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Editorial Board Member of *World Journal of Clinical Cases*, Jjiang-Huei Jeng, DDS, PhD, Professor, School of Dentistry and Department of Dentistry, National Taiwan University Medical College and National Taiwan University Hospital, School of Dentistry, College of Dental Medicine, Kaohsiung Medical University, Taipei 100, Taiwan. [jhjeng@ntu.edu.tw](mailto:jhjeng@ntu.edu.tw)

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## Primary intracranial synovial sarcoma with hemorrhage: A case report

Yang-Yang Wang, Man-Li Li, Zhi-Yong Zhang, Jiang-Wei Ding, Li-Fei Xiao, Wen-Chao Li, Lei Wang, Tao Sun

**ORCID number:** Yang-Yang Wang 0000-0001-5810-8928; Man-Li Li 0000-0002-3019-6142; Zhi-Yong Zhang 0000-0003-1532-4078; Jiang-Wei Ding 0000-0002-3231-9679; Li-Fei Xiao 0000-0003-2094-3615; Wen-Chao Li 0000-0002-9388-9261; Lei Wang 0000-0002-7449-0570; Tao Sun 0000-0001-8062-3432.

**Author contributions:** Wang YY and Zhang ZY were the patient's neurosurgeons, reviewed the literature and contributed to manuscript drafting; Li ML reviewed the literature and contributed to manuscript drafting; Ding JW contributed to manuscript drafting; Xiao LF, Li WC and Wang L analyzed and interpreted the imaging findings; Sun T was responsible for revision of the manuscript for important intellectual content; all authors have read and approved the final manuscript.

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**Yang-Yang Wang, Jiang-Wei Ding, Li-Fei Xiao, Wen-Chao Li, Lei Wang, Tao Sun**, Department of Neurosurgery, General Hospital of Ningxia Medical University, Yinchuan 750000, Ningxia Hui Autonomous Region, China

**Man-Li Li**, Sanquan College, Xinxiang Medical University, Xinxiang 453000, Henan Province, China

**Zhi-Yong Zhang**, Department of Neurosurgery, The First Affiliated Hospital of Xinxiang Medical University, Xinxiang 453000, Henan Province, China

**Corresponding author:** Tao Sun, MD, Chief Doctor, Department of Neurosurgery, General Hospital of Ningxia Medical University, No. 1160 Shengli Street, Xingqing District, Yinchuan 750000, Ningxia Hui Autonomous Region, China. [suntao\\_nxmu@163.com](mailto:suntao_nxmu@163.com)

### Abstract

#### BACKGROUND

Synovial sarcoma (SS) is a highly malignant tumor of unknown histological origin. This tumor can occur in various parts of the body, including those without synovial structures, but mainly in and around the joints, mostly in the lower extremities. Primary intracranial SSs are remarkably rare. This paper aims to report a case of primary intracranial SS with hemorrhage.

#### CASE SUMMARY

A 35-year-old male patient suffered a headache and slurred speech during manual labor and was sent to the emergency department. Through imaging examination, the patient was considered to have high-grade glioma complicated with hemorrhage and was treated with craniotomy. Postoperative pathology revealed SS. positron emission tomography/computed tomography was performed, which ruled out the possibility of metastasis to the intracranial from other parts of the body. Postoperative radiotherapy was given to the patient, during which radiation necrosis occurred. Sixteen months after craniotomy, cranial magnetic resonance imaging revealed recurrence of the tumor.

#### CONCLUSION

Primary intracranial SS is a rare malignant tumor. Primary intracranial SS with hemorrhage and radiation necrosis should be carefully monitored during postoperative radiotherapy. Surgical resection of the tumor combined with postoperative radiotherapy and chemotherapy is currently used, but the prognosis is poor.

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**Core Tip:** This paper presents a rare case of primary intracranial synovial sarcoma (SS) with hemorrhage. Through imaging examination, the patient was considered to have high-grade glioma complicated with hemorrhage and was treated with craniotomy. Postoperative pathology revealed SS. Postoperative radiotherapy was given, during which radiation necrosis occurred. By reviewing the diagnostic and therapeutic history and analyzing the clinical and radiological manifestations, a better understanding of the characteristics of primary intracranial SS can be achieved, helping to improve diagnosis and treatment.

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## INTRODUCTION

Synovial sarcoma (SS) is a rare malignant soft-tissue tumor that occurs at any age and in any part of the body, but mainly in and around the joints, mostly in the lower extremities of middle-aged patients. The tumor constitutes 5%-10% of soft-tissue sarcomas[1].

Intracranial SS, especially primary intracranial SS, is rare. This article reports a case of SS originating from intracranial SS. The authors believe that this is the second case of an SS patient with a tumor complicated with hemorrhage[2]. This article aims to understand the characteristics of primary intracranial SS and help improve its diagnosis and treatment by reviewing the diagnostic and treatment history and analyzing the clinical and imaging findings.

## CASE PRESENTATION

### Chief complaints

A 35-year-old man was sent to the emergency department due to headache and slurred speech.

### History of present illness

This patient was sent to the emergency department (The First Affiliated Hospital of Xixiang Medical University, Xixiang, Henan Province, China) for treatment due to a sudden headache and slurred speech during manual labor 11 h previously. Head computed tomography angiography (CTA) revealed left middle cerebral artery malformation with hemorrhage. The patient was hospitalized.

### History of past illness

The patient was physically healthy and had no abnormal medical history.

### Personal and family history

The patient has smoked approximately 20 cigarettes a day for 10 years and has not quit smoking. He and his younger brother are twins. His younger brother, who has a history of epilepsy since childhood, has been treated with oral carbamazepine.

### Physical examination

The patient was mildly lethargic and slurred. His right upper, right lower, and left limb muscle strength was Grade 3, 4 and 5, respectively. Physiological reflex was present, the Babinski sign on the right was positive, and other pathological signs were



not elicited.

### Laboratory examinations

White blood cell count was increased ( $12.2 \times 10^9/L$ ), platelet count was decreased ( $109 \times 10^9/L$ ), and phosphorus was increased in the electrolyte test ( $1.39 \text{ mmol/L}$ ). The results of other routine laboratory biochemical tests showed no abnormalities.

### Imaging examinations

CT and CTA indicated left intracerebral lesions with hemorrhage and left middle cerebral artery arteriovenous malformation, respectively (Figure 1). Preoperative magnetic resonance imaging (MRI) revealed a mass of mixed-signal shadows in the left frontotemporal parietal lobe, approximately  $51.3 \text{ mm} \times 54.2 \text{ mm} \times 60.0 \text{ mm}$ , and the wall and substantial part of the lesion were enhanced (Figure 2). Radiological diagnosis of the left frontotemporal parietal lobe was solid cystic mass, which was possibly high-grade glioma with hemorrhage.

## FINAL DIAGNOSIS

The patient was diagnosed with primary intracranial SS.

## TREATMENT

Whole cerebral arteriography was performed because of arteriovenous malformation of the left middle cerebral artery, as suggested by CTA, and no vascular malformation was found. Preoperative examinations were conducted to exclude surgical contraindications. Craniotomy was performed. The dura mater had high tension during the operation, and the brain tissue was expanded. The tumor was gray-red, soft in texture, with a rich blood supply, accompanied by hemorrhage. The M2 segment of the middle cerebral artery and its branches were located inside the tumor. The tumor was resected in pieces, and all the resected tumors were finally removed. Antibiotics, dehydration to lower intracranial pressure, and other related treatments were given after the operation.

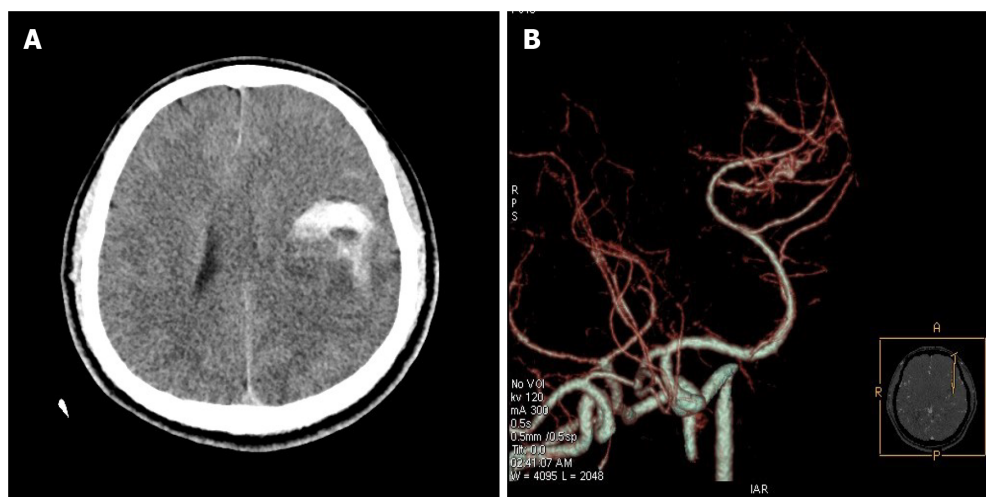
## OUTCOME AND FOLLOW-UP

Pathological examination of the tumor tissue and brain tissue in the surrounding edematous area was performed. Dark red-gray-white tissue was observed, approximately  $9.5 \text{ cm} \times 9 \text{ cm} \times 2 \text{ cm}$  in size. The section was purplish brown and partly gray-white. Microscopically, the tumor cells showed diffused growth with cellular atypia, and mitotic figures (Figure 3). The immunohistochemical results showed the following: GFAP(-), IDH1(R132H)(-), oligo-2(-), S-100(-), TARX(+), CK(-), EMA(-), Ki67(+70%), vimentin(+), CD56(+), NAPSIN-A(-), TTF-1(-), INI1(+), CD34(-), CD99(-), FLI-1(+), NSE(-), PLAP(-), PR(+/-), Sall-4(-), SATB-2(+), SMA(-), Syn(-), Bcl-2(+), and TLE1(+). Further genetic testing of SYT (SS18) was negative for gene breaks, suggesting that SS18 had no fusion with other genes. SS usually occurs in the extremities. Thus, a whole-body positron emission tomography/CT were performed to search for a potential primary tumor. However, no abnormality was found. Postoperative head MRI was also re-examined (Figure 4A).

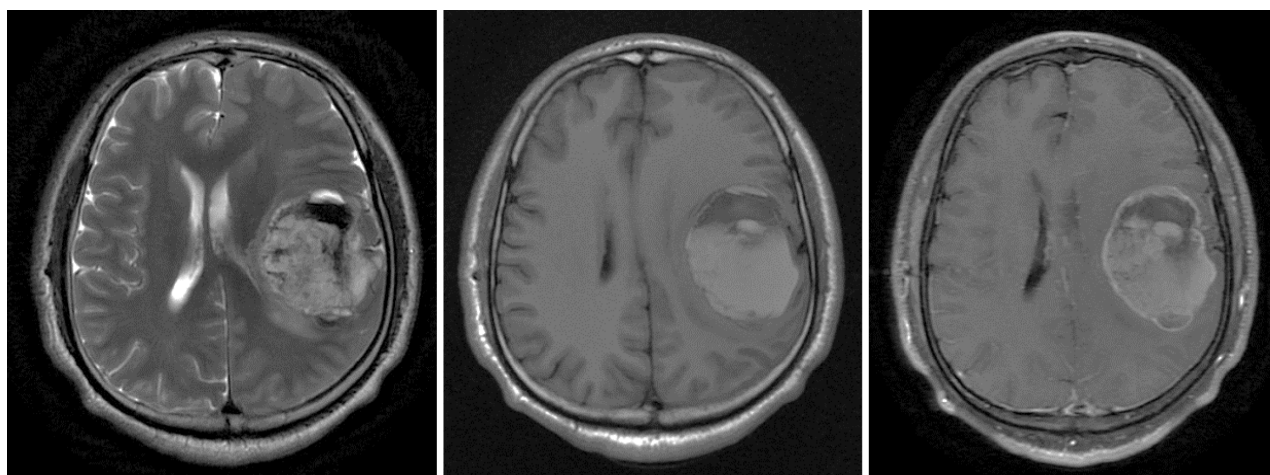
Radiotherapy was administered 2 wk after the operation with 95% PGTV  $2.2 \text{ Gy} \times 30 \text{ F}$ . Six months after craniotomy (3 mo after radiotherapy), perfusion-weighted imaging (PWI) was performed in the outpatient department (Figure 4B-D): The lesion was markedly enlarged, but this enhancement showed hypoperfusion, which was considered to be radiation necrosis, and bevacizumab was given. Sixteen months after craniotomy, the patient developed headaches, and PWI was repeated for detection of tumor recurrence (Figure 5).

## DISCUSSION

SS is a clinically rare malignant tumor. SS is generally believed to be a soft tissue



**Figure 1 Preoperative computed tomography and computed tomography angiography.** A: Computed tomography (CT) indicated a round space on the left with high-density shadows, suggesting tumor with hemorrhage; B: CT angiography indicated the possibility of arteriovenous malformation of the left middle cerebral artery.



**Figure 2 Preoperative magnetic resonance imaging.** Round abnormal signal occupying lesions on T2-weighted imaging (WI) and T1WI were long or short mixed signals, associated with cystic and minimal edema. The wall and substantial part of the lesion were enhanced.

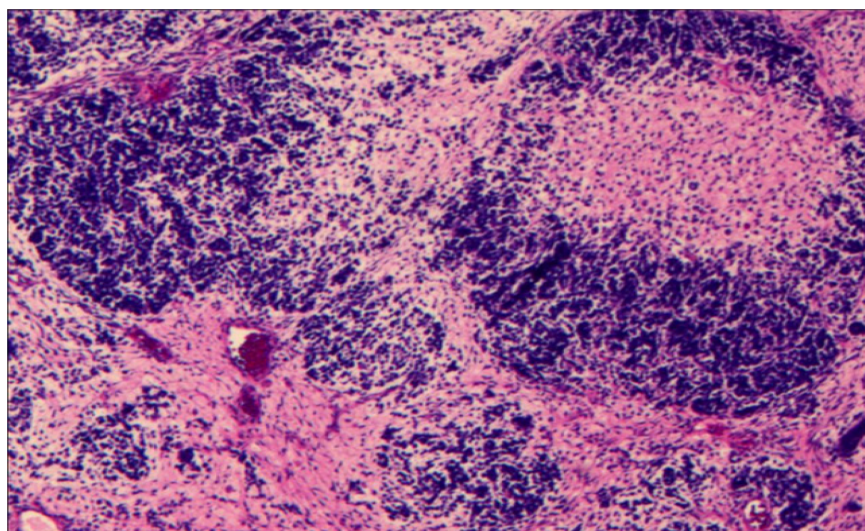
malignant tumor originating from the joints, synovia, and tendon sheath synovia, which usually occurs in the extremities of younger adults. However, SS may occur in any part of the body, such as the heart, kidney, throat and tongue, with an increase in case reports[3-6]. Therefore, SS is considered to be a malignant tumor with an unknown histological origin. The literature review revealed that cases of intracranial primary SS are remarkably rare, and diagnosis by surgery and pathology is uncommon. This article reports a case of SS originating from intracranial SS. The authors believe that this is the second case of SS complicated with hemorrhage. This tumor was initially misdiagnosed as glioma. The final diagnosis was SS after surgical and pathological examination.

The clinical manifestations of primary intracranial SS lack specificity, and diagnosis depends on the location, size and complications of the tumor. The clinical manifestations of primary intracranial SS include headache, nausea, vomiting, limb hemiplegia, and slurred speech. This patient was sent to the emergency department for treatment due to SS with hemorrhage. The clinical manifestations were headache, slurred speech, and hemiplegia. The literature on primary intracranial SSs with hemorrhage is summarized in Table 1. Similarly, imaging examination of intracranial primary SS also lacks specificity[2]. CT mainly manifests as dense or low-density shadows, such as nodular, lobulated masses, and most of the boundaries are clear. CT of this patient showed a round tumor with clear borders, and the part of the tumor with hemorrhage showed high-density shadows. MRI mostly reveals the shadow of soft-tissue masses

**Table 1 Primary intracranial synovial sarcoma with hemorrhage**

Characteristics	Present case	Patel <i>et al</i> [2]
Sex	Male	Male
Age (yr)	35	21
Clinical presentation	Headaches and slurred speech	Headaches, ataxia, left hemianopsia, left arm weakness
Intracranial findings	Solid and cystic mass with hemorrhage in the left frontotemporal parietal lobe	Right parietal mass with hemorrhage
Imaging examinations	CT (left intracerebral lesions with hemorrhage); MRI revealed a mass of mixed-signal shadows in the left frontotemporal parietal lobe, approximately 51.3 mm × 54.2 mm × 60.0 mm, and the wall and substantial part of the lesion were enhanced	CT (right parietal heterogeneous, hyperdense mass with a large medial hematoma)
Treatment	(1) Craniotomy, GTR; and (2) RT	(1) Decompression and clot evacuation; (2) RT; (3) Craniotomy, GTR; and (4) Chemotherapy
Outcome and Follow-up	The tumor recurred 16 mo after the craniotomy	Survived for at least two years

CT: Computed tomography; MRI: Magnetic resonance imaging; GTR: Gross total resection; RT: Radiotherapy.

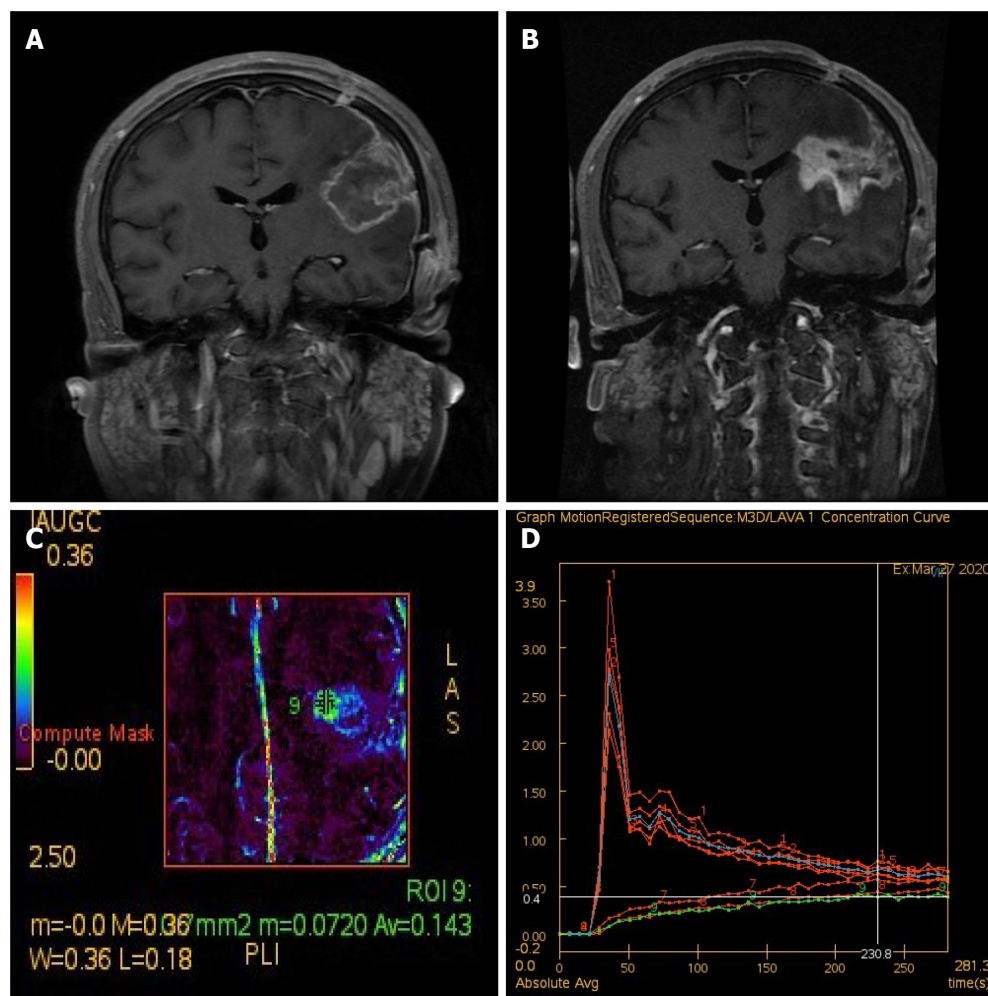


**Figure 3 Pathological examination of the tumor tissue.** Hematoxylin–eosin staining of the tumor cells showed diffuse growth with cellular atypia and mitotic figures. Image at 40 × magnification.

with clear boundaries, with high- or low-signal intensity on T1-weighted imaging (WI) and slightly high signal intensity on T2WI. Mixed signals of low, equal, and high may appear with the occurrence of cystic degeneration, necrosis, and hemorrhage in the mass. MRI of this patient showed long/short T1WI signals, and T2WI demonstrated mixed signals. The wall and substantial part of the lesion were enhanced, which was consistent with the MRI findings of previous cases reported in the literature[7].

Diagnosis of primary intracranial SS requires pathological examination and immunohistochemistry due to the lack of specificity in clinical manifestations and imaging findings. Primary intracranial SS is generally large in volume and usually lobulated and nodular. The section of the tumor is fish-like, gray white, or gray red. Gross observation of the present case showed that the tumor had dark red and gray and white tissues, approximately 9.5 cm × 9 cm × 2 cm in size, purple-brown in section, and partly gray-white, which is consistent with previous reports[7]. SS can be classified microscopically according to the composition of epithelial and/or spindle cells. The classification is as follows: (1) Biphasic type: Contains spindle and epithelioid cells; (2) Monophasic type: Comprises only spindle or epithelioid cells; and (3) Poorly differentiated SS type[7,8]. Vimentin and Bcl-2 are consistently expressed in SS tumor cells, as seen by immunohistochemistry. CD99 and CD56 are frequently expressed but not SMA, S-100, CD34 and GFAP[9,10]. The immunohistochemical





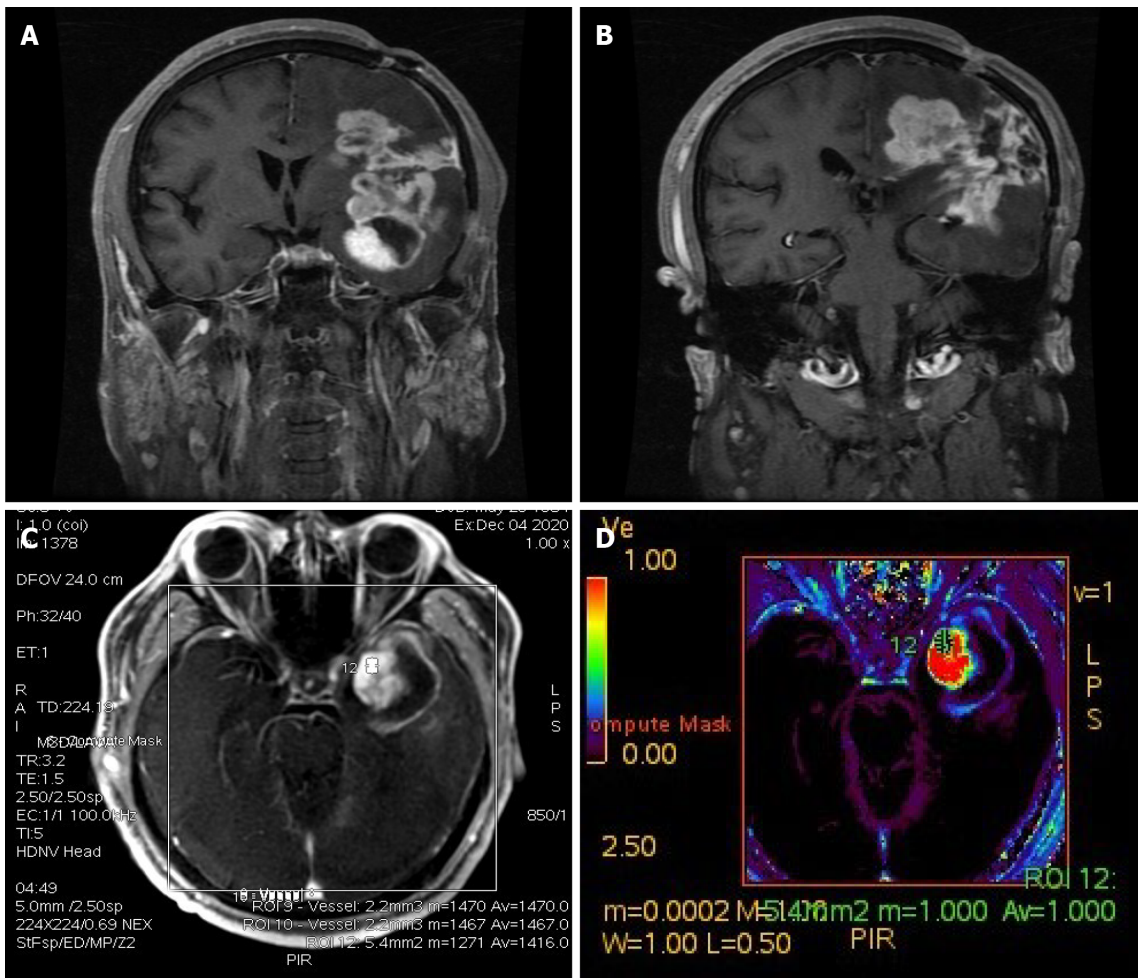
**Figure 4** Imaging changes after treatment of synovial sarcoma. A: Enhanced T1-weighted imaging (T1WI) after craniotomy indicated tumor removal; B: Enhanced T1WI 6 mo after craniotomy (3 mo after radiotherapy). Enhanced T1WI indicated that the lesion was larger than before (B). Analysis of the enlarged part of the lesion (the area marked by 9 in C) suggested that the enlarged lesion is hypoperfusion, and radiation necrosis was considered (C and D).

results of tumor cells in this patient were consistent with those reported in the literature, including vimentin(+), Bcl-2(+), CD56(+), SMA(-), GFAP(-), S-100(-) and CD34(-), as well as CD99(-). In addition, Ki-67 proliferation index of 70% was observed in the most proliferative zone of the tumor.

The cytogenetic marker of SS is currently believed to be the mutual translocation of the SYT gene on chromosome 18 and SSX1 or SSX2 gene on chromosome X and the formation of a new chimeric gene SYT-SSX1 or SYT-SSX2. More than 90% of SSs possess this molecular characteristic[11]. Unfortunately, the genetic test of our patient for SYT (SS18) was negative for gene breaks, suggesting the absence of fusion of other SS18 genes. However, the disease was finally concluded to be a primary intracranial SS after comprehensive consideration of patient history, clinical manifestations, imaging data, and pathological examination, as well as multidisciplinary discussion among neurosurgery, imaging and pathology departments.

The current recommendation for the treatment of primary intracranial SS is to remove the tumor as far as possible while ensuring the patient's safety, followed by postoperative adjuvant radiotherapy and chemotherapy. Reports indicate that the degree of tumor resection is closely related to prognosis[10]. The recommended dose of postoperative adjuvant radiotherapy is 60-70 Gy[12,13]. A 95% PGTV 2.2 Gy × 30 F radiotherapy plan for our patient was performed, but he developed radiation necrosis 3 mo after radiotherapy. In addition, SS is remarkably prone to metastasis, particularly to the lungs, because it is a highly malignant tumor, and related monitoring approaches, such as chest radiography, should be performed during follow-up[10].

Primary intracranial SS is a remarkably rare malignant tumor with atypical clinical and imaging manifestations. Diagnosis requires pathological and cytogenetic examination. The current treatment methods are surgical resection combined with postoperative radiotherapy and chemotherapy. Numerous experiences and lessons



**Figure 5** Imaging changes during patient follow-up. A and B: Enhanced T1-weighted imaging 16 mo after craniotomy indicated enlargement of the lesion; C and D: Perfusion-weighted imaging of the enlarged part of the lesion (area marked by 12 in C) suggested that the enlarged lesion had high perfusion, and tumor recurrence was considered.

can be learned from the treatment of our patient. He and his brother are twins, and his brother has had a history of epilepsy since childhood. The possible relation of this kind of primary intracranial SS to the history of the patient should be further studied. In addition, primary intracranial SS with hemorrhage and radiation necrosis should be carefully monitored during postoperative radiotherapy. Overall, there are few cases of primary intracranial SS, and the lack of long-term follow-up data complicates the diagnosis and identification. Therefore, the diagnosis, treatment and prognosis of primary intracranial SS need to be studied in further cases.

## CONCLUSION

Primary intracranial synovial sarcoma is a remarkably rare malignant tumor. Primary intracranial SS with hemorrhage and radiation necrosis should be carefully monitored during postoperative radiotherapy. Surgical resection of the tumor combined with postoperative radiotherapy and chemotherapy is currently used, but the prognosis is poor.

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