



Nocardia farcinica arthritis of the knee A case report

Emmanuel AUDENAERT, Amar ALMAFRAGI, Marc VUYLSTEKE, Christine VERDONCKT, René VERDONK

From St-Andries Hospital, Tielt, Belgium

Nocardia is a Gram positive, aerobic branched actinomycete; it is an ubiquitous soil saprophyte. As an infecting agent it has been increasingly identified in humans, especially in immuno-suppressed hosts. *Nocardia* as a cause of septic arthritis is very unusual. The described genus was *Nocardia asteroides* in nearly all previously reported cases. We report an unusual case of spontaneous *Nocardia farcinica* septic arthritis of the left knee in a 68-year-old man, who was under systemic corticosteroids for chronic obstructive pulmonary disease. The diagnosis was rapidly made by Gram and acid-fast stains and later confirmed by culture. PCR was used to identify the subtype. The patient was treated successfully with sulphametaxazole-trimethoprim for six months. Our case re-emphasises, especially in the immunocompromised patient, the importance of performing fungal and acid-fast bacilli cultures, besides the most common bacterial cultures for aerobic and anaerobic organisms, in order to identify less common organisms and to initiate early and adequate treatment.

INTRODUCTION

Nocardiosis can present as a cutaneous, pulmonary, or disseminated disease and can follow an acute, subacute, or chronic course. The most frequently reported form is *Nocardia pneumonia* in an immunocompromised host. Disseminated nocardiosis may involve any organ, mainly the brain and meninges (6). The aetiological agent is an aerobic actinomycete, an ubiquitous saprophyte in soil,

decaying organic matter and water. At least 12 species of the *Nocardia* genus have been identified. *Nocardia asteroides* is the most frequent human pathogen. Human disease has also been caused by *Nocardia brasiliensis*, *Nocardia farcinica*, *Nocardia nova*, *Nocardia transvalensis* and *Nocardia otitidis calvarium* (6). Infection usually occurs through inhalation; primary cutaneous disease can start from soil-contaminated wounds. Rarely does surgical transmission occur.

Risk groups for the development of nocardiosis are severely immuno-compromised patients (malignancy, connective tissue diseases, bone marrow and organ transplanted patients, oral corticosteroid users, HIV-infected patients, alcoholism and pulmonary proteinosis) (8). The male : female ratio is 3:1 (4).

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- Emmanuel Audenaert, MD, Registrar,
 - Marc Vuylsteke, MD, Consultant Surgeon.

Department of General Surgery.

- Amar Almafragi, MD, Registrar.

Department of Internal Medicine.

- Christine Verdonckt, MD, Head of Department.

Department of Clinical biology, St-Andries Hospital, Krommewal 9-11, B-8700 Tielt, Belgium.

- René Verdonk, MD, PhD, Professor and Chairman.

Department of Orthopaedic Surgery, Gent University Hospital, De Pintelaan 185, B-9000 Gent, Belgium.

Correspondence : Marc Vuylsteke, Department of General Surgery, St-Andries Hospital, Krommewal 9-11, B-8700 Tielt, Belgium. E-mail : marc.vuylsteke@sintandriestielt.be.

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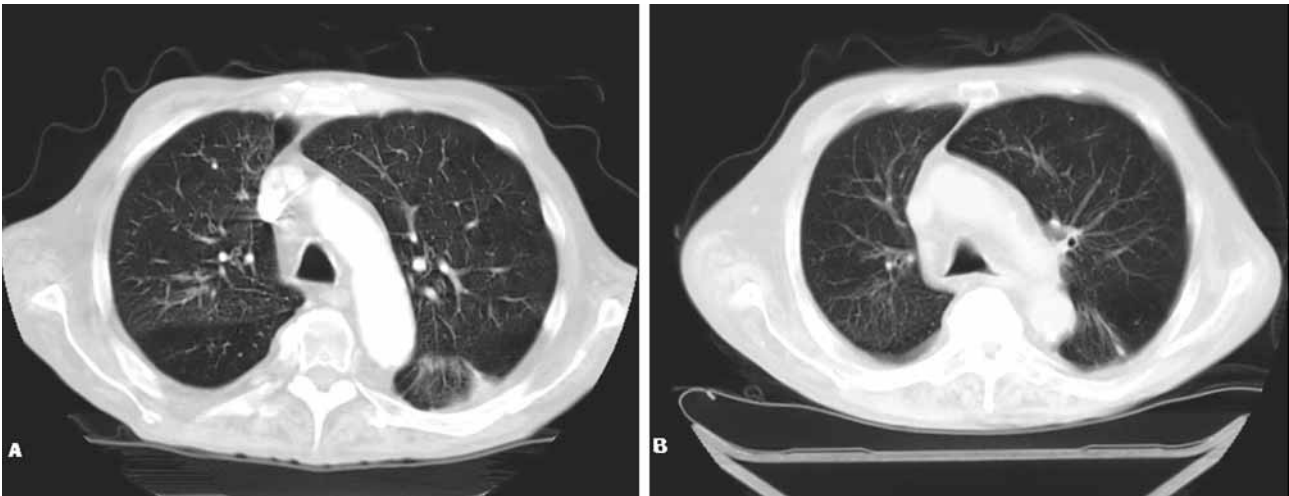


Fig. 1. — Sagittal CT-images of the lung before (A) and after six months of treatment (B) for nocardiosis due to *Nocardia farcinica*

Disseminated Nocardiosis presenting as septic arthritis has rarely been reported. In nearly all cases the described pathogen was *Nocardia asteroides*. We found no cases of *Nocardia farcinica* causing septic arthritis.

CASE REPORT

A 68-year-old man presented with deterioration of his general condition, shortness of breath, painful swelling of the left knee joint, tiredness, anorexia, nausea, vomiting and headache. The patient had an elevated temperature of 38.5° Celsius; other vital parameters were normal. Clinical examination of the left knee joint showed a painful intra-articular swelling, redness and elevated skin temperature. The range of motion was limited by pain and swelling. There was no history or clinical evidence of skin lesions or traumatic wounds. Lung auscultation revealed diffuse wheezes and bilateral basal crepitations.

The patient had a long medical history of uncontrolled type 2 diabetes mellitus, and chronic severe corticosteroid-dependent obstructive pulmonary disease (COPD) with progressive deterioration of respiratory function.

His main problem was recurrent respiratory infection with *Klebsiella* and *E. coli* (five times in one year) presenting with dyspnoea, fever, and

purulent sputum, necessitating repeated hospital admissions.

On this admission, haematological tests showed leucocytosis with neutrophilia and CRP elevation of 20.8 mg% (0.3). Chest radiographs and CT-scan of the thorax showed signs of interstitial lung disease with zones of panlobular emphysema mainly in the lower lobes. Pleuro-pulmonary fibrotic changes were present in the left upper lobe and interlobular apical septal fibrosis in the right upper lobe, consistent with pulmonary nocardiosis (fig 1a).

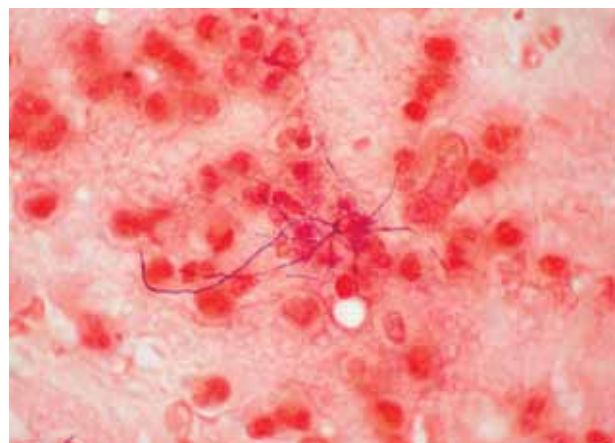


Fig. 2. — Gram-stain showing Gram-positive, branched bacterial filaments. The identification of the *Nocardia* was confirmed by culture.

Sputum examination and culture yielded *Escherichia coli* and *Pseudomonas aeruginosa*, sensitive to amikacin and ciprofloxacin.

Sonography and venous Doppler examination of the left lower limb disclosed a ruptured Baker cyst and left knee joint effusion. The left knee was punctured and a highly purulent fibro-viscous material was obtained, suggestive of septic arthritis. Gram and acid-fast staining were positive for gram-positive, branched bacterial filaments identified as *Nocardia* on culture (fig 2). The *Nocardia* was identified as *Nocardia farcinica* by sequence determination of the 16S rRNA gene.

Once the diagnosis of *Nocardia* had been made, the patient was treated with trimethoprim-sulphamethoxazole. Oral treatment was maintained for six months. During this period the patient showed progressive recovery of pulmonary function and subsidence of the septic arthritis. Chest radiograph and CT scan of the thorax after six months of treatment showed significant regression of the pleural and pulmonary fibrotic infiltration (fig 1b).

DISCUSSION

Nocardia species are organisms associated with environmental materials. They are ubiquitous saprophytes, including soil saprophytes, and only occasionally cause human disease, often in immunocompromised patients. The lung is the most commonly affected site (73%). Less frequently, the infection is disseminated, with an unknown entry portal. According to the literature these infections are usually of pulmonary origin (2,9).

Our patient was at specific risk for infection, because he was a diabetic, had chronic obstructive pulmonary disease, and was undergoing immunosuppressive treatment with corticosteroids.

The first stage in the diagnosis of nocardiosis involves direct examination of the specimen. Microscopic observation occasionally reveals the presence of Gram-positive, branched bacterial filaments. The development of *Nocardia* in culture media is rather slow; colonies are usually visible after 3 to 5 days as in our case, but the delay can be

as long as 2 to 3 weeks (7). PCR technique by sequence determination of the 16S rRNA gene demonstrated that the specific *Nocardia* type in our case was *Nocardia farcinica* (3).

With regard to the treatment of *Nocardia* arthritis, repeated thick-needle aspiration is the first line of management. Antibiotic treatment should not be started before the susceptibility pattern of the causal micro-organism is known from a diagnostic aspiration sample. In our case, the strain was sensitive to trimethoprim-sulphamethoxazole (1), which is the drug of choice for nocardiosis. The duration of treatment is 6 to 12 months. Minocycline is the best established alternative oral drug, amikacin the best alternative parenteral drug. Newer B-lactam antibiotics, including cefotaxime, ceftazidime, ceftriaxone and imipenem may be less effective in some cases caused by *Nocardia farcinica*.

REFERENCES

1. Berkey P, Moore D, Rolston K. In vitro susceptibilities of *Nocardia* species to newer antimicrobial agents. *Antimicrob Agents Chemotherapy* 1998; 32: 1078-1079.
2. Boiron P, Provost F, Chevrier G, Dupont B. Review of nocardial infections in France from 1987 to 1990. *Eur J Clin Microbiol Inf Dis* 1992; 11: 709-714.
3. Boiron P, Locci R, Goodfellow M. *Nocardia*, nocardiosis and mycetoma. *Med Myco* 1998; 36 suppl 1: 26-37.
4. Farina C, Borion P, Goglio A. Human nocardiosis in northern Italy from 1982 to 1992. Northern Italy Collaborative Group on Nocardiosis. *Scand J Infect Dis* 1995; 27: 23-27.
5. Laurent F, Provost F, Boiron P. Rapid identification of clinically relevant *Nocardia* species to genus level by 16S rRNA gene PCR. *J Clin Microbiol* 1999; 37: 99-102.
6. Lerner P. Nocardiosis. *Clin Infect Dis* 1996; 22: 891-903.
7. Roth A, Andrees S, Kroppenstedt RM, Harmsen D, Mauch H. Phylogeny of the genus *Nocardia* based on reassessed 16S rRNA gene sequences reveals underspeciation and division of strains classified as *Nocardia asteroides* into three established species and two unnamed taxa. *J Clin Microbiol* 2003; 41: 851-856.
8. Smego RA Jr, Moeller MB, Gallis HA. Trimethoprim-sulphamethoxazole therapy for *Nocardia* infections. *Arch Intern Med* 1983; 143: 711-718.
9. Uttamchandani RB, Diakos GL, Reyes RR. Nocardiosis in 30 patients with advanced human immunodeficiency virus infection: clinical features and outcome. *Clin Infect Dis* 1994; 18: 348-353.