Cerebral nocardiosis

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ABSTRACT

Cerebral Nocardiosis is a rare, challenging, opportunistic infectious disease of the central nervous system occurring in both immunocompetent and immunocompromised hosts. It often results in intraparenchymal abscess formation, which represents only 2% of all cerebral abscesses. The diagnosis of cerebral Nocardiosis is seldom based on imaging. Bacteriological diagnosis is often reached only after surgical excision of the abscess. We report a rare case of brain abscess caused by Nocardia species in a 20-year-old immunocompromised lady. Total surgical excision of the abscess, prompt bacteriological diagnosis based on smear and culture of the pus and initiation of specific antimicrobial therapy (trimethoprim and sulfamethoxazole) resulted in good clinical outcome.

Key words: Brain abscess, cerebral nocardiosis, Nocardia

Introduction

Brain abscess is one of the several forms of intracranial infection that is though uncommon, can be life-threatening. With the advent of better imaging modalities and effective antimicrobial therapy, there has been a trend toward reduced mortality. Brain abscess in immunocompetent patients are polymicrobial due to aerobic and anaerobic bacteria. However, widespread use of antibiotics has resulted in brain abscess caused by atypical organisms like Nocardia species.

Case Report

A 20-year-old female presented with gradual onset progressive weakness in her right upper and lower limbs of 3 weeks duration. She also had persistent holocranial headache associated with multiple episodes of vomiting since 1-week. She did not have convulsions or fever during the course of her illness. Three months prior to the onset of the present illness, she was diagnosed to have pulmonary tuberculosis and was on intensive regimen of antituberculous therapy (Isoniazid, Rifampcin, Ethambutol and Pyrizinamide). On clinical examination, her pulse was 56/min. She had bilateral sixth nerve palsy and right upper motor neuron facial palsy. Her fundus examination revealed gross papilledema. She had right spastic hemiparesis with power grade 3/5 and 2/5 in upper and lower limbs respectively. Right plantar response was extensor. Magnetic resonance imaging (MRI) of the brain revealed multiple multiloculated lesions in the left medial frontal lobe. Largest of the lesions measured 4 cm in diameter with multiple smaller lesions located posteriorly. The lesions demonstrated central hypointense signals on T1-weighted imaging (T1-WI) with corresponding hyperintense signals on T2-WI [Figure 1a and 1b]. The peripheral rim revealed iso-intense signals on T1-WI and hypointense signals on T2-WI [Figure 1a and 1b]. On gadolinium contrast administration, thick, smooth enhancement of the peripheral rim was noted [Figure 1c and 1d]. Diffusion-weighted imaging revealed hyperintense signal within the lesions [Figure 2a] with corresponding apparent diffusion coefficient revealing hypointense signals [Figure 2b] suggestive of restricted diffusion. Fluid attenuated inversion recovery sequence revealed hyperintense signals adjacent to the lesions suggestive of vasogenic edema [Figure 2c]. Chest ray revealed right apical infiltrates. Laboratory workup showed the leucocytosis (white blood cell 13000/cmm) and elevated sedimentation rate (70 mm/h in first 1 h). The presumptive imaging diagnosis was multiple tubercular abscesses. At surgery multiloculated, thick walled abscess was encountered in the left medial frontal region. Aspiration of the largest abscess cavity revealed non-foul smelling pus. Total excision of the abscess was performed. Aspirated pus was sent for g stain, aerobic, anaerobic and tubercular bacilli culture. Examination of pus smear revealed g positive...
branching filaments. The bacilli were partially acid alcohol fast suggestive of *Nocardia* species [Figure 3a]. Growth was obtained on blood agar and Lowenstein Jensen media after 96 h of incubation [Figure 3b]. Smears from culture also revealed g positive filaments which were partially acid fast. Biochemical reactions were performed which revealed catalase positive, urease positive, oxidase test negative and reduction of nitrates to nitrites. Blood culture remained sterile. Sputum culture did not reveal growth of *Nocardia* or tuberculous bacilli. Postoperatively, she received broad spectrum empirical antibiotic therapy and antituberculous drugs until culture reports were available. Following the growth of *Nocardia* on culture, she was started on trimethoprim/sulphamethoxazole (TMP/SMX) (960/4800 mg) daily. In the postoperative period, her right hemiparesis improved to grade 4/5 power. On Follow up at 3 months, her bilateral sixth nerve palsy had resolved partially and she was independent for her daily activities. MRI of the brain performed at 6 months did not show any signs of relapse [Figure 4a and b] following which TMP/SMZ was discontinued.

**Discussion**

Nocardiosis is a rare opportunistic infection occurring in the immunocompromised host.[1] The causative organism is *Nocardia* species - an aerobic, catalase positive, g variable to g positive, acid alcohol fast, nonmotile bacilli with branching filaments.[2-4] These soil-borne ubiquitous actinomycetes often cause localized or disseminated infection. Primary infection is acquired following inhalation of airborne bacilli or by direct cutaneous inoculation resulting in pulmonary or cutaneous manifestations respectively.[1-3] Hematogenous spread results in widespread systemic dissemination.[1,6,7] Only 25-40% of patients with systemic nocardiosis develop cerebral infection.[5]

Cerebral Nocardiosis is a rare clinical entity representing only 2% of all cerebral abscesses.[8,9] It usually results in...
Table 1: The cases of Nocardial brain abscesses published in recent literature

| Author            | Age/sex | Immunocompromised | Location                      | Primary Species | Surgery                          | Treatment                                      | Morbidity |
|-------------------|---------|-------------------|-------------------------------|-----------------|----------------------------------|-----------------------------------------------|-----------|
| Present case      | 20/female | Yes               | Left frontal lobe             | No              | N. asteroides                    | Craniotomy and total excision                | No        |
| Patil et al.      | 53/male  | No                | Right occipital lobe          | No              | N. brasiliensis                  | Craniotomy and total excision (twice)        | No        |
| Lin et al.        | 61/female | No                | Right cerebellar              | No              | N. asteroides                    | Craniotomy                                    | Death     |
| Patil et al.      | 52/male  | No                | Left frontal lobe             | No              | N. asteroides                    | Ceftriazone and metronidazole for 2 months followed by ampicillin for 1-month     | No        |
| Lin et al.        | 61/male  | No                | Right occipital lobe          | No              | N. asteroides                    | Craniotomy                                    | No        |
| Dias et al.       | 69/female | No                | Left frontal lobe             | No              | N. asteroides                    | Craniotomy and total excision                | No        |
| Zakaria et al.    | 67/male  | No                | Left frontal lobe             | No              | N. asteroides                    | Craniotomy and total excision                | No        |
| Kennedy et al.    | 79/male  | No                | Right occipital lobe          | No              | N. asteroides                    | Craniotomy                                    | No        |
| Kennedy et al.    | 79/male  | No                | Right cerebellum and left frontal lobe | No | N. asteroides | Craniotomy | Cotrimoxazole and ceftriazone for 45 days followed by cotrimoxazole for 18 months | No |
| Kennedy et al.    | 61/male  | No                | Right parietal lobe           | No              | N. asteroides                    | Craniotomy and excision                      | No        |
| Kennedy et al.    | 51/male  | No                | Left cerebellar               | No              | N. asteroides                    | Craniotomy                                    | No        |
| Braga et al.      | 58/male  | No                | Lung                           | No              | N. asteroides                    | Craniotomy and excision                      | No        |
| Barnaud et al.    | 33/male  | Yes               | Lungs                          | N. cyriacigeorgical | Stereotactic aspiration         | Imipenem and amikacin/ciprofloxacin for 1-month followed by cotrimoxazole for 8 months | No |
| Valarezo et al.   | 65/male  | Yes               | Right cerebellar               | No              | N. asteroides                    | Craniotomy and excision                      | No        |
| Valarezo et al.   | 42/male  | No                | Right occipital lobe           | No              | N. asteroides                    | Craniotomy                                    | No        |
| Valarezo et al.   | 68/male  | Yes               | Right occipital lobe           | No              | N. asteroides                    | Aspiration (twice) followed by craniotomy    | No        |
| Fleetwood et al.  | 58/male  | No                | Right frontal                 | No              | N. asteroides                    | Craniotomy and total excision                | No        |
| Fleetwood et al.  | 65/male  | No                | Right frontal                 | No              | N. asteroides                    | Craniotomy and total excision                | No        |
| Fleetwood et al.  | 29/male  | No                | Left frontal and right parietal | No | N. asteroides | Craniotomy and total excision | Cotrimoxazole for 1 month, imipenem 1-month, amikacin, cefotaxime, sulfadiazine 1-month, cefotaxime 4 months, ceftriazone 5 months | No |

N. asteroides - Nocardia asteroid; N. brasiliensis - Nocardia brasiliensis; N. cyriacigeorgical - Nocardia cyriacigeorgical
The optimal management of cerebral Nocardiosis has not been established. Nocardial brain abscess have been managed either by antimicrobial therapy alone, aspiration or total excision of the abscess wall.[2,15] Synergetic combination of TMP/SMX (cotrimoxazole) is the drug of choice often requiring prolonged therapy from 6 weeks to 1-year.[4,6] Total excision of the abscess followed by administration of appropriate antimicrobial therapy based on sensitivity pattern results in complete resolution of symptoms.[1] Mamelak et al. reviewed 120 reported cases of Nocardial brain abscess and concluded that the overall mortality rate was 33% among patients with single abscess.[8] Mortality rates were much higher following aspiration alone (50%) as compared to craniotomy and excision (24%) and after nonoperative antimicrobial therapy (30%).[10] On the contrary, Lee et al. have demonstrated that single or repeated aspiration alone is a safe and efficacious treatment for the majority of patients with Nocardial brain abscess.[14] Complete surgical excision and prolonged antimicrobial therapy are needed to prevent relapse of infection.[3,5] Table 1 briefly summarizes the cases of Nocardia brain abscesses published in recent literature.

Though cerebral Nocardiosis is often secondary to primary foci in lungs, primary infection could not be detected in our patient. One of the predisposing risk factor could have been her immunocompromised state following pulmonary tuberculosis. The relatively good outcome in our patient was due to her young age, total surgical excision of the abscess and early initiation of antimicrobial therapy with good compliance.

### Conclusion

Unusual central nervous system infections like Nocardiosis should be considered in the differential diagnosis of brain abscess especially in the immunocompromised state. In patients with multiple ring-enhancing lesions, high index of clinical suspicion is needed to establish the diagnosis of this atypical infection. Total surgical excision of the abscess reduces the mass effect, surrounding vasogenic edema and provides bacteriological diagnosis, thereby enabling early initiation of organism-specific antimicrobial therapy. Prolonged antimicrobial therapy and long-term surveillance are needed to prevent relapse, thereby ensuring excellent patient outcome.

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How to cite this article: Baldawa S, Nayak N, Kukreja S, D’souza D, Diyora B, Sharma A. Cerebral nocardiosis. Asian J Neurosurg 2014;9:245.

Source of Support: Nil, Conflict of Interest: None declared.