## MYCOPATHOLOGIA IMAGES



## A Large Deep Skin Ulcer as an Initial Manifestation of Systemic Cryptococcosis

Received: 16 January 2019/Accepted: 22 January 2019/Published online: 1 February 2019 © Springer Nature B.V. 2019

**Abstract** An 82-year-old woman presented to our hospital with a deep skin ulcer in her right lower limb. Although the skin biopsy showed necrosis and neutrophil infiltration, we could not initially detect any pathogen. Chest radiography showed multiple nodules despite the lack of respiratory symptoms or fever, and the serum latex agglutination test for cryptococcus showed an elevated titer (1:512). Considering these findings, we performed additional stains of periodic acid-Schiff reaction and Grocott to the skin-biopsy specimen and detected multiple yeast-like fungi. The cultures of the skin and lung-biopsy specimens revealed Cryptococcus neoformans. In this case, it is suggested that a large deep skin ulcer can be an initial manifestation of systemic cryptococcosis, as, in some cases, pulmonary cryptococcosis may be asymptomatic. Second, cutaneous cryptococcosis of an unexposed area such as the thigh can contribute to the diagnosis of systemic cryptococcal infection because it is attributable to bloodstream dissemination from other organs.

**Keywords** Skin ulcer · Systemic cryptococcosis · *Cryptococcus neoformans* · Glucocorticoids

An 82-year-old woman presented to our hospital with refractory skin ulcers in her right lower limb (Fig. 1), which had appeared approximately 3 months earlier and progressively worsened. She had been treated with glucocorticoids for 2 years under a diagnosis of remitting seronegative symmetrical synovitis with pitting edema syndrome.

Although the skin biopsy showed necrosis and neutrophil infiltration, we could not initially detect any pathogen. Chest radiography (Fig. 2) showed multiple nodules despite the lack of respiratory symptoms or fever, and the serum latex agglutination test for cryptococcus showed an elevated titer (1:512). Considering these findings, we performed additional stains of periodic acid–Schiff reaction and Grocott to the skin-biopsy specimen and detected multiple yeast-like fungi (Fig. 3). The cultures of the skin- and lung-biopsy specimens revealed *Cryptococcus neoformans*.

After we confirmed that the blood culture was negative for growth and the cerebrospinal fluid examination was normal, we finally diagnosed the patient with systemic Cryptococcus infection of the skin and lung. Her condition improved with intravenous liposomal amphotericin-B and oral flucytosine treatment for 2 weeks followed by oral fluconazole (Fig. 4).





Fig. 1 Patient's thigh with refractory skin ulcers



Fig. 2 Chest X-ray showing multiple nodules in the right lung



Fig. 4 Patient's skin ulcers improved after antifungal drug treatment

Two important observations can be made based on this case. First, a large deep skin ulcer can be an initial manifestation of systemic cryptococcosis, as, in some cases, pulmonary cryptococcosis may be asymptomatic. Second, cutaneous cryptococcosis of an unexposed area such as the thigh can contribute to the diagnosis of systemic cryptococcal infection because it is attributable to bloodstream dissemination from other organs (e.g., the lungs).

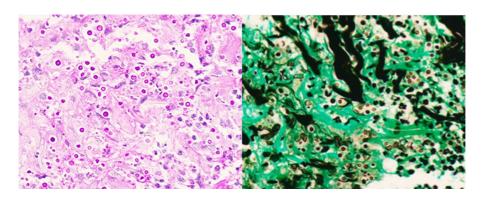


Fig. 3 Stains of periodic acid-Schiff reaction (left) and Grocott (right) to the skin-biopsy specimen



## **Compliance with Ethical Standards**

**Conflict of interest** The authors declare that they have no conflict of interest.

**Informed Consent** Informed consent was obtained from all individual participants included in the study.

**Publisher's Note** Springer Nature remains neutral with regard to jurisdictional claims in published maps and institutional affiliations.

<sup>1</sup>Kei Araki
<sup>1</sup>Yusuke Yoshida
<sup>1</sup>Hiroki Kohno
<sup>1</sup>Eiji Sugiyama

<sup>1</sup>Department of Clinical Immunology and Rheumatology, Hiroshima University Hospital, 1-2-3, Kasumi, Minami, Hiroshima 734-8551, Japan e-mail: fisher37@hiroshima-u.ac.jp

