



Tuberculosis of Frontal Bone—A Rare Entity: Case Report and Review of Literature

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Ind J Med Paediatr Oncol

Abstract

Tuberculosis of the frontal bone is a rare entity, most commonly occurring in childhood. In situations of painless scalp swelling coupled with or without a discharging sinus that has not responded to antibiotics, a strong index of suspicion must be raised. Generally, conventional radiography and computed tomography are used for establishing the diagnosis along with microbiological confirmation. We present an interesting case of a young boy who presented with a 2-month-old bone lesion in the frontal scalp, accompanied by mediastinal lymph node involvement. Histopathology revealed tuberculous origin and successful antitubercular therapy resulted in significant improvement within 3 months. The calvarial bones can rarely be affected by tuberculosis, which can present with swelling, sinus discharge, pain, or in rare cases as blindness. Prompt diagnosis of this disease is needed as it is potentially curable and has an excellent prognosis. The cornerstone of treatment is antitubercular therapy, while surgical intervention may sometimes be required. Our case highlights the importance of keeping tuberculosis as a differential at the back of the mind when dealing with scalp swellings, particularly in children.

Keywords

- ▶ case report
- ▶ tuberculosis
- ▶ calvaria
- ▶ CT
- ▶ PET-CT

Introduction

Infection with mycobacterium tuberculosis (TB) is still widespread in developing nations. With the resurgence of immunocompromised states like human immunodeficiency virus infection, malignancies, etc. there is a rise in the incidence of TB in developed countries. In the last 10 years, there has been a dramatic transformation in the clinical pattern and presentation of TB. Disseminated TB may occur either due to activation of a latent focus of infection or secondary to progressive primary pulmonary TB. The tubercular bacilli are said to erode the epithelial layer of the alveoli, and enter the pulmonary vein. Following this, they enter the left side of the heart, and get disseminated into the systemic circulation

thereby reaching bones, kidneys, brain, and other organs. Only 0.2 to 1.3% of skeletal TB cases involve calvarial bones, which is almost always secondary to disseminated TB. The first case was reported in Germany by Reid.¹ Strauss reported 220 cases in 1933.² Meng and Wu reported 40 cases.³

Case History

An 8-year-old boy, born of a nonconsanguineous marriage and with no family history of cancer, presented with a 2-month-old history of bulge in the frontal region of the scalp. It was initially peanut-sized, and showed subsequent increase in size. The swelling was associated with mild pain. Skin over the swelling, however, was normal. There were no

DOI <https://doi.org/10.1055/s-0043-1772234>.
ISSN 0971-5851.

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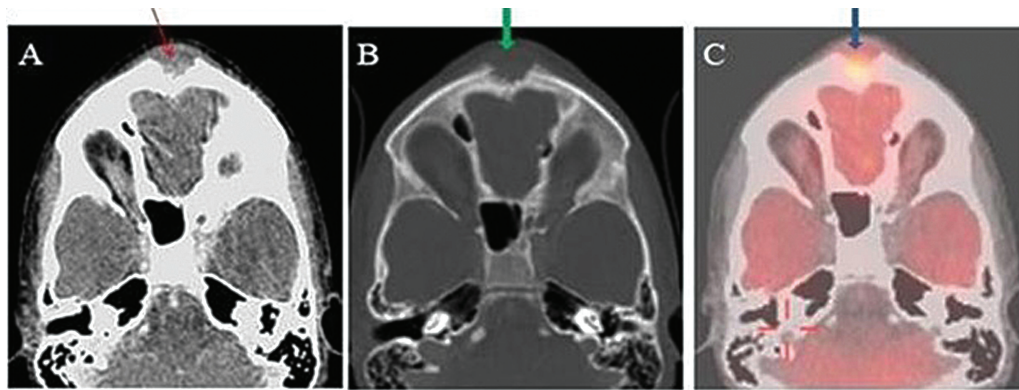


Fig. 1 (A) Axial computed tomography (CT) soft tissue window shows a lytic lesion in the frontal bone with associated soft tissue (red arrow). (B) Axial CT bone window shows soft tissue in the frontal bone associated with erosion of underlying bone. No periosteal reaction was seen. (C) The fused transverse image shows 18F-fluorodeoxyglucose avid lytic frontal bone lesion. Possibilities on imaging were Langhans cell histiocytosis, metastasis. Frontal bone lesion biopsy revealed necrotizing granulomatous inflammation of tuberculous etiology.

constitutional symptoms, no loss of weight, or loss of appetite. On examination, the vitals were stable. There was no sinus at the site of the swelling. No neurological deficits were present.

There was no pallor or organomegaly. Positron emission tomography-computed tomography was done outside our institute that revealed a hypermetabolic osteolytic lesion in the frontal bone in the midline as shown in **Fig. 1** and D10 vertebra body associated with soft tissue component. Pleural and parenchymal nodules with associated mediastinal lymph nodes (left hilar, left prevascular, aorta pulmonary, paratracheal, pretracheal and subcarinal nodes) were also seen as

shown in **Fig. 2**. A frontal bone lesion biopsy was performed under ultrasonography guidance for further assessment, and the sample was tested for acid-fast bacilli staining and polymerase chain reaction (cartridge based nucleic-acid amplification) testing. Results revealed necrotizing granulomatous inflammation of TB etiology. The patient took antitubercular drugs including rifampicin (10 mg/kg), ethambutol (15 mg/kg), isoniazid (5 mg/kg), and pyrazinamide (25 mg/kg) for a period of 3 months followed by rifampicin (10 mg/kg) and isoniazid (5mg/kg) for the next 9 months. The swelling size reduced and complete resolution of the scalp lesion was noted 3 months post initiating the therapy.

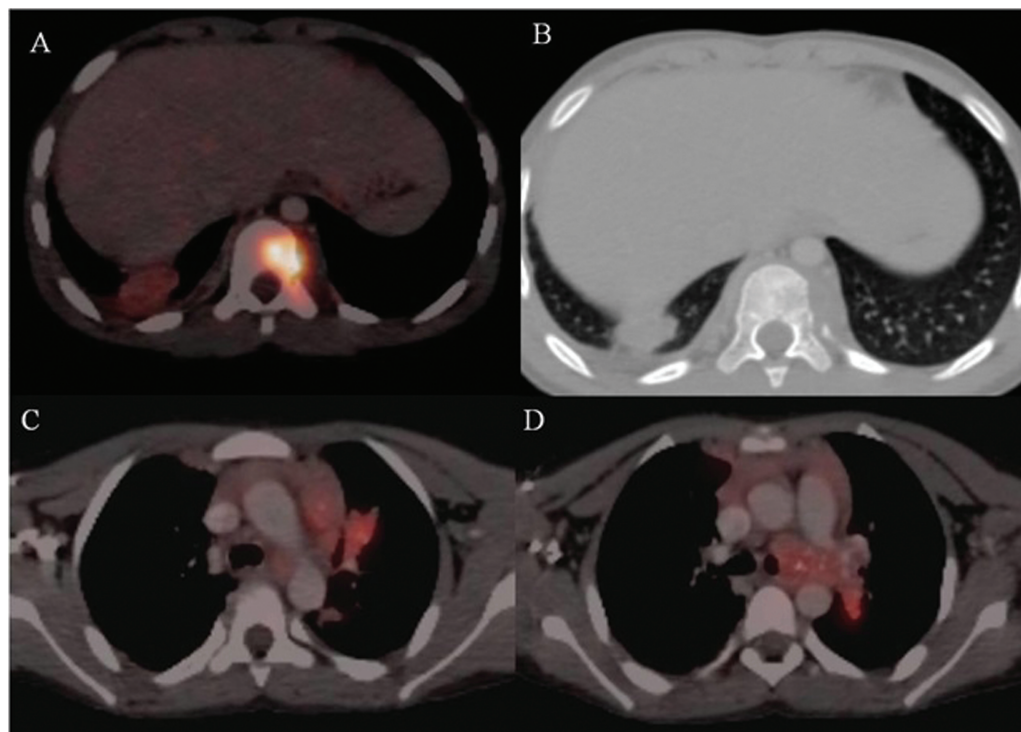


Fig. 2 (A, B) Axial positron emission tomography-computed tomography (PET-CT) shows a lytic lesion with associated soft tissue involving body of D10 vertebrae. (C, D) Axial PET-CT shows pleura and parenchymal nodules with associated mediastinal lymph nodes with foci of calcification within.

Table 1 Differential diagnosis of calvarial lesion

Pathology	Age	Radiograph	CT	MRI	PET	Additional features
Pyogenic osteomyelitis	Any age; however, more common in children and adolescents	ACUTE: Ill-defined lucent area with or without periosteal reaction, usually associated with sinusitis/mastoiditis CHRONIC: Osteolytic lesion with or without sequestrum and/or involucrum	An osteolytic lesion with or without dead necrotic bone / new bone formation	T1: Isointense to hypointense lesion Cortical destruction T2: Hyperintense lesion with surrounding high SI edema T1c + : Enhancement of periphery of associated collections and surrounding soft tissue	Hypermetabolic	Constitutional symptoms Raised WBC count Blood cultures are positive for bacterial/fungal organisms
Langhan's cell histiocytosis	Children	Geographic skull: Single (eosinophilic granuloma) or multiple osteolytic lesions without sclerotic rim Button sequestrum	An osteolytic lesion with a "bevelled edge" appearance (due to asymmetrical involvement of the inner and outer tables) ("hole within a hole" sign) Button sequestrum	T1: Isointense to hypointense T2: Heterogeneously hyperintense T1c + : Marked enhancement	Hypermetabolic	Asymptomatic or constitutional symptoms may be present depending on multisystem involvement. Histopathology shows "Langhans cells" and immunohistochemistry is positive for CD1a and/or CD207
Multiple myeloma	>40 years, With maximum cases between 50 and 70 years	Raindrop skull: Multiple, variable-sized well-defined punched out osteolytic lesions without sclerotic rim	Osteolytic lesions throughout the skull Axial and appendicular skeletal involvement in the form of diffuse osteopenia, osteolytic lesions with endosteal scalloping, and vertebral compression fractures	T1: Hypointense T2: Hyperintense T1C + : Post-contrast enhancement seen	Hypermetabolic	Monoclonal gammopathy (IgA and/or IgG) Reversal of albumin/globulin ratio Bence Jones protein proteinuria Hypercalcemia
Metastasis	Depend on primary cancer, however more common in the older age group. Commonly from neuroblastomas and Ewing's sarcoma in children	Commonly osteolytic, maybe be mixed/sclerotic depending upon the primary malignancy. May show spiculated ("hair-on-end") periosteal reaction	Commonly osteolytic, maybe be mixed/sclerotic.	T1: Isointense to hypointense T2: Hyperintense T1C + : Post-contrast enhancement seen	Hypermetabolic	
Dermoid/epidermoid cyst	Any age, however common in 3rd to 4th decades	Well-defined osteolytic lesion with lobulated margins and sclerotic rim	Well-demarcated osteolytic lesion with smooth/lobulated margins, affecting the inner table more affected than the outer table. Internal fat density and/or calcification	T1: Both cysts appear hyperintense T2: Varying signal depending on content T1C + : Dermoid cysts may show peripheral enhancement Intradiploic epidermoid cysts do not enhance	Variable metabolic activity	Dermoid cysts develop along the midline Painless swellings with or without associated sinus tract Epidermoid cysts show restricted diffusion on DWI

Abbreviations: CT, computed tomography; DWI, diffusion-weighted imaging; IgA, immunoglobulin A; MRI, magnetic resonance imaging; PET, positron emission tomography; WBC, white blood cell.

Discussion

TB of the calvarium is almost always secondary to primary tubercular infection at various other sites such as lungs, lymph nodes, bones, gastrointestinal tract, and central nervous system. It is highly unusual to have isolated calvarial TB. There is a male preponderance, with the male to female ratio being 2:1. The maximum numbers of cases (75–80%) occur in less than 20 years of age.³ It is extremely rare in infants, likely due to a paucity of cancellous bone. Due to the relatively higher proportion of cancellous bone, the frontal and parietal bones are most frequently damaged, followed by the occipital and sphenoid bones. Sphenoid bone involvement has been reported in a few cases.⁴ The tubercular bacilli from the primary focus hematogenously seed the diploe to begin the pathogenesis of calvarial TB. The virulence depends on the patient's state of immunity. In immunodeficiency states, there is increased pathogenicity of the organism that causes the infection to spread to both the outer and inner table, thus destroying them, causing obliteration of capillaries, and replacement of bony trabeculae by granulation tissue. Microscopic examination shows necrotic areas, caseation, granulation tissue, epithelial cells, and Langhans giant cells. Involvement of the outer table leads to the formation of soft tissue swelling in the subgaleal region also referred to as Pott's puffy tumor and discharging sinuses. On the other hand, extradural granulation tissue forms when the inner table is involved and its extension continues. Since cranial sutures do not prevent granulation tissue from spreading, substantial bone destruction may take place before a sinus or swelling manifests. The majority of patients with frontal bone TB have painless scalp edema and discharge-producing sinuses. Initial presentation with seizures, focal neurological deficits, and meningeal signs is uncommon. Despite the dura being an effective barrier, subdural empyemas, meningitis, and intraparenchymal extension can also occur. Usually, skeletal TB occurs along with pulmonary involvement.³ Sometimes, it may have local spread from adjacent mastoiditis. Trauma is also considered as one of the contributing factors due to increased vascularity and decreased resistance.⁵ Radiological features are not always classical and diagnostic. The earliest finding is an area of rarefaction. This is followed by a punched-out lytic lesion with a central sequestrum. Sometimes, the defect may be surrounded by a sclerotic rim. There can be single or multiple lesions. Diffuse involvement or base of the skull is less commonly involved.⁵ Since sometimes there may be multiple lytic lesions, multiple myeloma and metastases are considered differentials. Conventional radiography, computed tomography (CT), and magnetic resonance imaging (MRI) can be used for the diagnosis of frontal bone TB. The disintegration of one or both of the tables is visible on CT with associated soft tissue swelling.⁶ On conventional radiography, three types of tubercular osteitis lesions are identified based on the type of calvarial destruction. The most common subtype are small, punched-out lesions with granulation tissue covering the inner and outer tables of the calvaria also

known as "circumscribed lytic lesions." This term was used by Volkmann.⁷ It is not associated with periosteal reaction.

Our case likely belonged to this subtype of tubercular osteitis, based on its radiological features as seen in ►Fig. 1.

"Spreading-type" is the term coined for lesions causing extensive extradural granulation tissue in addition to substantial damage of the inner skull table. The least common is the "circumscribed sclerotic variant."⁸ Periosteal reaction is very rare but has been reported. CT is used to assess the degree of bone involvement, demonstration of the soft tissue component, sequestrum, and spread of disease to the extradural space meninges and brain parenchyma. A hypoattenuating lentiform or crescent-shaped collection describes the characteristic appearance of epidural granulation tissue on CT. The surrounding meninges show intense post-contrast enhancement on contrast-enhanced CT scan. MRI reveals a high signal intensity soft tissue mass within the bone defect projecting into the epidural space, with intense capsular enhancement on post-contrast sequences. MRI has higher sensitivity for the detection of meningeal enhancement, ventriculitis, and intraparenchymal granulomas or tuberculomas (rarely present along with calvarial TB). MRI was not required in our case as neurological symptoms suggestive of complications were absent.

Microbiological and histological confirmation is essential before starting Ziehl Neelsen staining. Biopsy in our case proved fruitful in clinching the diagnosis of TB as necrotizing granulomatous inflammation was detected on histopathological examination, which was further confirmed to be of tubercular etiology by Ziehl Neelsen staining and polymerase chain reaction testing. Other causes of necrotizing granulomas, which could be considered in cases of noninfective cases, include vasculitis such as Wegener's granulomatosis, rheumatoid nodules, and sarcoidosis.

Frontal bone or calvarial TB was primarily treated through surgery, before the advent of modern antitubercular therapy. Surgery is now reserved for patients with large epidural collections, extensive bone destruction, and sequestrum formation or in cases of mass effect on the brain or focal neurological deficits. The sinus tract is removed along with the complete excision of the damaged bone and debridement of the granulation tissue. However, the treatment of tubercular osteomyelitis of the skull is antitubercular treatment with appropriate antibiotic cover for secondary infections. Current guidelines advocate five drugs for at least 24 months. The important differentials to be considered in cases of TB of the frontal bone are pyogenic osteomyelitis; Langhans cell histiocytosis, multiple myeloma, epidermoid/dermoid cyst, and metastases. ►Table 1 shows differential diagnosis of osteolytic calvarial lesions that can mimic calvarial TB.

Conclusion

The frontal and parietal bones are rarely affected by calvarial TB, which is typically a complication of pulmonary TB. With this case report, we aim to highlight the fact that in a patient presenting with scalp swelling and a punched out lytic lesion, a suspicion of calvarial TB can be made over a

metastatic lesion. Conclusive diagnosis requires imaging findings to be combined with histological and microbiological evidence of the disease. In countries endemic to TB, the differential of tubercular etiology should always be kept in the mind of the clinicians and radiologists to administer prompt and effective care to the patient. This case report hopes to raise awareness that despite isolated calvarial TB being rare, the inconvenience of going through a biopsy for the patient may be avoided if there is a high level of suspicion and clinic-radiological correlation is made. Calvarial TB must be treated surgically and pharmaceutically, and continuous follow-up is essential.

Declaration of Patient Consent

The authors certify that they have obtained all appropriate patient consent forms. In the form, the patient has their consent for their images and other clinical information to be reported in the journal. The patient understands that their names and initials will not be published and due efforts will be made to conceal their identity, but anonymity cannot be guaranteed.

Financial Support and Sponsorship

Nil.

Source of Funding

Nil.

Conflict of Interest

None.

Acknowledgment

We acknowledge our institute Tata Memorial Hospital for providing us with the means and resources to be able to write this article and access the images.

References

- 1 Ram H, Kumar S, Atam V, Kumar M. Primary tuberculosis of zygomatic bone: a rare case report. *Journal of Infection and Public Health*. 2020 May 1;13(05):815–817
- 2 Strauss DC. Tuberculosis of the flat bones of the vault of the skull. *Surg Gynecol Obstet* 1933;57:384–398
- 3 Meng CM, Wu YK. Tuberculosis of the flat bones of the vault of the skull: a study of forty cases. *J Bone Jt Surg* 1942;24(02):341–353
- 4 Thomas ML, Reid BR. Cranial tuberculosis presenting with proptosis. *Radiology* 1971;100(01):91–92
- 5 LeRoux PD, Griffin GE, Marsh HT, Winn HR. Tuberculosis of the skull—a rare condition: case report and review of the literature. *Neurosurgery* 1990;26(05):851–855, discussion 855–856
- 6 Raut AA, Nagar AM, Muzumdar D, et al. Imaging features of calvarial tuberculosis: a study of 42 cases. *AJNR Am J Neuroradiol* 2004;25(03):409–414
- 7 Volkmann R. Die perforierende Tuberkulose der Knochen des Schädeldaches. . [Perforating tuberculosis of the skull bones] *Zentralbl Chir* 1880;7:3–7
- 8 Mohanty S, Rao CJ, Mukherjee KC. Tuberculosis of the skull. *Int Surg* 1981;66(01):81–83