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ABOUT COVER
Editorial Board Member of World Journal of Clinical Cases, Dr. Romano is Professor of Medicine-Gastroenterology at the University of Campania “Luigi Vanvitelli” in Naples, Italy. Dr. Romano received his MD degree cum Laude at the University Federico II in Naples, Italy in 1980 and, after 4 year of Post-Graduate course, he became Specialist in Gastroenterology and Gastrointestinal Endoscopy. Dr. Romano’s research interest was on the cross-talk between H. pylori and gastric epithelial cells, and presently is mainly focused on H. pylori eradication therapy and on the role of nutraceuticals in gastrointestinal diseases. Dr. Romano is presently the Chief of the Endoscopy and Chronic Inflammatory Gastrointestinal Disorders Unit, and Teacher at the University of Campania “Luigi Vanvitelli” in Naples, Italy.

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Imaging of hemorrhagic primary central nervous system lymphoma: A case report

Ya-Wei Wu, Jin Zheng, Lu-Lu Liu, Jun-Hui Cai, Hu Yuan, Jing Ye

ORCID number: Ya-Wei Wu 0000-0001-9925-4729; Jin Zheng 0000-0001-8183-9817; Lu-Lu Liu 0000-0002-2734-7488; Jun-Hui Cai 0000-0002-3569-9806; Hu Yuan 0000-0001-5981-6037; Jing Ye 0000-0003-2756-9988.

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Abstract

BACKGROUND
A primary central nervous system lymphoma (PCNSL) presenting with massive hemorrhage is a rare occurrence that is difficult to distinguish from a high-grade glioblastoma. Comprehensive descriptions of the imaging characteristics of such tumors have not yet been reported. Herein, we reported a case of a PCNSL with massive hemorrhage by presenting the imaging features of computed tomography (CT) imaging and structural and perfusion magnetic resonance imaging (MRI).

CASE SUMMARY
A 48-year-old man presented with headache lasting for 10 d. CT of the brain showed a round, heterogeneous, high-density lesion with surrounding edema in the right temporal lobe. For further diagnosis, a series of MRI examinations of the brain were subsequently performed, and a hemorrhagic lesion with ring-like enhancement was determined. The whole lesion was relatively hypoperfused on arterial spin labeling images. Surgical resection of the lesion and histopathological examination confirmed that the lesion was a diffuse large B-cell lymphoma with massive hemorrhage.

CONCLUSION
PCNSLs with hemorrhage occur very rarely, and structural and perfusion MRI examinations are requested exceedingly rarely. This case provided insight into some characteristics of a hemorrhagic lymphoma on CT and MRI examinations. Perfusion MRI examination may be useful for the differential diagnosis of PCNSLs and other brain tumors.

Key words: Primary central nervous system lymphoma; Massive hemorrhage; Perfusion; Multimodal magnetic resonance imaging; Lymphoma; Case report
INTRODUCTION

Primary central nervous system lymphoma (PCNSL) is a relatively rare tumor that accounts for approximately 2%-6% of all primary brain tumors. The majority of intracerebral lymphomas are non-Hodgkin’s lymphomas, and approximately 90% are diffuse large B-cell lymphomas[1,2]. PCNSL is less common in immunocompetent individuals. However, due to the increasing prevalence of human immunodeficiency virus infection and the growing number of organ transplantations, the incidence of PCNSL has been increasingly observed in both immunocompromised patients and the immunocompetent population over the last decades[3]. Mild to moderate edema and space-occupying effects can always be seen in PCNSL[4], but massive hemorrhage at presentation in PCNSL is extremely rare. The presence of hemorrhage is utilized to exclude primary cerebral lymphoma in the differential diagnosis[5].

The clinical treatment for PCNSLs is different from that of other tumors, and an early and accurate diagnosis is vital to improve the treatment outcomes. Herein, we present a particular case of PCNSL with multimodal magnetic resonance imaging (MRI) examinations. The purpose of this study was to provide insight into some MRI characteristics of hemorrhagic lymphomas and to facilitate the differentiation between PCNSLs and other brain tumors.

CASE PRESENTATION

Chief complaints

A 48-year-old man was admitted to a local hospital with a 10 d history of headache and dizziness, followed by aggravation of these symptoms for 2 d.

History of present illness

None.

History of past illness

The patient was previously in good condition and had no history of congenital or acquired immunodeficiency.

Laboratory examinations

A series of routine examinations before the surgical operation were unremarkable, including: (1) Routine blood testing of serum levels of platelet, alpha fetoprotein, Carcinoembryonic antigen, CA199, neuron-specific enolase and progastrin releasing peptide; and (2) Cerebrospinal fluid analysis.

Imaging examinations

Non-contrast-enhanced computed tomography (CT) examination was performed immediately after admission, and a round and high-density lesion approximately 56 mm × 46 mm in size was found in the right temporal lobe surrounded by moderate
edema (Figure 1A). It compressed the right basal ganglia and the right lateral ventricle, forcing a midline shift towards the left. For further diagnosis, CT angiography examination was performed, and the possibilities of vascular malformation and aneurysms were excluded. Multimodal MRI examinations of the brain were conducted on the same day. The gross appearance of the lesion was heterogeneous on MR images. The parenchyma of the lesion showed iso- to hypointensity on T1-weighted (T1W) and T2-weighted images, which can be interpreted as a relatively hyperintense signal on diffusion-weighted images (DWI). Contrast-enhanced T1W images showed the lesion with ring-like enhancement. The gross lesion displayed relatively low perfusion on arterial spin labeling (ASL) images (Figure 1).

**FINAL DIAGNOSIS**

According to the radiological findings, the initial imaging impression was glioblastoma combined with hemorrhage. Surgery was performed 3 d after admission to the hospital. The lesion was completely resected, and the hematoma (approximately 30 mL) was also removed. The parenchyma of the lesion was 30 mm × 25 mm × 20 mm in size; it looked like rotten fish, was yellow-gray in color and accompanied by some blood vessels.

Histopathological examination revealed that the nuclei of the tumor cells were round or elliptic, with prominent nucleoli, diffuse arrangement and necrotic tissue formed in the lesion. In the immunohistochemical stains, cells were positive for B-cell markers (CD20 and Pax-5; Figure 2), and the Ki-67 index was 70%. The lesion was identified as a diffuse large B-cell lymphoma with acute hemorrhage.

**TREATMENT**

The lesion was completely resected.

**OUTCOME AND FOLLOW-UP**

The patient was treated with chemotherapy following surgery. He had no imaging findings of recurrence at 6 and 12 mo after treatment.

**DISCUSSION**

Even though PCNSLs are relatively rare, and represent 1%-2% of all primary CNS malignancies, their incidence has risen over recent years[3]. In immunocompetent populations, the median age of PCNSL occurrence is 53-years-old to 57-years-old, with a male to female ratio of 1.5:1[6]. PCNSLs can occur in the brain parenchyma, meninges, eyes or spinal cord. Approximately 70% are restricted to the supratentorial brain[5]. The most common presentation of a PCNSL is a single intracranial mass. The clinical manifestations of PCNSLs are similar to those of other intracranial tumors, including high intracranial pressure and focal neurological deficits. To date, six cases of lymphoma with hemorrhage have been reported. Massive bleeding hemorrhage at the first presentation of lymphoma has been reported in only three cases thus far, and this is the only case with detailed structural and perfusion MRI examinations.

In this case, the structural MR images revealed that the parenchyma of the lesion was consistent with a previous study on PCNSLs. PCNSLs always present with a defined margin and are surrounded by mild to moderate edema[2,7]. They appear as homogeneous, iso- to high-density lesions on CT images and as iso- to hypointense lesions relative to the gray matter in T1W and T2-weighted images due to the hypercellularity of the lymphomatous deposits. Approximately 85% of lesions exhibit homogeneous enhancement both on CT and MRI following contrast administration[8]. Ring-like enhancement is rarely seen unless necrosis occurs in the center of the mass, which can always be seen in acquired immunodeficiency syndrome-related PCNSLs[2,8]. On perfusion images, the lesions show relatively lower perfusion as a whole. The proliferation pattern of lymphomas includes vasocentric growth, and vessels in lymphomas are few and small. Compared with the fulminant
Figure 1  Brain computed tomography and magnetic resonance imaging showed a heterogeneous hyperintense signal in the right temporal lobe. A: Non-contrast-enhanced computed tomography images showed massive hemorrhage (orange star) with perilesional edema; B: The diffusion-weighted image (b = 1000 mm/s) demonstrates a relatively hyperintense signal of the parenchyma (orange arrow); C: T1-weighted image; D: T2W image showing the parenchyma (orange arrow) of the lesion as iso- to hypointense; E: Contrast-enhanced T1-weighted image revealed a ring-like enhancing pattern; and F: Arterial spin labeling showed relatively low perfusion of the whole lesion.

Figure 2  Histological examination and immunohistochemical analysis of the lesion. A: Hematoxylin-eosin staining revealed diffuse lymphoid cells with prominent nuclei; B, C: Immunohistochemistry analysis of the specimen shows positive staining of the cells for CD20 and Pax-5.

neovascularization of glioblastomas, the blood flow of PCNSLs is relatively lower[4].

Recently, some advanced imaging techniques, such as ASL perfusion imaging and DWI, which respectively reflect tumor vascularity and cellularity, have been successfully applied for the differential diagnosis between lymphomas and other brain tumors[10]. The majority of PCNSLs demonstrate relatively lower perfusion in ASL images and higher intensity on DWI images than other brain tumors[7,11], which is consistent with our findings. As previous studies have shown, ASL and DWI are potential diagnostic tools for differentiating PCNSL from glioblastoma due to their differing hemodynamics and tumor density[12,13].

Hemorrhage is very rarely observed in untreated CNS lymphoma[5,14]. The mechanism of the occurrence of primary lymphoma with hemorrhage remains unclear, and it could be potentially explained by high immunoreactivity for a vascular endothelial growth factor may account for it[5,13,14]. Another explanation is that fragile vessels
traversing necrotic areas or tumor invasion of large vessels lead to the breakdown of the vessel wall, resulting in bleeding[5].

CONCLUSION

Accurate diagnosis of lymphoma is essential in clinical practice and is related to therapeutic decision-making and the patients’ prognosis, whereas the diagnosis of atypical PCNSLs is difficult. We presented a special case of a PCNSL with acute massive intracerebral hemorrhage: a report of 58 cases. 10.1054/jocn.2002.1131

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