Pituitary abscess: report of two cases and review of the literature

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Abstract: Pituitary abscess is a rare but critical disorder caused by an infectious process where purulent material accumulates inside the sella turcica. Since symptoms, signs and radiographic characteristics of pituitary abscess are similar to several other pituitary lesions, correct diagnosis before surgery is challenging. In this article, two cases of pituitary abscess treated in our department are reported, followed by a literature review. In these two cases, both patients presented with intermittent fever. Magnetic resonance imaging revealed a suprasellar lesion with rim enhancement after contrast injection. After transsphenoidal surgery, the diagnosis of pituitary abscess was confirmed. The patients were then given antibiotic treatment and recovered fully in less than 2 months. Findings of this article support timely diagnosis and proper treatment including transsphenoidal surgery and antibiotic therapy for pituitary abscess, leading to lower mortality rates and higher probability of pituitary hormone function recovery.

Keywords: pituitary abscess, transsphenoidal resection, antibiotic therapy, diagnosis

Introduction

Pituitary abscess (PA) is a rare intracranial mass with high mortality,¹ which explains less than 1% of all pituitary disease.² In most cases, pathogens are not clear. Its treatment is mainly surgical, combined with the use of antibiotics.³ It may occur either as a primary disease or as a result of infection such as sphenoiditis and meningitis.⁴ The clinical manifestations of PA are similar to other pituitary lesions. Because of the rarity of PA and the non-specific symptoms, it is difficult to make a diagnosis before surgery.⁵ Here, we report two cases of PA. Written informed consent has been provided by both patients to have their case details and any accompanying images published.

Case series

Case 1

A 37-year-old man presented with intermittent fever for 2 months. Upon admission, the patient was given antibiotic over a 5 day period. Each time after antibiotic treatment his body temperature returned to a normal level. Nevertheless, upon stopping the antibiotic therapy he became febrile again. He had no medical history and was not taking any regular medication, other than antibiotics for his fever.

On clinical examination, his temperature was 39°C. Neurological examination revealed bitemporal hemianopsia without meningeal signs. Breathing sounds were normal and the abdomen was soft with no mass.

Examination of routine blood work showed mildly high white cell count ($12.13 \times 10^9/L$ with 56.8% neutrophils). Laboratory investigation of inflammatory

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markers, showed elevated erythrocyte sedimentation rate (ESR) of 31 mm/h (normal: 0–15 mm/h), and elevated hypersensitive c-reactive protein (HCRP) levels of 5 mg/L (normal: 0–3 mg/L). Examination of the cerebrospinal fluid (CSF) showed increased protein levels of 787 mg/L (normal: 150–450 mg/L), low glucose levels of 1.93 mmol/L (normal: 2.5–4.5 mmol/L) and high white cell count ($170 \times 10^6/L$ with 80% neutrophils). No pathogen was isolated from blood or CSF cultures. A routine urine test indicated a urine-specific gravity of 1.010 and no signs of infection. Urinary excretion was 3 to 3.5 L/24h. Hormonal evaluation revealed elevated serum prolactin (PRL) levels of 34.9 ng/mL (normal 4.13–18.40), decreased serum thyroid-stimulating hormone

(TSH) of 0.115 $\mu IU/mL$ (normal: 0.35–5.5) with normal free thyroxine (fT4) of 0.82 ng/dL (normal: 0.8–1.76 ng/dL), and a low morning serum cortisol of 0.819 $\mu g/dL$ (normal: 6.2–19.4 $\mu g/dL$) with normal adrenocorticotrophic hormone (ACTH) of 19 (normal: 5–50 pg/mL).

Magnetic resonance imaging (MRI) scan showed a more marked thickening of the stalk and a sellar mass measuring 1.6×1.4×1.3 cm with suprasellar extension. After administration of gadolinium, there was a partial rim enhancement appearance of the lesion on T1-weighted images. The pituitary mass had smooth borders and it compressed the optic chiasm. The patient was diagnosed with possible pituitary adenoma (Figure 1A and B).

Figure 1 Preoperative T1-weighted coronal (A) and sagittal (B) images after intravenous gadolinium contrast administration demonstrated a suprasellar partial rim enhancing the mass. Six months after surgery T1-weighted coronal (C) and sagittal (D) images after intravenous gadolinium contrast administration revealed resolution of the pituitary abscess after surgery.

Empirical antibiotic therapy with intravenous ceftriaxone and hydrocortisone replacement therapy were given to the patient. Thirty-five days following intravenous antibiotics, the patient's temperature, examination of routine blood work, and CSF were all at a normal level. Transsphenoidal resection was performed subsequently. Intraoperatively, a significant amount of yellowish pus was encountered upon surgical incision of the dura mater that forms the capsule of the pituitary gland. The pus was drained and samples were sent for histological analysis and culture. The pituitary gland was found to be intact after the removal of the lesion. No pathogen was isolated and the pathology showed benign anterior pituitary tissue with marked lymphoplasmacytic inflammation.

After the diagnosis of PA, antibiotic treatment continued. The patient was treated with ceftriaxone (Shanghai Roche Pharmaceutical Co., Ltd, Shanghai, People's Republic of China) and teicoplanin (Sanofi-aventis S.p.A, Milano, Italy) intravenously for 2 weeks and oral antibiotic therapy for 4 weeks. Postoperatively, the patient was afebrile, but he developed polyuria (>6 L per day) and was treated with desmopressin (0.3 mg orally thrice daily). Postoperative MRI demonstrated a complete removal of the intrasellar mass (Figure 1C and D).

Two months after surgery, the endocrine laboratory test results were all normal. Visual field examination was normal and there was no bitemporal hemianopsia. Hydrocortisone was discontinued and he was prescribed desmopressin (Ferring Pharmaceuticals Inc., Parsippany, NJ, USA) once daily.

Case 2

A 36-year-old man with no significant past medical history was investigated for headache and intermittent fever lasting for 1 month. He did not receive antibiotic treatment for his fever prior to admission.

At the time of admission he was afebrile. There were no meningeal signs and no neurological or cranial nerve deficits. Visual evaluation was normal.

Routine blood test, ESR, HCRP, PRL, ACTH and routine urine tests were all within the normal range, apart from a decrease in the level of TSH (0.271 μ IU/mL, normal: 0.35–5.5 μ IU/mL), fT4 (0.75 ng/dL, normal: 0.8–1.76 ng/dL), and morning serum cortisol (1.022 μ g/dL, normal: 6.2–19.4 μ g/dL).

A brain MRI of hypothalamo-pituitary area revealed 1.2 $\times$ 1.2 $\times$ 1.3 cm mass which had isointense signal in T1-weighted images and hyperintense signals in T2-weighted images with peripheral rim enhancement after administration

of gadolinium. An initial diagnosis of PA was considered (Figure 2A and B).

The patient underwent endoscopic transsphenoidal surgery. Upon dura incision, a yellowish purulent material was found (Figure 3). After evacuating the abscess, the pituitary was found preserved intact. A culture of the pus was positive for *Staphylococcus aureus*. Histological examination revealed fibrocollagenous tissue with infiltration by lymphoplasmacytoid cells.

After surgery the patient was given intravenous ceftriaxone and teicoplanin for 2 weeks. Afterwards, oral antibiotic therapy was continued for another 4 weeks. There were no postoperative complications. Postoperative MRI showed no irregular changes post-surgery and complete evacuation of the abscess (Figure 2C and D).

Six weeks after surgery, the laboratory investigation results were all within the normal range. The patient recovered fully. Hormone replacement treatment was discontinued.

Discussion

PAs are uncommon, with only about 270 cases globally reported to date.⁶ PA has an incidence of 1% among pituitary lesions.^{2,6–9}

PA may occur de novo, such as in our patients, in a previously healthy normal pituitary gland in 70% of cases.^{2,10} It also can occur secondarily to a pre-existing lesion in the pituitary region, such as pituitary adenoma, craniopharyngioma, or Rathke's cleft cyst.^{11–13} Moreover, immunosuppression, previous irradiation or surgical procedures to the pituitary gland are also known risk factors for PA.^{7,9} In most cases of PA, no apparent causes are generally found.⁷

Our patients presented with fever, headache, visual disorder and symptoms of pituitary hormone deficiency. Anterior pituitary hypofunction is the most common symptom of PA, such as anorexia, weakness, amenorrhea, vomiting, and hypogonadism. In one series of 33 patients, 28 reported anterior pituitary hypofunction symptom.⁷ Headache is the second-most common symptom. In another series of 66 patients, 46 reported a headache.⁶ Visual disorders, such as hemianopsia, are also commonly observed. In one series¹⁴ 57% of patients with PA presented with visual disorders. Central diabetes insipidus has an incidence of 41.4%–69.7% among PA patients.^{4,7} However, it is uncommon for pituitary adenomas to present with diabetes insipidus.¹⁵ Heary reported that diabetes insipidus was only present in 10% of pituitary adenoma cases.¹⁶ Therefore, diabetes insipidus in the diagnosis of PA is of great significance. Fever with signs of meningeal irritation was inspected in only 25% of patients.⁹

Figure 2 Preoperative T1-weighted coronal (A) and sagittal (B) images after intravenous gadolinium contrast administration revealed a suprasellar rim enhancing mass. Five days after surgery T1-weighted coronal (C) and sagittal (D) images after intravenous gadolinium contrast administration revealed resolution of the pituitary abscess after surgery.

Microorganisms such as *Staphylococcus* and *Streptococcus* are the most common isolated pathogens from pituitary abscess.⁹ Additionally, plenty of different microorganisms have been seen in surgical cultures from PA such as *Neisseria* spp., *Micrococcus*, *Citrobacter* spp., *Escherichia coli*, *Brucella*, *Salmonella*, *Corynebacterium*, and *Mycobacterium*.⁹ However, *Aspergillus fumigatus* was most frequently noted in cases of secondary PA.⁶ In our first case, no infectious agent was isolated, which might be due to previous antibiotic treatment.

MRI is one of the most effective imaging approaches for pituitary lesions. Typical features of PA include a single cystic or partially cystic mass that appears hypointense in T1-weighted and hyperintense in T2-weighted imaging,

and shows rim enhancement after gadolinium injection.^{7,17} However, in one series only 66.7% of the patients exhibited such typical features on MRI.⁶ Our patients presented with a rim enhancing suprasellar mass, which was not specific for PA. On the basis of these observations, MRI has limitations in recognizing PA from other pathology of the suprasellar region.

Transsphenoidal excision (TSS) and antibiotic therapy are the main treatment for PA.^{4,6,7} Transsphenoidal resection is preferred over craniotomy, as TSS has the advantages of efficacy, safety, and minimal invasiveness and craniotomy can cause infectious substances to spill into cerebrospinal fluid.^{4,12} Antibiotic therapy should be instituted for approximately 4–6 weeks. It is indicated that empirical treatment should be

Figure 3 Intraoperative picture of transsphenoidal excision showing a significant amount of yellowish pus (white arrow) that was encountered upon surgical incision of the dura mater that forms the capsule of the pituitary gland.

given to the patients who are waiting for microbiology and histological confirmation.^{9,12} Hormone replacement therapy is needed based on pituitary hormone deficiency.^{12,18} Our patients received antibiotic and hormone replacement therapy after transsphenoidal excision.

Timely diagnosis and appropriate treatment of PA typically result in low mortality, and patients may eventually have a good quality of life. In one series of 66 patients, 81.8% of patients had fully recovered, 12.1% of patients had undergone at least one operation for recurrence, and only one patient had died, 4 years after the operation.⁶ Treatment is effective for typical symptoms such as fever, headache and visual changes. In one series of 24 patients, headaches and visual problems were alleviated for over 90% and 75% of patients, respectively.⁹ Two patients had completely recovered from fever, headache and bitemporal hemianopsia. As seen in our patients, proper postoperative hormone replacement is needed considering the high rate of hypopituitarism (62.5%) even after surgery.⁹

In conclusion, these case reports emphasize the importance of timely diagnosis and treatment of PA. Because of similarities in clinical symptoms, signs, imaging and laboratory findings with other sellar lesions, PA is highly difficult to diagnose before surgery. Therefore, the diagnosis of PA should be considered when patients have fever, headache and signs of pituitary dysfunction, or for patients with sellar lesion presenting with signs of meningeal inflammation. Appropriate surgery and medical treatment leads to a lower

mortality rate and higher probability of pituitary hormone function recovery.

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Disclosure

The authors report no conflicts of interest in this work.

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