



Case study

Disseminated *Mycobacterium abscessus* infection in a peritoneal dialysis patient



Vincent H.J.F. Mooren^{a,b,*}, Michiel W.P. Bleeker^a, Jakko van Ingen^c, Mirjam H.A. Hermans^d, Peter C. Wever^e

^a Department of Internal Medicine and Nephrology, Bernhoven Hospital, Uden, The Netherlands

^b Department of Internal Medicine, Radboud University Nijmegen Medical Centre, Nijmegen, The Netherlands

^c Department of Medical Microbiology, Radboud University Nijmegen Medical Centre, Nijmegen, The Netherlands

^d Molecular Diagnostics, Jeroen Bosch Hospital, 's-Hertogenbosch, The Netherlands

^e Department of Medical Microbiology and Infection Control, Jeroen Bosch Hospital, 's-Hertogenbosch, The Netherlands

ARTICLE INFO

Keywords:

Mycobacterium abscessus
Peritoneal dialysis
Disseminated infection
Exit-site infection

ABSTRACT

A disseminated peritoneal dialysis-related *Mycobacterium abscessus* infection is very rare. *M. abscessus* belongs to the rapidly growing mycobacteria and can be misidentified as a diphtheroid bacterium, which in our case delayed diagnosis and optimal treatment. Due to intrinsic resistance to most antimicrobials, therapeutic options in *M. abscessus* infections are limited. Infection often leads to catheter loss. A fatal outcome, like in our case, is not exceptional.

Introduction

Despite numerous preventive measures, exit-site infections (ESIs) are a common problem in the peritoneal dialysis (PD) population. ESIs usually respond well to empiric antimicrobial therapy aimed at Gram-positive and Gram-negative bacteria. However, if no clinical improvement occurs one should consider other infectious causes. On the basis of our case, we would like to address important features and pitfalls of PD-related *Mycobacterium abscessus* infections.

Case report

A 73-year-old Caucasian woman with end-stage renal disease who had been treated with continuous ambulatory peritoneal dialysis (CAPD) for 12 months presented on routine evaluation with local tenderness and erythema at the exit-site of the PD catheter. Her medical history was significant for a right-sided nephrectomy because of recurrent episodes of pyelonephritis due to nephrolithiasis, ischaemic heart disease, and two caesarean sections.

On physical examination there was no purulent discharge from the exit-site. The peritoneal fluid was clear and the patient had no fever. Patient history and physical examination revealed no further abnormalities. According to protocol she was treated as an outpatient for an exit-site infection (ESI) with oral flucloxacillin. After ten days of treatment there was no clinical improvement. As the exit-site culture

seemed to grow *Corynebacterium* species, therapy was switched to oral amoxicillin-clavulanic acid. On day 15, it was reported that the isolate was resistant to all tested antimicrobial agents including vancomycin. Since there was still no sign of clinical improvement the infected catheter was surgically removed on day 24 and in the same session a new catheter was inserted in a different abdominal region. She was switched to hemodialysis (HD) via a tunnelled catheter. On day 29, 16S ribosomal RNA polymerase chain reaction (16S rRNA PCR) identification performed on the presumed *Corynebacterium* isolate revealed 100% homology with *Mycobacterium chelonae-abscessus* complex. Further species identification by the National Institute for Public Health and the Environment (RIVM) revealed *M. abscessus*. As the PD catheter had been replaced, no further therapeutic actions were undertaken.

Ten days after surgical PD catheter-replacement the patient presented with fever and abdominal complaints. This time the new PD exit-site was inflamed. Laboratory studies showed an elevated C-reactive protein level: 130 mg/L (ref. < 6 mg/L) and white blood cell count: $12.7 \times 10^9/L$ (ref. $4.0\text{--}10.0 \times 10^9/L$). Ultrasonography showed no evidence for abdominal or tunnel abscesses. Flucloxacillin was commenced intravenously and the patient was admitted to the nephrology ward. Bacterial cultures taken from the exit-site, peripheral blood and jugular venous catheter remained negative. Despite prolonged antimicrobial therapy the patient had recurrent fever and chills, especially during HD sessions. While initial blood cultures obtained on admission and on day 5 remained negative, blood cultures obtained on day 11

* Corresponding author at: Department of Internal Medicine, Radboudumc, P.O. Box 9101, Route 463, 6500 HB, Nijmegen, The Netherlands.
E-mail address: vincent.mooren@radboudumc.nl (V.H.J.F. Mooren).

<http://dx.doi.org/10.1016/j.idcr.2017.05.001>

Received 22 April 2017; Received in revised form 2 May 2017; Accepted 2 May 2017

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Table 1
Previously reported cases of peritoneal dialysis peritonitis and exit-site infections by *Mycobacterium abscessus*.

Article (Ref)	Country or region	Number of PD-peritonitis cases	Number of ESI cases	Number of cases with bloodstream infection
Lo et al., Perit Dial Int 2013 [2]	Hong Kong	2	4	0
Renaud et al., Nephrology 2011 [3]	Singapore	4	3	1
Kameyama et al., Ther Apher Dial 2007 [4]	Japan	1	–	0
Ellis et al., Pediatr Nephrol 2005 [5]	USA	–	1	0
Tsai, Ther Apher Dial 2013 [6]	Taiwan	–	1	0
Yang et al., Perit Dial Int 2015 [7]	Taiwan	2	–	0
Jiang et al., Int Urol Nephrol 2013 [8]	Australia	3	–	0
Siddiqi et al., Saudi J Kidney Dis Transpl 2012 [9]	Saudi Arabia	2	–	0

Abbreviations: PD, peritoneal dialysis; ESI, exit-site infection.

after admission ultimately revealed growth of *M. abscessus*. Immediate removal of both the new PD catheter and the HD catheter was ordered. Flucloxacillin was discontinued and switched to tigecycline and clarithromycin. Unfortunately, bilateral thrombosis of the jugular veins occurred, which caused major access problems for adequate dialysis. The patient was, therefore, transferred to a nearby academic hospital. Broth microdilution susceptibility testing of the bloodstream isolate revealed amikacin susceptibility, intermediate imipenem susceptibility (minimum inhibitory concentration [MIC] 8 mg/L), inducible macrolide resistance and a tigecycline MIC of 0.25 mg/L [1]. Based hereupon, clarithromycin was switched to imipenem. Additional imaging studies showed no source of infection explaining the bacteraemia. Follow up blood cultures remained negative. The patient unexpectedly died because of ventricular fibrillation directly after insertion of a central venous catheter in the left subclavian vein.

Discussion

Among the rapidly growing nontuberculous mycobacteria (RGM), *Mycobacterium fortuitum*, *M. chelonae* and *M. abscessus* are considered clinically most relevant. They are ubiquitous in the environment, including water, soil and dust, and survive nutritional deprivation and extreme temperatures. The most common clinical manifestations comprise skin and soft tissue infections, catheter-related infections and pulmonary infections. RGM are infrequent causative agents of ESIs and PD-peritonitis; *M. abscessus* infections are particularly rare in this context with only eight previous reports comprising 23 cases (14 peritonitis; 9 ESIs) (Table 1) [2–9]. The vast majority of cases are described in Asian countries. To the best of our knowledge, this is the second case in literature of a *M. abscessus* bloodstream infection as a complication of PD-related infection [3].

Predisposing factors for development of PD-related infections with RGM include additional immunosuppressive conditions, recurrent episodes of ESIs or peritonitis with multiple courses of broad-spectrum antimicrobials, and underdialysis [5]. In addition, the use of topical gentamicin cream is postulated as a risk factor in two reports [2,10]. None of these applied to our patient.

The microbiological similarities of RGM and *Corynebacterium* species impose potential for misidentification in the laboratory. This phenomenon, which also happened in our case, has been previously reported [11,12]. In order to diagnose RGM infection, performance of auramine or Ziehl-Neelsen staining and mycobacterial culture are essential. An earlier auramine or Ziehl-Neelsen staining might, in our case, have led to earlier adequate antimicrobial therapy and possibly a better clinical outcome.

M. abscessus is intrinsically resistant to most antimicrobials [1]. Treatment consists of combination therapy, usually with cefoxitin or imipenem, amikacin and clarithromycin [1]. The minimum duration and optimal regimen for treatment for PD-related RGM infection are not known, but a prolonged course of antimicrobial treatment is recommended, especially when the plan is to salvage the PD catheter [13].

There is no guideline recommending catheter removal following diagnosis of PD-related RGM infection. High rates of catheter loss and mortality are reported in PD-related RGM disease [3]. Extensive comorbidities, as in our case, contribute to fatal outcomes [13].

Conclusion

We present a case with a disseminated PD-related *M. abscessus* infection. To prevent delay in optimal antimicrobial treatment, RGM infections should be considered when initial therapy does not lead to clinical improvement.

Funding

This research did not receive any specific grant from funding agencies in the public, commercial, or not-for-profit sectors.

Conflicts of interest

None.

References

- [1] Griffith DE, Aksamit T, Brown-Elliott BA, Catanzaro A, Daley C, Gordin F. An official ATS/IDSA statement: diagnosis, treatment, and prevention of nontuberculous mycobacterial diseases. *Am J Respir Crit Care Med* 2007;175:367–416.
- [2] Lo MW, Mak SK, Wong YY, Lo KC, Chan SF, Tong GM, et al. Atypical mycobacterial exit-site infection and peritonitis in peritoneal dialysis patients on prophylactic exit-site gentamicin cream. *Perit Dial Int* 2013;33(3):267–72.
- [3] Renaud CJ, Subramanian S, Tambyah PA, Lee EJ. The clinical course of rapidly growing nontuberculous mycobacterial peritoneal dialysis infections in Asians: a case series and literature review. *Nephrology* 2011;16(2):174–9.
- [4] Kameyama H, Mori Y, Kimura T, Sugishita C, Adachi T, Sonomura K, et al. A case report of *Mycobacterium abscessus* peritonitis in a peritoneal dialysis patient. *Ther Apher Dial* 2007;11(6):449–51.
- [5] Ellis EN, Schutze GE, Wheeler JG. Nontuberculous mycobacterial exit-site infection and abscess in a peritoneal dialysis patient. *Pediatr Nephrol* 2005;20(7):1016–8.
- [6] Tsai SF. Catheter related infection due to *Mycobacterium abscessus* in a patient under peritoneal dialysis. *Ther Apher Dial* 2013;17(3):349–50.
- [7] Yang TK, Lee JJ, Lu PL, Kuo HT, Chen HC. Peritoneal dialysis-associated peritonitis caused by *Mycobacterium abscessus*. *Perit Dial Int* 2015;35(3):369–71.
- [8] Jiang SH, Roberts DM, Clayton PA, Jardine M. Non-tuberculous mycobacterial PD peritonitis in Australia. *Int Urol Nephrol* 2013;45:1423–8.
- [9] Siddiqi N, Sheikh I. Peritonitis caused by *Mycobacterium abscessus* in patients on continuous ambulatory peritoneal dialysis. *Saudi J Kidney Dis Transplant* 2012;23(2):321–4.
- [10] Tse KC, Lui SL, Cheng VC, Yip TP, Lo WK. A cluster of rapidly growing mycobacterial peritoneal dialysis catheter exit-site infections. *Am J Kidney Dis* 2007;50(1):e1–5.
- [11] Williamson JC, Miano TA, Morgan MR, Palavecino EL. Fatal *Mycobacterium abscessus* endocarditis misidentified as *Corynebacterium* spp. *Scand J Infect Dis* 2010;42(3):222–4.
- [12] Larkin JA, Shashy RG, Gonzalez CA. Difficulties in differentiating a rapidly growing mycobacterial species from diphtheroids in an immunocompromised patient. *Clin Microbiol News* 1997;19(14):109–11.
- [13] Huang YC, Liu MF, Shen GH, Lin CF, Kao CC, Liu PY. Clinical outcome of *Mycobacterium abscessus* infection and antimicrobial susceptibility testing. *J Microbiol Immunol Infect* 2010;43(5):401–6.