CASE REPORT

A.I. Alomari
P. Jiarakongmun
C. Dejthevaporn
S. Pongpech

Multiple Symptomatic Cerebral Arteriovenous Malformations in a Patient with HIV

SUMMARY: This brief case report describes an unusual presumed association of chronic HIV infection with multiple cerebral arteriovenous malformations. An adult male patient presented with recurrent spontaneous brain intracranial hemorrhages and neurologic deficits requiring surgical evacuation. The diagnosis was delayed because of lack of a high index of suspicion of this unexpected and previously unrecognized association.

HIV-infected patients are at a higher risk for stroke, both ischemic and hemorrhagic. However, symptomatic multiple cerebral arteriovenous malformations are not a reported presentation of HIV.

We herein report the clinical and imaging findings in an adult with HIV and multiple arteriovenous shunts affecting both cerebral hemispheres with 3 documented episodes of brain hemorrhage.

Case Report

The institutional ethical clearance committee granted approval for publishing this case report. A 45-year-old Thai man known to have HIV infection, diabetes, and hypertension presented with a sudden onset of motor weakness. Neurologic examination revealed right hemiparesis and hyperreflexia. An urgent head CT scan showed an acute hematoma in the left frontoparietal region with subarachnoid and intraventricular bleeds. The left cerebral hemisphere appeared diffusely edematous with some subfalcine herniation. Postcontrast images showed multiple scattered enhancing foci bilaterally within the cerebral hemispheres. The differential diagnoses entertained in light of the patient's history included spontaneous bleed, opportunistic infection, and metastatic lesions. Laboratory results showed mild anemia (hemoglobin, 10.4–13.7 g/L) and thrombocytopenia (platelet count, 65,000–123,000/ μ L). Otherwise, results of liver function tests, renal function tests, coagulation tests, and lipid profiles were within normal ranges. The hematoma was urgently evacuated.

Medical history was significant for HIV infection contracted at age 33 years. After contracting *Pneumocystis carinii* infection at age 38 years, the patient was initiated on a triple antiretroviral regimen and remained without any AIDS-defining findings, with CD4+ T cell counts of 300 to 500 cells/ μ L. The patient also had a 10-year history of noninsulin-dependent diabetes (on glibenclamide), essential hypertension, and smoking (10-pack-year).

Follow-up enhanced CT and MR imaging scans at 1 month demonstrated resolution of the parenchymal hematoma and some postoperative changes. Several parenchymal hypodensities were also noted in both hemispheres, suggestive of previous insults, possibly of the same etiology. In addition, there were some small adjacent enhancing foci around these lesions, best seen after contrast administra-

tion. Brain MR imaging study confirmed the presence of small flow voids within the white matter of both cerebral hemispheres and the left thalamus (Fig 1A). Gradient-echo sequence revealed hypointense signal intensity adjacent to these lesions, presumably because of previous bleeds and hemosiderin deposits. Brain MR venography showed small venous pouches associated with dilated cortical veins (Fig 1B).

The patient had 2 additional episodes of right hemispheric bleeds at 8 and 19 months after discharge and eventually he agreed to undergo a cerebral angiography. Under conscious sedation, 5-vessel cerebral angiography (bilateral internal carotid artery, bilateral external carotid artery and left vertebra artery) was uneventfully performed. The study showed multiple small arteriovenous shunts within the temporal, parietal, and frontal lobes (Fig 2A, -B), representing deeply seated pial microarteriovenous malformations (AVMs) within the brain parenchyma. There were multiple feeding arteries from the anterior and middle cerebral arteries. There was a complex pattern of draining veins associated with multiple focal aneurysms. The vertebral and the external carotid circulations were normal.

The patient is currently scheduled to undergo staged embolization of the AVMs, primarily targeting the larger venous aneurysms to reduce the risk for additional hemorrhage.

Discussion

Cerebral AVMs are typically solitary lesions. Multiplicity, which is more frequently seen in association with syndromes, such as hereditary hemorrhagic telangiectasia and Wyburn-Mason syndrome, is rarely encountered in sporadic cases.¹⁻³

HIV-related neurologic complications (HIV-RNC) are common, affecting approximately 40% of patients.⁴ The complex cause of most HIV-RNC includes non-HIV CNS infection, CNS lymphoma, cardioembolic conditions, drug misuse, and cerebral infarcts occurring in the absence of these conditions.⁵ However, intracerebral hemorrhage is a rare (1%), late, and serious complication of AIDS with a mortality rate of 62.5%.⁶ In children, the risk is primarily related to coagulopathy, cardioembolism, vascular ectasia or aneurysms, subacute necrotizing encephalopathy, and arteriopathy.^{7,8}

HIV-associated cerebral vasculopathy was proposed as an important mechanism of stroke in the absence of a known cause, possibly related to deposition of a circulating immune complex or a direct toxic effect on the vascular endothelium.⁹ Kure et al¹⁰ demonstrated the presence of HIV within the arterial wall of the circle of Willis and in the cerebral parenchyma in a child who died of HIV-related cerebral vascular ectasia and multiple infarcts. The vascular histopathologic changes in HIV-infected patients include medial fibrosis, loss of muscularis with destruction of the internal elastic lamina, and intimal hyperplasia.¹¹

We only found 1 case report by Wolf et al¹² describing a

Received November 18, 2008; accepted after revision November 25.

From the Division of Vascular and Interventional Radiology (A.I.A.), Department of Radiology, Children's Hospital Boston, Boston, Mass; Department of Radiology (P.J., S.P.), Ramathibodi Hospital, Mahidol University, Bangkok, Thailand; and Division of Neurology (C.D.), Department of Medicine, Ramathibodi Hospital, Mahidol University, Bangkok, Thailand.

Please address correspondence to Ahmad I. Alomari, MD, Division of Vascular and Interventional Radiology, Children's Hospital Boston, 300 Longwood Ave, Boston, MA 02115; e-mail: ahmad.alomari@childrens.harvard.edu

DOI 10.3174/ajnr.A1475

Fig 1. A, Axial T2-weighted MR imaging of the brain. Evidence of previous parenchymal bleeding of different ages is seen in both cerebral hemispheres (white arrows). Multiple flow voids are seen within the white matter (black bent arrows), subependymal area (black arrow), and gray-white matter junction (white bent arrow). B, MR venography shows small venous aneurysms draining into slightly dilated cortical veins (white arrows).

Fig 2. Cerebral angiography of the internal cerebral arteries. Arterial phase of the right (A, lateral projection) and left (B, frontal projection) depicting the presence of multiple pial arteriovenous nodi with associated venous aneurysms (transparent arrows) and early opacification of the draining veins (arrowheads) and dural sinuses (black arrows). C, 3D rotational angiography, arterial phase, of the right internal cerebral artery. Note the early opacification of the venous aneurysm (bent arrows) and draining veins (straight arrow).

53-year-old patient with HIV infection who had a right Rolandic cerebral AVM leading to recurring focal seizures. There was a single draining vein with a large varix and a flow-related arterial aneurysm. After 2 years of highly active antiretroviral therapy, the AVM was completely and spontaneously occluded. The authors hypothesized that the suppression of the angiogenic activity of HIV led to resolution of the shunt.

The case presented in our report is unusual for many reasons. First, the presence of multiple cerebral AVMs is rare. Second, the occurrence of multiple episodes of cerebral hemorrhage secondary to multiple micro-AVMs has never been reported before in HIV. Third, most of the cerebrovascular complications in the literature were reported in patients with advanced immunosuppression. Our patient maintained a moderate immunosuppression status while receiving triple antiretroviral therapy without signs of AIDS. In contrast to the multiple cerebral AVMs seen with hereditary hemorrhagic telangiectasia, which are superficially located within the cerebral cortex and are not associated with hemorrhage or venous pouches, the lesions encountered in our patient were deeply situated lesions within the cerebral parenchyma and developed multiple hemorrhages.

In conclusion, this unusual case highlights a rare, previously unreported complication presumably related to a long-standing HIV infection. Cerebral AVM, though rare, might be considered in the differential diagnosis of acute cerebral hemorrhage in patients with HIV within the proper context and prompt a proper diagnostic and therapeutic approach. Early

diagnosis of this rare manifestation of HIV could potentially prevent a disabling and potentially lethal course.

References

1. Kawaguchi T, Fujita S, Yamada H, et al. Multiple cerebral and pulmonary arteriovenous malformations associated with brain and subcutaneous abscesses: a possible variant of hereditary hemorrhagic telangiectasia, case report. *Neurol Med Chir (Tokyo)* 1990;30:272–76
2. Matsubara S, Mandzia JL, ter Brugge K, et al. Angiographic and clinical characteristics of patients with cerebral arteriovenous malformations associated with hereditary hemorrhagic telangiectasia. *AJNR Am J Neuroradiol* 2000;21:1016–20
3. Willinsky RA, Lasjaunias P, Terbrugge K, et al. Multiple cerebral arteriovenous malformations (AVMs). Review of our experience from 203 patients with cerebral vascular lesions. *Neuroradiology* 1990;32:207–10
4. Pinto AN. AIDS and cerebrovascular disease. *Stroke* 1996;27:538–43
5. Connor MD, Lammie GA, Bell JE, et al. Cerebral infarction in adult AIDS patients: observations from the Edinburgh HIV Autopsy Cohort. *Stroke* 2000;31:2117–26
6. Roquer J, Palomeras E, Knobel H, et al. Intracerebral haemorrhage in AIDS. *Cerebrovasc Dis* 1998;8:222–27
7. Narayan P, Samuels OB, Barrow DL. Stroke and pediatric human immunodeficiency virus infection. Case report and review of the literature. *Pediatr Neurosurg* 2002;37:158–63
8. Shah SS, Zimmerman RA, Rorke LB, et al. Cerebrovascular complications of HIV in children. *AJNR Am J Neuroradiol* 1996;17:1913–17
9. Stapf C, Mohr JP, Pile-Spellman J, et al. Epidemiology and natural history of arteriovenous malformations. *Neurosurg Focus* 2001;11:e1
10. Kure K, Park YD, Kim TS, et al. Immunohistochemical localization of an HIV epitope in cerebral aneurysmal arteriopathy in pediatric acquired immunodeficiency syndrome (AIDS). *Pediatr Pathol* 1989;9:655–67
11. Dubrovsky T, Curless R, Scott G, et al. Cerebral aneurysmal arteriopathy in childhood AIDS. *Neurology* 1998;51:560–65
12. Wolf K, Kirsch EC, Sponagel L, et al. Disappearance of an intracerebral arteriovenous malformation in an HIV-infected patient after initiation of HAART. *Infection* 2002;30:101–03