamblle Disease (epidemic pleurodynia) in Norway | Vogelsang                                | 1967 |
| Epidemic pleurodynia, orchitis, and myocarditis in an adult due to Coxsackie virus, group B, type 4 | Swann | 1961 |
| Epidemic pleurodynia (Bornholm disease) due to Coxsackie B-5 virus. The inter-relationship of pleurodynia, benign pericarditis and aseptic meningitis | Bain et al.                               | 1961 |
| A case of epidemic pleurodynia or Bornholm disease | Hale and Pillai                          | 1956 |
| Studies on the etiology of Bornholm disease (epidemic pleurodynia). II. Epidemiological observations | Johnson                                   | 1954 |
| Oxford epidemic of Bornholm disease | Warin et al.                              | 1951 |
| Epidemic pleurodynia with special reference to the differential diagnosis in acute abdominal pain | Ekman | 1953 |
| Isolation of Coxsackie virus from patients with epidemic pleurodynia | Thordarson et al.                          | 1953 |
| Epidemic pleurodynia in Texas; a study of 22 cases | Huebner et al.                           | 1953 |
| Bornholm disease in children | Duney et al.                              | 1953 |
| Studies on the etiology of epidemic pleurodynia (Bornholm disease). I. Clinical and virological observations | Galbinus et al.                           | 1952 |
| The importance of Coxsackie viruses in human disease, particularly herpangina and epidemic pleurodynia | Huebner et al.                           | 1952 |
| An outbreak of epidemic pleurodynia, with special reference to the laboratory diagnosis of Coxsackie virus infections | Lazarus et al.                           | 1952 |
| A virus isolated from a case resembling epidemic pleurodynia; a preliminary report | Howes | 1951 |
| Epidemic myalgia in Cape Town; pleurodynia or Bornholm disease | Prisman and Shrand | 1950 |
| Coxsackie viruses and Bornholm disease | Findlay and Howard | 1950 |
| The etiology of epidemic pleurodynia: a study of two viruses isolated from a typical outbreak | Weller et al.                           | 1950 |
| Epidemic pleurodynia; clinical and etiologic studies based on 114 cases | Finn et al.                               | 1949 |
| Acute benign dry pleurisy in the Middle East | Scadding                                  | 1946 |

Importance of recognizing this disease in not merely an academic pursuit. According to a CDC report, in 2016 CB4 claimed for 1.9% of all enterovirus infections charting at least 54 cases all over the United States [38]. The differential diagnosis for fever and chest pain or abdominal pain includes serious illnesses such as acute appendicitis,
pancreatitis, cholecystitis, pulmonary embolism and acute coronary syndromes amongst many others. Some of these can be lethal if not diagnosed with time. Awareness of this entity and inclusion in the differential diagnoses, can avoid the pitfall of surgical intervention at times. Moreover, an understanding of the disease and its natural course can avoid unnecessary and expensive investigations. Knowledge of its viral agent will also avoid unwarranted use of broad-spectrum antibiotics. Lastly it will significantly allay the fear and anxiety of the patients and even the doctor of a serious condition due to its self-limiting course and complete recovery in large majority.

**Authors' contribution**

AL: Conception of study + Manuscript draft + Data collection + Review.  
JA: Image Selection + Data collection + Critical Review.  
SI: Image Selection + Data collection + Critical Review.  
MN: Conception of study + Manuscript draft + Critical Review.  
MSK: Manuscript draft + Critical Review.  
GMA: Manuscript draft + Critical Review + Expert Opinion.

**Conflict of interest**

Authors claim no conflict of interest.

**Funding**

This work required no external funding.

**Appendix A. Supplementary data**

Supplementary data to this article can be found online at [https://doi.org/10.1016/j.rmcr.2018.10.005](https://doi.org/10.1016/j.rmcr.2018.10.005).

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