An unexpected cause of a swollen pacemaker pocket: a case report

Bert Popelier 1*, Ruben Vanheste 2, Sofie Cuypers 1, and Ward Hegermont 1

1Cardiovascular Center, OLV Hospital, Moorselbaan 164, 9300 Aalst, Belgium; and 2Department of Radiology, OLV Hospital, Moorselbaan 164, 9300 Aalst, Belgium

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Background
A complication originating from the pacemaker pocket after device implantation can most often be explained by a post-operative pocket haematoma, or, less frequently, by a pocket infection. Both conditions need immediate assessment, dedicated treatment, and specialized follow-up. In rare cases, however, a swollen pacemaker pocket has an alternative diagnosis, which is exemplified by the following case.

Case summary
A 70-year-old male patient had non-specific symptoms of fatigue, dyspnoea, and coughing for some weeks. He also noted an evident, new swelling of his pacemaker pocket several months after pacemaker implantation, a procedure that was performed in a high-volume center and without any complication. Ultrasound imaging of the pocket suggested the presence of a soft tissue mass with increased vascularity, rather than a fluid collection or a late organized haematoma. Ultrasound-guided biopsy of the mass was obtained for histopathology analysis and revealed a well-differentiated invasive squamous cell carcinoma. Additional PET-CT imaging demonstrated multiple fluorodeoxyglucose-avid hotspots: a voluminous lesion in the left lung hilum, smaller lesions in the liver, some mediastinal lymph nodes, several bone lesions, and a large mass surrounding the pacemaker. The multidisciplinary oncologic specialty team concluded that the patient had an aggressive metastatic lung carcinoma. The patient refused to undergo further treatment and died 1.5 months after diagnosis.

Discussion
To the best of our knowledge, we did not find any earlier reports of a squamous cell carcinoma of the lung spreading to a pacemaker pocket. Presentation of a primary tumour or a metastasis in a pacemaker pocket is extremely rare. Ultrasound imaging with ultrasound-guided biopsy is a fast and reliable method to sample the tissue and to obtain a reliable diagnosis.

Keywords
Case report • Pocket haematoma • Pacemaker • Soft-tissue mass • Squamous cell carcinoma

ESC Curriculum
5.9 Pacemakers • 9.9 Cardiological consultations

Learning points
• When observing a soft-tissue mass in or around a pacemaker pocket several months after pacemaker implantation, a malignant process should be part of the differential diagnosis.
• The key to a fast and accurate diagnosis of a soft-tissue mass in or around a pacemaker pocket is ultrasound imaging and eventually ultrasound-guided tissue sampling.
• PET-CT imaging is of added value to detect the eventuality of a primary tumour and for staging of the oncologic disease.

* Corresponding author. Tel: +32 53 72 44 39, Email: bert.popelier@olvz-aalst.be
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Introduction

Implantation of a cardiac implantable device is a routine procedure that should go along with careful post-operative monitoring of the device pocket in search for possible complications. Among the most frequent post-operative complications are: pocket haematoma,1 pocket infection,1,2 and cardiac device-related infective endocarditis (CDRiE).2 Presentation of a primary tumour or a metastasis in a pacemaker pocket is rare, yet several case reports are available in the literature.3–17 Reporting these rare cases is particularly relevant to understand the metastatic behaviour of tumours and their predilection for prosthetic material.

Timeline

| Day 0 | Successful two-chamber pacemaker implantation |
|------|-----------------------------------------------|
| Day 1, 2 day 84 | Normal routine outpatient check-up with normal pacemaker function and clinically unremarkable pacemaker pocket |
| Day 272 | Routine outpatient check-up, patient reports no problems except a prominent mass in the pacemaker pocket, painless but annoying |
| Day 272 | Ultrasound imaging of soft-tissue mass, CT imaging, reveals soft-tissue mass around the pacemaker, haematoma or infection is excluded |
| Day 282 | Ultrasound-guided biopsy of soft-tissue mass is performed and sent for anatomopathological analysis |
| Day 289 | PET-CT imaging of total body reveals active oncologic process in the left lung hilum, the liver, mediastinal lymph nodes, bone lesions and a soft tissue mass around the pacemaker |
| Day 290 | Histopathological results of biopsy sample show evidence of a metastatic lesion of a squamous cell carcinoma |
| Day 302 | Multidisciplinary oncological assessment concludes that the patient suffers from an aggressive squamous cell carcinoma probably originating from the lung; patient refuses any form of anti-cancer treatment and consciously opts for palliative care |
| Day 318 | Patient deceased |

Case description

A 70-year-old male patient with a history of extensive nicotine abuse, laryngeal squamous cell carcinoma seven years ago, bladder urothelial carcinoma ten years ago, and a dual-chamber pacemaker (type Medtronic Azure XT W2DR01 MRI Surescan, an RV lead type Medtronic 5076 Capsure Fix Novus MRI 59 cm and an RA lead type Medtronic 5076 Capsure Fix Novus MRI 52 cm) implanted 9 months ago for sick sinus syndrome, was seen routinely in the outpatient cardiology clinic. He had complaints of persisting fatigue, dyspnoea, and coughing. He had been tested positive for COVID-19 infection three weeks earlier, but strangely his symptoms did not abate with time and were rather progressive. He also noted an evident, new swelling of his pacemaker pocket (Figure 1A and B). At the outpatient clinic, the patient had normal blood pressure, normal heart rate, no signs of cardiac decompensation or cachexia. Chest ultrasound suggested the presence of a soft tissue mass with increased vascularity, rather than a collection or a late haematoma (Figure 1C and D). The mass was also identified using conventional CT imaging (Figure 2A–C). Ultrasound-guided biopsy of the mass was obtained for histopathology and revealed a well differentiated invasive squamous cell carcinoma (Figure 3A–C). Additional PD-L1 staining was moderately positive (Figure 3D). PET-CT scan demonstrated FDG-avid lesions in the left lung hilum (Figure 2B–D, arrowhead) with multiple FDG-avid lesions present in the liver, as well as mediastinal lymph nodes and bone lesions. Furthermore, several soft tissue masses, among which a large mass surrounding the pacemaker pocket, were identified (Figure 2).

The patient was subsequently referred for an oncologic assessment, but he refused to undergo further treatment and chose for palliative care. He died 1.5 months after his cardiology outpatient consultation where the mass was first noticed.

Discussion

To the best of our knowledge, we did not find any earlier reports of a squamous cell carcinoma of the lung metastasizing to a pacemaker pocket. Based on the tissue sampling and the PET-CT images, the multidisciplinary oncology team decided that the most likely primary origin of the metastasis was the tumour in the left lung. There are case reports and small case series describing metastatic lesions of melanoma,3 malignant lesions in the proximity of breast carcinoma,4,5 sarcoma,6 rhabdomyosarcoma,7 cutaneous leiomyosarcoma,7 squamous cell and basocellular carcinoma of the skin,8,9 non-Hodgkin lymphoma,10 plasmablastic lymphoma,11 plasmacytoma,12 and multiple myeloma.13

The detection of an active oncologic process, a primary tumour or a metastasis in a pacemaker pocket is a rare phenomenon. Depending on the time frame, after implantation of the device, primary causes of pocket swelling are pocket haematoma, inflammatory fluid collection, and pocket infection. A rare presentation of a merely local pocket infection with important fluid collection but without systemic inflammation has been described.14 In the first place, ultrasound imaging of the swelling is the best tool to differentiate a soft-tissue mass from a blood collection. Whenever a soft-tissue mass is identified, an ultrasound-guided biopsy is a fast and reliable method to sample the tissue and a fast route towards a definite diagnosis. The result of the histopathology will then guide the oncologic staging process and the best additional examinations to be performed. In Figure 4, we propose a comprehensive diagnostic workup, but additional or different tests might be necessary based on the identification and characterization of the primary tumour.

Based on other case reports,5,16 the time frame between implantation of the device and occurrence of a tumour metastasis in this case is, rather short: in these other reports5,16 it took years for this phenomenon to take place. Another interesting question that remains unanswered, is precisely why this pacemaker pocket was a predilection site for early metastasis. Literature is rather scarce, however two cases of a squamous lung carcinoma spreading to sites of 'recent physical trauma' have been described.18
**Figure 1** Clinical appearance and ultrasound imaging of the pocket. (A) Anterior view of the macroscopic aspect of the swollen pocket, with patient in prone position. Remark the old, healed scar. (B) Profile view of the macroscopic aspect of the swollen pocket with the patient in supine position. (C) Colour-doppler ultrasound of the tissue mass surrounding the pacemaker. (D) Ultrasound imaging of the soft-tissue mass with a clear echogenic structure below which is the device battery.

**Figure 2** CT and PET-CT imaging of the soft-tissue mass and total body. (A) Anteroposterior CT image showing the pacemaker battery and the soft-tissue mass (arrow), and a left upper lobe tumour (arrowhead). (B) PET-CT image comparable to (A), showing the metabolically active soft-tissue mass in the pocket (arrow), the lung tumour (arrowhead) and several lesions in the liver. (C) CT image of the tissue mass surrounding the pacemaker (arrow). (D) PET-CT image of the lung tumour (arrowhead).
Figure 3  Histopathology imaging of the soft tissue. Cylindrical biopsy of the soft tissue, with haematoxylin/eosin staining at 20× (A), 100× (B), 200× (C). PD-L1 staining at 200× magnification is depicted in panel (D).

Figure 4  Proposed diagnostic work-up for pocket swelling, with an important role for the time frame between implantation of the device, and the diagnosis of the swelling. A stepwise approach is proposed, with an important role for ultrasound as the first-in-line diagnostic examination.
In the last decades, there were some reports on ‘oncotaxis’, the phenomenon describing the propensity of certain solid-tissue tumours to spread to zones of recent surgery (‘surgical oncotaxis’) or chronic inflammation (‘inflammatory oncotaxis’). Furthermore, it remains elusive whether the presence of titanium or other substances constituting a pacemaker battery, might have pro-oncogenic properties. As far as titanium is concerned, in vitro reports and experiments in human subjects have shown that titanium can induce chronic low-grade inflammation, e.g. by activating tissue macrophages overexpressing TNF-alpha. Potentially, this low-grade inflammation with an altered micro-environment in the pacemaker pocket is an attractive site for metastatic tumour cells.

Conclusions

When observing a soft-tissue mass in or around a pacemaker pocket months or years after pacemaker implantation, a malignant process should be part of the differential diagnosis. The key to a fast and accurate diagnosis is ultrasound imaging and eventually ultrasound-guided tissue sampling. PET-CT imaging is of added value to detect the eventuality of a primary tumour and for staging of the oncologic disease. Further research is warranted to understand which signals drive metastasis of an active cancer to a pacemaker pocket.

Lead author biography

Dr. Ward Heggermont is a board certified cardiologist, specialized in chronic heart failure, and device implantations, especially cardiac resynchronization therapy. He combines his clinical activities in the OLV Hospital Aalst (Belgium) with basic and translational research at the Cardiovascular Center Aalst (Belgium) and the Katholieke Universiteit Leuven.

Supplementary material

Supplementary material is available at European Heart Journal – Case Reports online.

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Slide sets: A fully edited slide set detailing these cases and suitable for local presentation is available online as Supplementary data.

Consent: The authors confirm that written consent for submission and publication of this case report including images and associated text has been obtained from the patient’s next-of-kin in line with COPE guidance.

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