Image Report

Multiple cerebral hemorrhages in sepsis-disseminated intravascular coagulation versus septic embolism: An image report

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ABSTRACT

Background: Septic emboli are commonly attributed to infective endocarditis and can present with a variety of symptoms including altered mental status and focal neurological deficits. Here, we reviewed images of septic emboli with hemorrhagic conversion in a patient with sepsis and a psoas abscess. We aim to show the classical image findings in septic embolism to brain, which is sparsely described in literature and the report differentiates the septic embolism from disseminated intravascular coagulation which can present with almost identical image findings.

Case Description: A 53-year-old male patient who was operated on for a right inguinal hernia developed a postoperative wound infection 2 weeks after surgery and was started on IV antibiotics. Despite medical management, his infection did not improve, prompting a computed tomography (CT) scan which revealed a psoas abscess. The abscess was drained, and antibiotics continued. A few days later, he developed altered sensorium prompting a head CT which revealed septic emboli and hemorrhage at the gray-white junction. Cultures grew multidrug-resistant Escherichia coli; the patient was treated with IV tigecycline and improved over the following 4 weeks.

Conclusion: In patients with a known ongoing infectious process with hemodynamic stability who develop altered mental status in the setting of a normal coagulation profile, D-dimer, positive blood cultures, and absent signs of multiorgan failure, a diagnosis of septic emboli should be entertained. Although CT can reveal macrobleeds, MRI is more sensitive in confirming cerebral microbleeds. Thus, patients in sepsis with unexplained altered sensorium should undergo an MRI of the brain to rule out septic emboli and microbleeds.

Keywords: Cerebral bleed, Disseminated intravascular coagulation, Psoas abscess, Sepsis, Septic embolism

INTRODUCTION

Septic embolism to the brain from an infective focus is well known. It can present with micro or macro hemorrhages or abscesses involving the cortical grey and white matter junction. The hemorrhagic type of septic embolism can be confused or overlapping with the multiple cerebral hemorrhages of DIC. Here, we compare the imaging findings of septic embolism and DIC with an illustrative case.

CASE REPORT

A 53-year-old male patient who was operated on for a right inguinal hernia subsequently developed a postoperative wound infection 2 weeks after surgery and was started on IV
antibiotics. Despite medical management, the patient's wound discharge did not improve and therefore a computed tomography (CT) scan was conducted which revealed a right psoas abscess. The abscess was partially drained, and he was continued on IV antibiotics. After a few days, the patient demonstrated progressive deterioration of his sensorium. Biochemical and hematological parameters revealed hemoglobin: 14 g/dL, WBC: 18,000 cells/mcL, platelet count: 280,000 cells/mcL, peripheral smear: normocytic and normochromic, neutrophils showing multiple toxic granules, no hemolysis or schistocytes, CRP:14 mg/L, INR: 0.9, D-dimer: normal, and fibrinogen: 2.4 g/suggestive of sepsis. Based on the imaging findings, a diagnosis of septic embolism to the brain was made. Culture sensitivity of the purulent discharge revealed a multidrug-resistant Escherichia coli and was managed with IV tigecycline (100 mg IV, followed by 50 mg IV q12 hourly for 14 days). Over the next 4 weeks, the patient improved dramatically, and repeat imaging showed significant resolution of the hemorrhages.

Imaging findings

CT abdomen with contrast showed an enlarged right psoas muscle with multiple hypodense loculations enhancing peripherally on contrast suggestive of a psoas abscess. CT brain imaging showed diffuse multiple micro- and macro-hemorrhages of varied sizes involving the cortical gray-white matter junction [Figure 1]. However, there were no hemorrhages within the cerebellum or deep nuclei.

Differential diagnoses of multiple cerebral microbleed (CMB)

The detection of CMB alone is nonspecific and can be found incidentally, thus clinical correlation is imperative. The two most common causes are hypertensive small vessel disease and congophilic amyloid angiopathy (CAA). In hypertensive small-vessel disease, CMBs are found in the basal ganglia, thalamus, brainstem, and cerebellum. In contrast, bleed in CAA tends to occur at the corticomedullary junction and can lead to lobar hemorrhage.

Less common causes include diffuse axonal injury leading to microhemorrhages near the gray-white matter junction. In patients with a known history of malignancy, hemorrhagic micrometastasis from melanoma and renal cell carcinoma can also cause microbleeds. CMBs can also be associated with critical medical illnesses such as sepsis and infective endocarditis; hereditary and idiopathic diseases such as cerebral autosomal dominant arteriopathy with subcortical infarcts and leukoencephalopathy, Fabry disease, Moyamoya disease, and radiation-induced vascular injury.

DISCUSSION

Definition of micro- and macrobleed

Various cutoffs have been utilized when describing CMB. CMBs have conventionally been defined in the literature as having a maximum of about 5–10 mm, with a minimum of 2 mm.

Differentiating disseminated intravascular coagulation (DIC) versus septic embolism

A septic embolism is the obstruction of a blood vessel by an infectious nidus, which causes occlusion of the vessel and leads to ischemia and infarction. Septic emboli present a unique challenge because the impact is 2-fold: the ischemic insult caused by the occlusion of the vessel and the infectious/inflammatory insult leading to erosive vasculitis, mycotic aneurysms, or end-organ abscess formation. The overwhelming inflammatory response associated with sepsis may lead to hemostatic changes, most notably DIC, which is characterized by microvascular thrombosis, consumption of platelets and coagulation proteins, and in most severe cases,
bleeding and multiorgan dysfunction.\textsuperscript{13} Diagnosing DIC in the setting of sepsis is critical, and several scoring systems have sought to quantify the severity of DIC. Laboratory values suggestive of DIC include prolonged prothrombin time (PT), reduction in platelet count, reduced fibrinogen levels, and elevated D-dimer.\textsuperscript{15}

**Pathogenesis of bleed in septic embolism and DIC**

The pathogenesis of massive bleeding in DIC results from hyperfibrinolysis leading to consumption of clotting factors; without adequate replacement, massive bleeding occurs. Among the four types of DIC, fibrinogen, PT, and platelets are important parameters for diagnosis of the consumptive hemorrhagic type. While no single laboratory finding is diagnostic, the collective trend of the parameters discussed suggests the diagnosis.\textsuperscript{15} In the setting of septic embolisms, hemorrhage can also occur secondary to ischemia and hemorrhagic transformation, but often present as petechial hemorrhage as opposed to larger parenchymal hematomas.\textsuperscript{17} Sepsis in and of itself may contribute to microcirculatory changes, leading to endothelial dysfunction, loss of smooth muscle regulation, increased platelet aggregation, and increased aggregation of red blood cells to endothelial cells.\textsuperscript{10,11} Furthermore, intra-CMBs have been reported in patients with sepsis without evidence of septic emboli or DIC, highlighting the potential clinical implication of microcirculatory dysfunction.\textsuperscript{4} Mycotic aneurysms, although rare, are another potential cause of bleeding in the setting of septic emboli. However, these present with subarachnoid hemorrhage rather than multiple parenchymal bleeds.\textsuperscript{5}

**MRI versus CT findings in septic embolism**

In evaluating a patient with altered mental status and/or signs and symptoms of neurological insult in sepsis, the imaging modality of choice will depend on the suspected lesion as well as acuity. CT scans have shown a high incidence in detecting cerebral lesions; this incidence, however, is even higher with the use of MRI.\textsuperscript{4}

Septic emboli as previously discussed can lead to ischemic strokes with hemorrhagic transformation, mycotic aneurysms, and focal arteritis. In identifying hemorrhagic lesions, such as lobar hemorrhage and hemorrhagic transformation of an ischemic insult, MRI has showed limited additional value. MRI angiography is also as effective as CT angiography in diagnosing these microbial aneurysms when the diameter is >5 mm.\textsuperscript{2}

MRI use in the setting of sepsis, DIC, and/or septic emboli has shown to be most useful when diagnosing small neurological lesions, such as microinfarcts and CMBs that would otherwise go undetected. One example is the use of susceptibility-weighted MRI in evaluating unexplained neurological and cognitive dysfunction in the context of DIC secondary to sepsis. Neligan et al. reported widespread microhemorrhages in the cortex, subcortex, brainstem, and cerebellum in a septic patient with sickle cell disease presenting in DIC, identifying the etiology of their patient’s neurological deterioration.\textsuperscript{12} Another value of brain MRI is guidance for medical management. Lesions indicative of meningitis, microabscesses, microischemia, or CMBs may guide antibiotic therapy, as evidence of central nervous system infections require antimicrobial drugs with high cerebral diffusion. This can alter management leading to changes in therapy or prognosis.\textsuperscript{2}

**Treatment**

Septic emboli to the brain from any source can travel to the cerebral circulation leading to embolic infarcts, cerebral hemorrhages, mycotic aneurysms, and cerebral abscesses.\textsuperscript{17} Adequate antimicrobial therapy remains the treatment of choice. The source of infection and identification of the infectious agent influence the choice of antimicrobial therapy.

Anticoagulation has been a topic of controversy and is avoided with evidence of hemorrhage; to the best of our knowledge, no randomized control trials have been conducted to assess the role of anticoagulation in the setting of septic emboli.\textsuperscript{16} Embolectomy is an option especially if the embolism is involving major blood vessels like the middle cerebral artery. The benefit of this intervention is subject to controversy and requires evaluation through randomized controlled trials.\textsuperscript{9} However, this has no role in patients like ours with multiple parenchymal bleeds.

**CONCLUSION**

In patients with a known ongoing infectious process with hemodynamic stability who develop altered mental status in the setting of a normal coagulation profile, D-dimer, positive blood cultures, and absent signs of multiorgan failure, a diagnosis of septic emboli should be entertained. Although CT can reveal macrobleeds, MRI is more sensitive in confirming CMB. Thus, patients in sepsis with unexplained altered sensorium should undergo an MRI of the brain to rule out septic emboli and microbleeds.

**Declaration of patient consent**

Patient’s consent not required as patients identity is not disclosed or compromised.

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Conflicts of interest

There are no conflicts of interest.

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