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Pacemaker endocarditis caused by *Propionibacterium acnes*: a case report

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Abstract

A 74-year old man, with a permanent pacemaker placed 2 years ago for high grade atrioventricular block, was admitted for worsening fatigue, confusion and thrombocytopenic purpura without fever. White blood cell count and C-reactive protein were elevated and echocardiography revealed a 6 x 3 cm echogenic mass surrounding the pacemaker leads. Multiple blood cultures were performed and one bottle only grew *Propionibacterium acnes* at 93 hours. The patient underwent surgery and 16S rRNA gene PCR amplification confirmed the presence of *P. acnes* in the removed vegetation.

Patients with late onset device-related endocarditis often present with vague symptoms and fever may be absent, obscuring the clinical diagnosis. Blood cultures and transesophageal echocardiography are key diagnostic tests. As a slow-growing, low virulent and common human skin germ, *P. acnes* can be wrongly considered as a blood culture contaminant.

Key words: Propionibacterium; artificial pacemaker; bacterial endocarditis; prosthesis related infection.

Abbreviations:

CIED: Cardiovascular implantable electronic device

CRP: C-Reactive Protein

CT: computed tomography

PCR: polymerase chain reaction

INTRODUCTION

Propionibacterium acnes is a slow growing anaerobic gram-positive bacillus that belongs to the human cutaneous flora. Despite its low virulence, it can cause various life-threatening infections including brain abcesses, spondylodiscitis, endophtalmitis and bone infections.¹ Infective endocarditis have been reported, typically late post-operative prosthetic valve infections.² When *P. acnes* is isolated in blood cultures, it can be wrongly regarded as a skin contaminant. We report a case that illustrates the diagnostic challenges of pacemaker-related endocarditis due to *P. acnes*.

CASE REPORT

A 74-year old man was admitted because of worsening fatigue for 15 days. He had a history of type 2 diabetes and hypertension, and a pacemaker was placed 2 years ago for high grade atrioventricular block. The temperature on admission was 35.6° C. Clinical examination was otherwise unremarkable. The white blood cell count was $15 \ 10^{9}$ /l, platelet count $40 \ 10^{9}$ /l and the C-Reactive Protein (CRP) concentration 109 mg/l. A whole body CT scan did not find any lesion but transthoracic echocardiography revealed a hyperechogenic mass floating in the right ventricle. Three samples were collected for blood culture but remained sterile. After 7 days, the patient left the hospital against medical advice.

Four days later he was admitted in our institution with delirium. His temperature was 35.4° C and he had a macular scattered purpuric eruption over the arms, legs and trunk. CRP was still high at 137 mg/l and white blood cell count at 11 10⁹/l, platelet count was stable at 39 10⁹/l and there was no disseminated intravascular coagulation. Bone marrow aspiration showed a normal number of megakaryocytes. A whole body CT scan was provisionally interpreted as normal except for several unremarkable pulmonary micronodules. The spinal fluid was normal.

In the next days, the purpura became palpable. One among six blood cultures revealed a Gram positive anaerobic bacillus at 93 hours of incubation, which was identified as *P. acnes*. The micronodules on chest CT were reinterpreted as septic pulmonary embolism. The skin

biopsy was also compatible with cutaneous septic embolisms. Transthoracic and transoesophageal echocardiography confirmed a large echogenic mass of 6×3 cm surrounding the lead of the pacemaker in the right atrium and prolapsing in the right ventricle during diastole (Figure 1A, Movies 1 and 2).

Still unsure if the *P. acnes* was the culprit or a skin contaminant, an antibiotic regimen active against *P. acnes* but also staphylococcal species was commenced (amoxicillin 12 g + clavulanic acid 600 mg + gentamicin 180 mg a day). The patient underwent cardiac surgery to remove the pacemaker with the vegetation on its leads (Figure 1B). The tricuspid valve was replaced by a biological prosthesis because of a perivalvular abcess found intraoperatively. An epicardial pace-maker was placed. Microscopy and culture of the vegetation were negative, but the 16S rRNA gene PCR amplification confirmed the presence of *P. acnes*. Antibiotics were changed for amoxicillin 12 g + rifampicin 1200 mg a day for 6 weeks. Five days after surgery, the platelet count increased to 300 $10^9/1$ and the purpura disappeared. The three blood cultures drawn at the first hospital and the five other blood culture drawn at our hospital before antibiotics onset remained sterile at four weeks.

DISCUSSION

Cardiovascular implantable electronic device (CIED) infections have been the subject of a recent scientific statement by the American Heart Association.³ CIED implantation rates, especially permanent pacemaker, have strongly increased over the past years and devices are more frequently placed in elderly and frail patients, ⁴ explaining higher rates of CIED infections. Pocket infections with or without associated bloodstream infection are more frequent than definite device infection (endocarditis). A recent study evaluated the rate of device-related endocarditis at 1.9/1000 device-years. ⁵ The main risk factors of device-related endocarditis are immunomodulatory therapy, chronic corticosteroid use, hemodialysis and presence of remote focus of primary infection. ⁶ About one third of device-related endocarditis occur more than 1 year after the last implantation procedure; clinical and microbiological features are similar in early- and late-onset endocarditis, with staphylococcal species causing up to 90% of cases. ⁷ Although device-related endocarditis may present without fever, biological markers of inflammation are almost always elevated. ⁷ As in right-sided endocarditis, pulmonary abnormalities are a clue to the diagnosis. The two most useful

diagnostic tests are blood cultures and transesophageal echocardiography, both with a sensitivity of 90 to 95%.^{3,7} Patients with suspected CIED should therefore have at least 2 sets of blood cultures before antibiotherapy is commenced. ³ However, falsely positive hemocultures due to skin contaminants are a well known issue and as much as 70% of lead-associated masses are non-infectious incidental findings. ⁸ Appropriate antibiotherapy and device removal are both indicated when device-related endocarditis is diagnosed.³

A recent review reports 58 detailed cases of endocarditis due to Propionibacterium species, including three associated to pacemakers.² This disease may be underdiagnosed and underreported because Propionibacterium species are often disregarded as skin contaminants when isolated in blood cultures and are sensible to empirical therapy with betalactams or glycopeptides. In this series, endocarditis due to Propionibacterium species involved a prosthetic valve in 67%, a permanent pacemaker or defibrillator in 7% and another prosthetic device in 5% of cases. The median delay between valvular replacement or device insertion and endocarditis was 4 years. *Propionibacterium* species are able to cause late post operative prosthetic valve endocarditis due to their ability to adhere to foreign body surfaces and produce a biofilm.^{9,10} They are low virulence bacteria, associated with minimal clinical and biological signs of infection. Their growth is slow and median time to positive blood or valve culture is 7 days (range 4-14); blood culture remain negative in almost 40% of cases.² Much of the antimicrobial regimen choices are extrapolated from in vitro or in vivo models of foreign body infections. Six weeks of a penicillin associated with a non-betalactam antibiotic should be prescribed. Aminoglycosids are possible choices but rifampicin and linezolid is increasingly used to take advantage of their biofilm penetrating ability.^{11,12} Second or third generation cephalosporins may be used in patients with reported penicillin allergy without severe or anaphylactic reactions; ¹³ combining rifampicin with linezolid is another option.¹²

CONCLUSION

Although uncommon, the diagnosis of *Propionibacterium* endocarditis should be considered when isolated in blood cultures even if the delay to positive culture is long and if only a few bottles are positive. The index of suspicion should be especially high in patients carrying a prosthetic valve or an implantable electronic device.

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Figure 1. A. Transesophageal echo (20°) showing a large echogenic mass surrounding the leads of the pacemaker in the right atrium (arrow). RA: right atrium, RV: right ventricle, LA: left atrium, LV: left ventricle. B. 6 x 3 cm vegetation on the pacemaker leads.

Movie 1: 2D Transesophageal echo (20°) showing a large echogenic mass surrounding the lead of the pacemaker in the right atrium and prolapsed largely in diastole in the right ventricle

Movie 2: 3D Transesophageal echo (20°) showing a large echogenic mass surrounding the lead of the pacemaker in the right atrium and prolapsed largely in diastole in the right ventricle

