Sepsis caused by *Myroides* spp. associated with bronchopneumonia and coinfection with *Pseudomonas aeruginosa* and *Proteus vulgaris*: A case report and review of *Myroides* spp. infections.

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Abstract

Background: *Myroides* spp. is considered an opportunistic germ, possibly pathogenic in immunocompromised patients, in whom it may cause varied clinical pictures in terms of severity, from skin and soft tissue infections, fasciitis, to urinary or respiratory infections, central nervous system infections, endocarditis etc. Few cases of sepsis caused by *Myroides* spp. have been described.

Case presentation: We present the case of an immunocompromised patient with diabetes mellitus, chronic renal failure, diagnosed with sepsis caused by *Myroides* spp. with a chronic leg ulcer as starting point, superinfected with *Pseudomonas aeruginosa* and *Proteus vulgaris*, and associating bronchopneumonia.

Conclusion: The presence of *Myroides* spp. or of other opportunistic pathogens in an immunocompromised patient, especially in those with renal failure, diabetes and persistent skin lesions, should be considered in cases with lack of response to commonly recommended antibiotics in these situations, as a possible emergence of an opportunistic pathogen that creates difficulties for treatment.

Keywords: Myroides spp., Sepsis, Bronchopneumonia, Co-infections, Biofilm.

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Introduction

Myroides spp. is an opportunistic infection-related pathogen in immunocompromised patients, most frequently being described in skin and soft tissue infections, rarely in pneumonia, bacteremia, venous-catheter related bacteremia, meningo-cerebral infections, endocarditis, and ventriculitis. The most common causes of associated immunosuppression are: diabetes mellitus, renal impairment, malignancy, and immunosuppressive therapy. Varied susceptibility to β -lactams, aminoglycosides, fluoroquinolones make it difficult to choose the first-line therapy. The association of an infection caused by *Myroides* with other bacteria in biofilm formation skin infections have not been described so far.

Case Report

We present the case of a 68 y old Caucasian male patient that has been hospitalized in the Cardiology Department for severe dyspnea and orthopnoea, chills, fever at home, coughing, and impaired general condition. Patient's medical history revealed anteroseptal myocardial infarction, atrial fibrillation, dilated cardiomyopathy, pulmonary hypertension, NYHA heart failure class IV, hypertension, insulin-dependent diabetes mellitus (diagnosed 20 y ago), diabetic nephropathy, chronic venous insufficiency CEAP C6. In the last two years, the patient had multiple admissions within the Dermatology Department for chronic bilateral leg ulceration, infected with a multidrug-resistant *Pseudomonas aeruginosa*, the last admission being 3 months prior to this episode.

On physical examination upon admission, the following could be noticed: fever 38.9°C, right shank ulcer of about 1/0.5 cm, left leg skin graft (performed 6 months ago), bilateral leg stasis dermatitis, body mass index 38.16 kg/m², emphysematous chest, diffusely disseminated crackles, SaO₂ 86%, Heart Rate (HR) of 130 beats per min (bpm), permanent atrial fibrillation, systolic murmur grade III/VI in all auscultation sites, Blood Pressure (BP) of 150/80 mmHg, hepatomegaly, and oliguria.

Laboratory investigations revealed the following alterations: C-reactive protein 303.06 mg/L (references values 0-5), fibrinogen 793.9 mg/dL (references values 170-420), erythrocyte sedimentation rate 48 mm/h (references values 0-15), white blood cells $15.14 \times 10^3/\mu$ l, neutrophils $12.23 \times 10^3/\mu$ l, platelets $438 \times 10^3/\mu$ l, blood urea nitrogen 100 mg/dL (references values 18-55), creatinine 1.80 mg/dL (references values 0,7-1-3), and blood glucose 124 mg/dl. Blood cultures

taken on admission, from a central venous catheter, were positive for Myroides spp. (cultures for aerobic bacteria, incubated in BacT/ALERT® 3D Microbial Detection Systems (bioMerieux, France). Direct Sensitivity Test (DST) in automatic VITEK® 2 Compact 15 system (bioMerieux, France) revealed susceptibility to cefepime, ciprofloxacin, imipenem, pefloxacin, meropenem, minocycline, and resistance to aztreonam, amikacin, tobramycin, gentamicin, piperacillin, piperacillin/tazobactam, ticarcillin, ceftazidime, colistin, and trimethoprim/sulfamethoxazole. The MICs (minimum inhibitory concentrations) were assessed according to the European Committee on Antimicrobial Susceptibility Testing breakpoints. Bacteriological examination of shank ulceration was positive for Proteus vulgaris, sensitive to amikacin, cefepime, ciprofloxacin, gentamicin, pefloxacin, tobramycin, trimethoprim/sulfamethoxazole, meropenem and Pseudomonas aeruginosa, sensitive to amikacin, colistin, gentamicin, imipenem, meropenem, tobramycin, resistant to cefepime, ceftazidime, ciprofloxacin, pefloxacin, piperacillin, ticarcillin, ticarcillin/clavulanic acid, and trimethoprim/ sulfamethoxazole.

Chest radiography was performed and revealed the presence of numerous inhomogeneous opacities in both lungs, and minimal basal bilateral pleural effusions. Also, a chest computed tomography scan was performed, that concluded with the diagnosis of bronchopneumonia. The transthoracic echocardiogram revealed moderate mitral regurgitation and moderate tricuspid insufficiency, with circumferential pericardial fluid with a diameter of 12 mm (Figure 1).

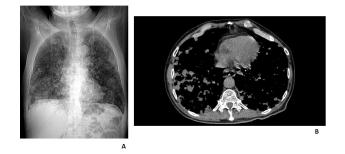


Figure 1. CT scan images (A: CT scout view. B: CT axial view).

Clinical data in conjunction with laboratory examinations supported the final diagnosis of sepsis caused by *Myroides* spp., bronchopneumonia, dilated cardiomyopathy, permanent atrial fibrillation, moderate pericarditis, moderate tricuspid insufficiency, moderate mitral regurgitation, NYHA heart failure class IV, stage 3 chronic kidney disease, insulindependent type 2 diabetes mellitus, leg ulcers superinfected with *Pseudomonas aeruginosa* and *Proteus vulgaris*.

The evolution was favourable under treatment with meropenem 2 g/d for 12 d, started on the 4^{th} d of hospitalization, and replaced 4 d after with ceftazidime 3 g/d, after obtaining the results of sensitivity testing, associated to the treatment for the underlying diseases: digoxin, furosemide, spironolactone, acenocoumarol, and rimenidine. On discharge,

the recommendation was to continue treatment with ciprofloxacin 2 \times 250 mg/d, for another 7 d.

Discussions

Myroides genus was known as *Flavobacterium odoratum* and belongs to the family Flavobacteriaceae [1], tribe Chromobacteriaceae. It is represented by non-fermentative gram-negative bacilli, non-motile, catalase-positive, oxidase-positive, yellow-pigmented because of the presence of flexirubin pigment [2] and fruity odor. Phenotypically, there are two different species, *M. odoratus* (which does not grow on MacConkey agar culture medium) and *M. odoratimimus*. In recent years, other 3 species were isolated from seawater [3-5], respectively *M. marinus*, *M. profundi* and *M. pelagicus*. It is considered that infection in humans is due to the takeover of bacteria from the environment, contaminated soil or water; in humans, *Myroides* spp. was isolated from blood, stool, urine, infected wounds; sometimes its role in pathogenesis is not very clear.

In immunocompromised patients, *Myroides* may be responsible for skin infections in patients with diabetes [6,7], urinary tract infections, lung infections [8-10], endocarditis [11], ventriculitis [12,13], bacteremia [14] or infections of the central nervous system. Rarely, there have been reported cases of infection in immunocompetent hosts [12,13]; the described cases were associated with hepatic, renal insufficiency, malignancies or co-infections with *Oerskovia turbata* in alithiasic cholecystitis [15].

So far, there has not been described any case of co-infections with other gram-negative bacteria, such as the case that we presented, which we believe that Myroides co-aggregated at the level of leg ulcer together with Proteus vulgaris and Pseudomonas aeruginosa to produce biofilm, allowing the multiplication and hematogenous dissemination of Myroides spp. The absence of the germ, from the bacteriological examinations of leg ulcers, does not exclude this entrance gate for bacteremia, the samples that were harvest on admissions were grown on usual growth medium. A growing on MacConkey agar medium was not used. Identification in a blood culture of an opportunist germ, commonly absent on the skin or mucous membranes, is sufficient to discuss about bacteremia with this germ. The patient had multiple admissions, which explains antibiotic resistance of Pseudomonas aeruginosa and Proteus vulgaris isolates. However, isolated Myroides strain was sensitive to carbapenems, fluoroquinolones, minocycline and cefepime, compared to other cases cited in the literature, that generally recorded resistance to β -lactams through chromosomally encoded the subclass B1 of metallo-\beta-lactamases [16] or variable susceptibility to aminoglycosides or quinolones [8,12,17-20]. Common susceptibility of isolated strains to meropenem allowed monotherapy in doses appropriate to renal impairment, recommending at discharge another 7 d of ciprofloxacin. Myroides susceptibility to antibiotics suggests a contamination subsequent to the last hospitalization, from the environment.

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Conclusion

The presence of *Myroides* spp. in immunocompromised patients, especially in those with renal failure, diabetes and persistent skin lesions, should be considered, and also other opportunistic pathogens, in cases with lack of response to commonly recommended antibiotics in these situations, as a possible emergence of an opportunistic germ that creates difficulties for treatment.

Ethics Approval and Consent to Participate

Written informed consent was obtained from the patient for publication of this case report and any accompanying images. The study was accepted by the Ethics Committee of the hospital and they encouraged publishing the article. A copy of the written consent is available for review by the Editor-in-Chief of this journal.

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