Nocardial Cerebral Abscess Associated with Mycetoma, Pneumonia, and Membranoproliferative Glomerulonephritis

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Nocardial brain abscesses remain a clinical challenge. We successfully treated a patient with nocardial brain abscess, mycetoma, pneumonia, and membranoproliferative glomerulonephritis. Nocardial soft tissue involvement, mycetoma, is well known. However, the fact that actinomycetoma can metastasize may not be as well appreciated. The association between nocardiosis and membranoproliferative glomerulonephritis should be better clarified.

CASE REPORT

A 49-year-old male, a businessman, was admitted to our hospital with complaints of severe edema of the lower extremities. Clinically, the diagnosis was nephrotic syndrome. Light microscopy of a renal biopsy showed fibrosis with mesangial hypercellularity and tubular atrophy. Immunofluorescence microscopy of a renal biopsy showed fibrosis with mesangial intiities. Clinically, the diagnosis was nephrotic syndrome. Light microscopy of a renal biopsy showed fibrosis with mesangial hypercellularity and tubular atrophy. Immunofluorescence microscopy of a renal biopsy showed fibrosis with mesangial intiities. Clinically, the diagnosis was nephrotic syndrome. Light microscopy of a renal biopsy showed fibrosis with mesangial hypercellularity and tubular atrophy. Immunofluorescence microscopy of a renal biopsy showed fibrosis with mesangial intiities. Clinically, the diagnosis was nephrotic syndrome. Light microscopy of a renal biopsy showed fibrosis with mesangial hypercellularity and tubular atrophy. Immunofluorescence microscopy of a renal biopsy showed fibrosis with mesangial intiities. Clinically, the diagnosis was nephrotic syndrome. Light microscopy of a renal biopsy showed fibrosis with mesangial hypercellularity and tubular atrophy.

When the patient was hospitalized for his third course of cyclophosphamide therapy 9 months later, he was febrile, and physical examination revealed a tender mass in the anterolateral region of the left thigh. Laboratory findings revealed the following: white blood cell count, 13,360/mm³ (normal range, 4 × 10⁹ to 10 × 10⁹/liter); hemoglobin, 10.4 g/dl (normal range, 11 to 16 g/dl); hematocrit, 32.1% (normal range, 37 to 50%); erythrocyte sedimentation rate, 63 mm/h (normal range, <25 mm/h); cyclic AMP receptor protein, 16.2 mg/dl (normal range, <0.8 mg/dl); urea, 35 mg/dl (normal range, 10 to 50 mg/dl); creatinine, 0.8 mg/dl (normal range, 0.7 to 1.2 mg/dl); aspartate aminotransferase, 68 U/liter (normal range, 14 to 36 U/liter); and alanine aminotransferase, 116/liter (normal range, 9 to 52 U/liter). There was 5 g/day of proteinuria. Magnetic resonance imaging of the left lower extremity showed a mass lesion within the vastus lateralis muscle (Fig. 1). The mass had cystic characteristics and was multiloculated. The magnetic resonance imaging scan of the thorax revealed areas of consolidation and aterectasia in the basal segments of the left lung and centrally necrotizing lymphadenopathy in the left para-aortic region. Since there was no adequate response to medical therapy with TMP/SMX for 2 months, the patient underwent a left tempo-occipital craniotomy and the abscess was excised.

The patient was diagnosed as having a mycetoma, and antibiotic therapy was started with orally administered trimethoprim-sulfamethoxazole (TMP/SMX) and intravenously administered cephazolin. The following day, the patient had a seizure and developed amnesia. Cranial magnetic resonance imaging studies were consistent with abscess formation in the left tempo-occipital region (Fig. 2). Because of suspicion of multisystem nocardiosis, the patient underwent computed tomography scanning of the thorax. It revealed areas of consolidation and aterectasia in the basal segments of the left lung and centrally necrotizing lymphadenopathy in the left para-aortic region. Since there was no adequate response to medical therapy with TMP/SMX for 2 months, the patient underwent a left tempo-occipital craniotomy and the abscess was excised.

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with its capsule (Fig. 3). Using the same isolation methods as those mentioned above (including the smears), the *Nocardia asteroides* complex was again identified in cultures of the drainage specimen obtained from the cerebral abscess excised on 26 July 2004. After the surgery, the patient recovered progressively. At the fourth month after the surgery, medical therapy with TMP/SMX continued with a dose of 10 mg/kg trimethoprim per day, and the patient had no complaints. Laboratory findings remained normal. Postoperative cranial magnetic resonance images showed no abscess in the brain.

**Discussion.** Nocardiae are gram-positive, aerobic actinomycetes found naturally in the soil, air, and sewage (1, 12). Members of the *Nocardia asteroides* complex account for 80% of cases of noncutaneous invasive nocardiosis and most systemic or central nervous system nocardiosis (7). Nocardial brain abscesses remain a clinical challenge associated with high mortality and morbidity rates (2, 7, 10, 12, 14).

The patient was a businessman, and hence inoculation of the nocardia could not be directly related to his occupation. It is generally accepted that the primary infection by nocardiae is most commonly acquired via the respiratory tract (7). Primary pulmonary lesions may be subclinical or overt, chronic, or, rarely, acute (3). *Nocardia asteroides* may cause mycetomas. Mycetoma is a chronic granulomatous process resulting from the implantation of one of various fungi or actinomycetes into soft tissues, usually of the foot (8, 11). The disease is almost painless, and there are few or no constitutional symptoms unless there is a secondary bacterial infection (8). The microbial colonies form grains and sinuses in soft tissues. A literature survey revealed that the flora causing mycetoma in Turkey consists of various pathogens, such as *Nocardia asteroides*, *Nocardia brasiliensis*, *Scedosporium apiospermum*, *Aspergillus niger*, and *Streptomyces madurae* (4).

Nocardiosis is an opportunistic infection that has been noted in patients with malignancies, systemic lupus erythematosus, or HIV infection and in long-term steroid users and transplant recipients (2, 6, 9). However, recent data demonstrate that central nervous system nocardiosis is as common or more prevalent in patients with normal immune systems (12). Therefore, the nocardiae may have a predilection for some, but not all, immunocompromised patients. For example, most transplant patients reported to have nocardiosis are renal-transplant recipients. In 1998, Jose et al. reported an association between mesangiocapillary glomerulonephritis and nocardiosis (5).
There are no similarities between our patient and their patient. There were no primary immune deficiency disorders in either case. Both patients had glomerulonephritis and received corticosteroids, although the patient in the case report of Jose et al. received up to 100 mg daily of prednisolone for 6 years because of cluster headaches, not for glomerulonephritis. In our case, it is unclear whether there was a predisposition to nocardia infection directly linked to the glomerulonephritis or if it was a secondary consequence of prolonged immunosuppression due to treatment with prednisolone (60 mg daily) and cyclophosphamide for 9 months. This therapy could have aggravated a silent chronic infection in the soft tissue of the leg, a mycetoma by Nocardia asteroides. In the literature, the average duration of the symptoms in mycetomas has been calculated as 8 years (8). However, in our case, lesions in the lungs and in the brain were revealed in the relatively short time period after the symptoms of mycetoma appeared. Also, Jose et al. made the diagnoses of glomerulonephritis and nocardiosis in their case nearly simultaneously. Therefore, there may be a link between glomerulonephritis and nocardiosis, since the diagnoses of these clinical entities could be made sequentially in a relatively short time in both cases.

In our case, the surgical excision of the nocardial brain abscess was successful. In 2002, Lee et al. recommended that nocardial cerebral abscesses be initially treated via aspiration, with aggressive surgical management being reserved for the small proportion of patients who do not respond to minimally invasive surgery (7). However, only one patient in their series was discharged in good condition after a single aspiration procedure. In their other patients, more than one aspiration or excision was less than half (50%) among patients undergoing aspiration and drainage alone (10).

Mycetomas may be confused with granulomas, abscesses, or benign soft-tissue tumors, and this may result in delayed diagnosis of a possible nocardial infection. It is well understood that nocardial mycetoma may lead to new lesions in vital organs, such as the lungs and brain. However, the fact that actinomyctoma can metastasize may not be as well appreciated. Additionally, the relationship between nocardiosis and glomerulonephritis should not be regarded solely as an effect of immunosuppression but should also be clarified in terms of specific pathophysiological mechanisms. Surgical excision of nocardial cerebral abscesses instead of an aspiration procedure may be more beneficial in some patients.

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REFERENCES