

88137\_Auto\_Edited.docx

**Multiple inflammatory pseudotumor formation after craniopharyngioma resection via an extended nasal endoscopic approach: A case report**

Multiple inflammatory pseudotumors.

**Abstract****BACKGROUND**

This report describes a case of an intracranial multiple inflammatory pseudotumor (MIP) after endoscopic resection of a craniopharyngioma, which is relatively rarely reported in the literature, and neurosurgeons should be aware of its existence.

**CASE SUMMARY**

Herein, we report the case of a 56-year-old man who developed decreased visual acuity and blurred vision without obvious cause or inducement on Monday, April 27, 2020. To seek further treatment, he went to the Department of Neurosurgery, Clinical Medical College, Yangzhou University. After falling ill, there was no nausea, vomiting, limb convulsions, obvious disturbance of consciousness, speech disorders, cough or persistent fever. The neurological examination findings were normal, and pituitary magnetic resonance imaging (MRI) revealed multiple nodules with abnormal signals in the sellar region. The diagnosis was craniopharyngioma. We performed total resection of the tumor via transnasal endoscopy, and the postoperative pathology suggested that the type of tumor was craniopharyngioma. Six months after the operation, the patient experienced sudden hearing loss in the right ear, tinnitus in both ears, and numbness on the right side of the face and head. Meanwhile, cranial MRI showed multiple inflammatory pseudotumors. After steroid hormone and anti-inflammatory

therapy, the above symptoms did not significantly improve. Finally, the patient's symptoms were well improved by surgery, and the postoperative pathological diagnosis was multiple inflammatory pseudotumors.

## CONCLUSION

Intracranial inflammatory pseudotumor is a benign disease with slow progression, but the clinical symptoms and imaging findings are not typical, no pathological findings, and the diagnosis is relatively difficult. Most of them are treated by surgical resection, and the prognosis is good after surgery.

**Key Words:** MIP; Craniopharyngioma; Treatment; Brain, Oncology; Case report

Wu H, Ding Y, Yan Z, Wei M, Wang X, Zhang H. Multiple inflammatory pseudotumor formation after craniopharyngioma resection *via* an extended nasal endoscopic approach: A case report and literature review. *World J Clin Cases* 2023; In press

**Core Tip:** The diagnosis of MIP was established using MRI, pathology and clinical features. The purpose of this report is to describe the patient's experience and facilitate the observation of postoperative MIP in the future.

## INTRODUCTION

Inflammatory pseudotumors (IP) are localized, isolated, nonneoplastic, occupying lesions characterized histopathologically by a mixture of collagenous fibers and inflammatory cells, such as lymphocytes and plasma cells<sup>[1]</sup>. These tumors have been reported to occur most commonly in the lungs, mesentery, greater omentum, retroperitoneum, genitourinary tract, or upper respiratory tract. However, inflammatory pseudotumors of the central nervous system are rare<sup>[2]</sup>. Most intracranial inflammatory pseudotumors originate from the dura mater and meningeal structures<sup>[3]</sup>.

There have been few reports of multiple inflammatory pseudotumors in the intracranial space after surgery for craniopharyngioma resection.

## **CASE PRESENTATION**

### ***Chief complaints***

The patient, XXX, was a 56-year-old male who developed decreased visual acuity and blurred vision without obvious cause or inducement that lasted for 15 days.

### ***History of present illness***

A 56-year-old male presented to the Department of Neurosurgery, Clinical Medical College, Yangzhou University, with decreased visual acuity and blurred vision that lasted for 15 days. The neurological examination findings were normal, and he had previously been in good health without any infectious diseases or immune disorders. Enhanced pituitary MRI revealed multiple nodules in the suprasellar region with slightly shorter T1 signals, compression of the optic chiasma and compression of the superior pituitary margin in a region of approximately 19 mm×12 mm×21 mm. The preoperative diagnosis was craniopharyngioma (Figure 1A and B). The patient's routine laboratory results were normal. The tumor was removed by an endoscopic transnasal transsellar tubercle sphenoid platform approach. Intraoperatively, the tumor was located in the suprasellar suboptic space with cystic changes and was connected to the posterior part of the pituitary gland. Postoperative MRI showed a patchy mixed signal in the sellar region (Figure 1C and D). Postoperative histopathology revealed hyperplasia, showing nests and a papillary arrangement of epithelial nests with a fenestrated arrangement of marginal cells in some areas; additionally, ghost cells were seen with calcification (Figure 1E and F). The final diagnosis was craniopharyngioma of the sellar region (enamel-forming cell type). Postoperative review MRI showed a patchy mixed signal in the sellar region, and the sellar base was incomplete. A nodal shadow was visible in the sellar region, with significant enhancement; the shadow was approximately 10×9 mm in extent and poorly demarcated from the visual cross. The

patient recovered well after surgery without special discomfort. Six months after the operation, the patient suffered from sudden hearing loss in the right ear, tinnitus in both ears, and numbness of the right side of the face and head, without any signs of infection. Physical examination revealed hearing loss in the right ear. Enhanced MRI of the inner ear showed multiple patchy and nodular, distinctly enhancing abnormal shadows around the right pontocerebellar horn, right cerebellum, and tetrasthanic pool; the largest lesion was in the right pontocerebellar horn, approximately 17×14 mm in size, with gross margins and significant thickening of the right rocky meninges. An inflammatory pseudotumor was considered (Figure 2 A-D). The patient was treated with steroid pulse therapy, and the patient received an initial high-dose regimen of 80 mg of methylprednisolone for four days, followed by a one-day drug cessation and then a maintenance dose of 40 mg of methylprednisolone for another four days. Despite this treatment approach, the patient's primary symptoms remained unresolved, and subsequent cranial MRI indicated no significant change in the size of the pseudotumor. Ultimately, we opted for surgical resection to remove the tumor, and during follow-up and review, there was no recurrence of the inflammatory pseudotumor. A lesion in the right cerebellum and pontocerebellar horn was resected under neuronavigation. Postoperative pathological examination showed fibrocollagenous tissue hyperplasia and extensive infiltration of inflammatory cells, including lymphocytes (Figure 2E and F). Immunohistochemistry suggested the following: KP1 (scattered+), CD3 (scattered+), CD20 (scattered+), S100 (neural+), CD1a (-), GFAP (brain+), Ki67 (scattered 20%), SMA (vascular+), CD34 (vascular+), MPO (scattered+); special staining results: PAS (-), antacid (-), silver stain (-). The postoperative diagnosis was an inflammatory pseudotumor. The patient was discharged 7 days after surgery and was reviewed 3 mo after surgery, without recurrence and no significant abnormalities on neurological examination. The patient continues to be under close follow-up (Figure 2 G-J).

1

### *History of past illness*

The patient had no previous history of infectious diseases, traumas, or bad habits (such as smoking or drinking).

### *Personal and family history*

The patient denied any family history of tumors.

### *Physical examination*

On physical examination, the vital signs were as follows: body temperature, 36.4 °C; blood pressure, 110/88 mmHg; heart rate, 88 beats per min; and respiratory rate, 19 breaths per min. Annual physical examinations (including various laboratory tests and imaging examinations) indicated that the patient was healthy.

### *Laboratory examinations*

White blood cell count:  $7.78 \times 10^9/L$ , neutrophil count:  $7.37 \times 10^9/L$ . Endocrine test results were normal.

### *Imaging examinations*

First pre-operation : Pituitary magnetic resonance imaging (MRI) suggested that there were multiple nodules with abnormal signal in the sellar region. The diagnosis was craniopharyngioma.

Second pre-operation : MRI enhancement of the inner ear showed multiple patchy and nodular distinctly enhancing abnormal shadows around the right pontocerebellar horn, right cerebellum, and tetraspanic pool, with the largest lesion in the right pontocerebellar horn, approximately  $17 \times 14$  mm, with gross margins and significant thickening of the right rocky meninges. Inflammatory pseudotumor was considered

1

### **FINAL DIAGNOSIS**

Combined with the patient's medical history, the final diagnosis was MIP.

## **TREATMENT**

Medical and surgical treatment.

1

## **OUTCOME AND FOLLOW-UP**

Symptoms were relieved gradually after treatment, and complete resolution of symptoms was observed after five days. To date, there has been no recurrence, and the patient is doing well.

## **DISCUSSION**

Pathogenesis

The etiology of inflammatory pseudotumors remains uncertain, with three prevailing theories attempting to explain their origin<sup>[4]</sup>. In terms of their composition, inflammatory pseudotumors exhibit internal fibrosis and infiltration of various inflammatory cells<sup>[3]</sup>, including macrophages, plasma cells, and lymphocytes, suggesting an association with inflammation<sup>[5]</sup>. Furthermore, minor traumas, surgical interventions, smoking, vasculitis and malignancies have been proposed as potential triggers<sup>[6]</sup>, indicating an inflammatory response as a causative factor. Infections caused by viruses such as Epstein–Barr virus, cytomegalovirus, and herpes simplex virus have also been linked to the development of inflammatory pseudotumors<sup>[3,7]</sup>. Moreover, inflammatory pseudotumors have been associated with IgG-4-related diseases, specifically characterized by fibrotic tissue masses infiltrated by IgG-4-positive lymphoplasmacytic cells<sup>[8]</sup>. In the present case, an intracranial inflammatory pseudotumor emerged subsequent to the transnasal endoscopic resection of a craniopharyngioma, with the patient experiencing an intracranial infection postsurgery. Consequently, it is believed that the inflammatory response triggered by an infection following the initial surgery may be responsible for the subsequent development of the intracranial inflammatory pseudotumor. Neuroendoscopic transsphenoidal craniopharyngotomy has been suggested to enhance the rate of complete tumor resection; however, it also increases the incidence of cerebrospinal fluid

leakage and intracranial infection and prolongs the operative time and hospital stay. Thus, for patients undergoing endoscopic transsphenoidal craniopharyngotomy, the administration of adequate antibiotics throughout the postoperative period is recommended to prevent intracranial infection. Elderly patients may require an extended duration of antibiotic treatment, and modifications to the surgical approach may be necessary if deemed appropriate. Additionally, postoperative endoscopic evaluation of the saddle base can facilitate the early detection and treatment of occult cerebrospinal fluid leaks.

### Pathology

Due to the limited number of cases, there is currently no standardized pathological classification for intracranial inflammatory pseudotumors, and most reported cases exhibit a mixed predominance of cell types<sup>[9]</sup>. Microscopically, these lesions are characterized by the presence of polyclonal plasma cells intermingled with varying numbers of lymphocytes, neutrophils, eosinophils, histiocytes and myofibroblasts<sup>[10]</sup>. Immunohistochemical analysis has shown positive expression of smooth muscle actin (SMA) in the majority of intracranial inflammatory pseudotumors<sup>[9]</sup>. In the present case of an intracranial inflammatory pseudotumor, microscopic examination revealed the proliferation of fibrillar collagenous tissue with significant infiltration of inflammatory cells, particularly lymphocytes. Immunohistochemistry also confirmed positive SMA expression. These pathological findings were consistent with those typically observed in conventional intracranial inflammatory pseudotumors. Thus, it is essential to consider the possibility of inflammatory pseudotumors in all cases of intracranial tumor-like lesions exhibiting lymphoplasmacytic infiltrates and varying densities of collagenous interstitium<sup>[11]</sup>. Pathological examination remains the gold standard for the diagnosis of intracranial inflammatory pseudotumors.

### Clinical presentation

Intracranial inflammatory pseudotumors are currently categorized into five distinct types<sup>[11]</sup>: intracerebral parenchymal, meningeal, mixed parenchymal-meningeal,

intracerebroventricular, and extracranial communicating types. The clinical manifestations of these tumors differ slightly depending on their locations. Inflammatory pseudotumors located in the skull base typically manifest as pain on the affected side of the head and face, cranial nerve palsy, or cerebrospinal fluid leakage. Inflammatory pseudotumors within the brain ventricles often present with symptoms such as hydrocephalus, headache, nausea, vomiting, and vision loss. Inflammatory pseudotumors in the brain parenchyma commonly result in increased intracranial pressure and varying degrees of neurological dysfunction based on the specific location. Lesions involving the meninges can exhibit a range of neurological manifestations, with mild cases being asymptomatic and severe cases leading to dysfunction due to compression of the brain parenchyma<sup>[12]</sup>.

#### Imaging manifestations

Brain CT scans reveal a well-defined circumferential shadow with isointense or high-density characteristics, with clear margins and uniform enhancement. MRI of the head demonstrates <sup>4</sup> low signal intensity on T1-weighted imaging (T1WI) and T2-weighted imaging (T2WI), with substantial enhancement observed following contrast administration. Delayed enhancement has also been reported in certain cases. It is crucial to note that the diagnosis primarily relies on histopathological examination<sup>[13]</sup>.

#### Differential diagnosis

Intracranial inflammatory pseudotumors require differentiation from several other conditions<sup>[4]</sup>, including lymphoplasmacyte-rich meningiomas, lymphomas, and plasmacytomas. Given that most intracranial inflammatory pseudotumors originate from dural and meningeal structures<sup>[14]</sup>, distinguishing them from lymphoplasmacyte-rich meningiomas can be challenging, often resulting in misinterpretation as meningiomas with lymphoplasmacytic infiltrates. However, careful pathological examination can differentiate between the two, primarily through the detection of the tumorigenic proliferation of meningeal cells and immunohistochemical staining for EMA (+)<sup>[11]</sup>.

Intracranial primary or secondary malignant lymphomas are malignant neoplasms originating from the central or peripheral lymphatic system, and secondary cases are more common, particularly B-cell non-Hodgkin's lymphoma. Immunohistochemical analysis of lymphomas reveals monoclonal expression of B or T lymphocyte antigens along with positive staining for EMA, which significantly distinguishes them from inflammatory pseudotumors<sup>[11]</sup>.

Plasmacytomas can be discerned through immunohistological flow cytometry, allowing the determination of whether the plasma cells present are polyclonal or monoclonal. Polyclonal plasma cells indicate a plasmacytoid granuloma, while monoclonal plasma cells indicate a plasmacytoma<sup>[4]</sup>.

#### Treatment

The optimal treatment approach for intracranial inflammatory pseudotumors remains uncertain, as there is a lack of a general consensus<sup>[4]</sup>. Existing treatment options include surgical resection, corticosteroid therapy, immunosuppressive therapy, and radiotherapy. Studies have indicated that a significant number of patients exhibit favorable outcomes following surgical resection, with low rates of recurrence or malignant progression<sup>[5,15]</sup>. Surgery offers the advantage of alleviating the mass effect of the pseudotumor and potentially reducing the risk of distant recurrence<sup>[14]</sup>.

Many experts consider corticosteroids to be the preferred treatment for inflammatory pseudotumors, particularly in the initial stages<sup>[10]</sup>. Oral corticosteroids have shown positive effects in the majority of patients, effectively improving symptoms and reducing the size of inflammatory pseudotumors. A recommended initial oral steroid dose of 40 mg per day, gradually tapered to 5 to 10 mg per day over a month, has demonstrated efficacy. However, it is important to note that studies have revealed a relatively high recurrence rate of inflammatory pseudotumors after the discontinuation of oral corticosteroids<sup>[10]</sup>.

Furthermore, while hormone therapy can significantly alleviate patient symptoms, there is evidence of a relatively high recurrence rate following the discontinuation of oral hormone treatment.

In this particular case, hormone therapy using methylprednisolone was initiated upon the diagnosis of an inflammatory pseudotumor. The patient received an initial high-dose regimen of 80 mg of methylprednisolone for four days, followed by a one-day drug cessation and then a maintenance dose of 40 mg of methylprednisolone for another four days. Despite this treatment approach, the patient's primary symptoms remained unresolved, and subsequent cranial MRI indicated no significant change in the size of the pseudotumor. Ultimately, we opted for surgical resection to remove the tumor, and during follow-up and review, there was no recurrence of the inflammatory pseudotumor.

The ineffectiveness of hormone therapy in this case may be attributed to an incorrect hormone dosage and an insufficient treatment duration. The absence of a standardized dose and duration for oral hormone therapy further complicates treatment decisions, highlighting the need for comprehensive case reports to guide future practices and recommendations.

Radiation therapy has been proposed as an alternative treatment option for intracranial inflammatory pseudotumors in cases where the tumors are insensitive to hormone therapy, there are contraindications to high-dose hormone therapy, relapse occurs after hormone therapy, or the tumors are unsuitable for surgery due to the location of the lesion or the possibility of incomplete surgical resection. In such scenarios, low-dose high-potential fractionated therapy is typically recommended, with a total radiation dose ranging from 20 to 40 Gy<sup>[16]</sup>.

In summary, the treatment of intracranial inflammatory pseudotumors remains a complex matter without a definitive treatment approach. Surgical resection and corticosteroid therapy, particularly oral glucocorticosteroid therapy, have shown positive results in improving symptoms and reducing tumor size. However, the risk of recurrence remains a concern, necessitating further research and evaluation of treatment strategies.

## **CONCLUSION**

Intracranial inflammatory pseudotumors lack specific imaging manifestations and are easily misdiagnosed as other tumors, and pathology is currently the gold standard for the diagnosis of intracranial inflammatory pseudotumors. Due to the small number of cases and the diverse histopathological classification, the best treatment plan is still inconclusive. In our opinion, surgery is preferred for inflammatory pseudotumors of the central nervous system that can be completely resected without damaging important surrounding structures. If the lesion is not suitable for surgery, steroid hormone therapy or hormone therapy combined with radiotherapy can be administered as appropriate. Patients with intracranial inflammatory pseudotumors should be treated individually according to the location of the lesion and the patient's own characteristics.

ORIGINALITY REPORT

7%

SIMILARITY INDEX

PRIMARY SOURCES

1	<a href="http://www.wjgnet.com">www.wjgnet.com</a> Internet	114 words — 4%
2	Vaishali Suri. "Intracranial inflammatory pseudotumor: Report of a rare case", Neuropathology, 2/16/2008 Crossref	16 words — 1%
3	<a href="http://f6publishing.blob.core.windows.net">f6publishing.blob.core.windows.net</a> Internet	14 words — 1%
4	<a href="http://www.tandfonline.com">www.tandfonline.com</a> Internet	13 words — < 1%
5	<a href="http://ia903208.us.archive.org">ia903208.us.archive.org</a> Internet	12 words — < 1%
6	<a href="http://www.ncbi.nlm.nih.gov">www.ncbi.nlm.nih.gov</a> Internet	12 words — < 1%

EXCLUDE QUOTES ON  
EXCLUDE BIBLIOGRAPHY ON

EXCLUDE SOURCES < 12 WORDS  
EXCLUDE MATCHES < 12 WORDS