Case Report

Catheter-Related Sepsis by *Candida pararugosa* in an Adult Patient under Chemotherapy Regimen

Gabriella Piatti¹,², Sara Feltrin³, Emanuela Fellini³, Valentina Barbero¹, and Alberto Ballestrero¹,³

¹Ospedale Policlinico San Martino, 10 Largo Benzi, Genoa 16132, Italy
²Department of Surgical and Diagnostic Sciences (DISC), Section of Microbiology, University of Genoa, 8 Largo Benzi, Genoa 16132, Italy
³Department of Internal Medicine and Medical Specialties (DiMI), University of Genoa, 8 Largo Benzi, Genoa 16132, Italy

Correspondence should be addressed to Gabriella Piatti; piatti@unige.it

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*Candida pararugosa* is present in animals and humans in different organs and biological liquids, usually as a saprophyte. We report the case of a 61-year-old woman diagnosed with *de novo* stage IV metastatic lobular breast cancer, carrying a central venous catheter (port-a-cath) and bilateral stents for perirenal infiltration by malignancy. During chemotherapy regimen, a febrile episode occurred, along with a high level of serum glucan. The port-a-cath was removed after blood collection for culture, which gave isolation of *Candida pararugosa* strains. Given high glucan level and the patient’s frailty, empirical treatment with fluconazole was started with load-dose, 800 mg orally, at day 1 and, afterwards, with 400 mg daily for two weeks. The phenotype of susceptibility to antibiotics of the strain demonstrated lower minimal inhibitory concentration to fluconazole than that reported in the literature. The patient remained asymptomatic, and inflammation parameters showed normalization. Unfortunately, three weeks later, meningeal localization of cancer caused rapid deterioration and death.

1. Introduction

*Candida pararugosa* are ascomycetes first isolated from human feces in 1978 [1]. Later, it was found in vegetable food [2–5] and anatomic sites of animals and humans [6, 7], where it is usually a harmless commensal organism and rarely involved in infections. A persistent detection of *C. pararugosa* from saliva of a two-year-old girl affected by rhabdomyosarcoma and relatives, and from the oral cavity of a healthy asymptomatic subject, was reported in 2004 [8, 9]. Recently, *C. pararugosa* was isolated from exterior infections, i.e., from onychomycosis [10]. The yeast was isolated as a cause of sepsis in Brazil and Qatar, respectively, in a child and in a newborn, in 2014 [11, 12]. Only one case report describes *C. pararugosa* isolated from blood and cause of sepsis in adult [13].

Since definition of *Candida* genus is highly polyphyletic and based on artificial morphologies, some species were moved to different genera, leading to a clinical linkage between phylogenetic placement and antifungal intrinsic resistance [7]. Interestingly, *C. pararugosa* belongs to fungi grouped as “rare yeasts” that, indeed, occur in low prevalence and express elevated minimal inhibitory concentrations (MICs) for at least one class of antifungals. These features make the microorganism worthy of particular attention, given the scarce clinical expertise and the difficulty of treatment [7].

2. Case Report

The patient was a woman in her sixties diagnosed with lobular breast cancer about six months earlier. She presented
carcinomatous mastitis with lymph nodes infiltration of ipsilateral axillary region, widespread bone and liver metastasis, and perirenal infiltration by malignancy. Tru-Cut of breast lump demonstrated hormone-sensitive invasive lobular carcinoma. Immunohistochemistry assay showed ER = 90%, PR = 60%, Ki-67 = 75% Her2 neg (score 1+). The patient started first-line hormone-therapy with CDK 4/6 inhibitor (ribociclib) plus aromatase inhibitor (letrozole). In spite of two months therapy, the visceral disease progression and bilateral hydrenephrosis necessitated the placement of bilateral stents and the beginning of first-line poly-chemotherapy with cyclophosphamide-epirubicin (EC75 q21) regimen and parental nutrition, after positioning of the central venous catheter (port-a-cath).

One week after the second of six courses of chemotherapy, the patient developed fever without apparent foci of infection. Laboratory blood tests showed normal leukocyte count (5.10/μm³) and low hemoglobin (82.0 g/L), while C-reactive protein (CRP) was 43.6 mg/L and erythrocyte sedimentation rate (ESR) was 109 mm/h. Central and peripheral blood cultures were repeatedly negative. Empirical antibiotic therapy with ceftiraxone and levofloxacin was undertaken, and the clinical response occurred after five days of treatment. In the suspicion of endocarditis, we performed echocardiography, which was negative for vegetation. Eleven days after the first febrile episode, the patient developed fever with normal leukocyte count (6.5/μm³), CRP of 20.5 mg/L, and serum glucan level of 128 pg/mL and the same of 183 pg/mL six days later. Consequently, bilateral stents, in spite of negativity of urine, were substituted and the same of 183 pg/mL six days later. 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Consequently, bilateral stents, in spite of negativ
In our report, the patient presented stage IV metastatic lobular breast cancer. She was under chemotherapy treatment, was carrying a port-a-cath central venous catheter and bilateral urinary stents, was fed with parenteral nutrition, and had been recently exposed to broad-spectrum antibacterial therapy. All these are clinical aspects worthy of being considered as predisposing factors for candidemia [6]. The lack of leukopenia makes it difficult to correlate the infection with chemotherapy, which can affect the bowel mucosa and favor invasion by the yeasts. Bilateral stents were replaced after detection of elevated serum glucan levels without being cultured. The urine was negative as was the search for C. pararugosa in patient’s feces. Although the infection itself lasted a very short period and isolation of C. pararugosa occurred only from the central venous catheter, the latter was confirmed in the clinical relevance by high glucan levels. The port-a-cath had been placed sixty-six days prior to the second febrile episode, and its removal coincided with the start of therapy and resolution of fever. These circumstances made us think of a single day of fluconazole treatment as a chance event in relation to clinical resolution. Like other Candida species, C. pararugosa produces biofilms, especially in prothetic materials [16]. Therefore, we think that the removal of the central catheter, along with possible biofilm, was instrumental in resolution of the infection, maintained with continuance of antifungal therapy. From the microbiological point of view, our report confirms the inability of biochemical testing to identify pararugosa species and the need to utilize molecular methods or spectrometry. Instead, we repeatedly obtained MIC values for both azole and echinocandin different from those described in the literature. In particular, MIC for fluconazole was lower than that obtained by Pérez-Hansen et al. and similar to that obtained by Paredes et al. [7, 15] and was supported by the persistence of the response to treatment.

Ethical Approval

This study adhered to ethical requirements.

Consent

Written informed consent for publication of clinical details was obtained from the patient.

Conflicts of Interest

The authors declare no conflicts of interest.

Authors’ Contributions

All authors had access to the data and contributed to writing the manuscript.

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