Cerebral Venous Thrombosis
Diagnosis and Management

Deepak Gulati, MD; Daniel Strbian, MD, PhD; Sophia Sundararajan, MD, PhD

A 42-year-old woman with a history of Crohn disease presented with left temporal headache for 1 week. On the day of presentation, she had transient word finding difficulty, lasting for 1 to 2 minutes and worsening headache. The patient reported starting prednisone 10 days earlier for a flare of Crohn’s disease associated with dehydration. She had been using the Nuva ring (ethinyl estradiol vaginal ring) for birth control. Neurological examination was significant for moderate expressive aphasia. Emergent computed tomography (CT) head revealed a hyperdense focus with surrounding hypodensity in the left temporoparietal lobe. Subsequent MRI of the brain revealed a venous infarct in the anterior left temporal lobe with surrounding edema and areas of hemorrhagic transformation. Magnetic resonance venography (MRV) showed left transverse and sigmoid thrombosis. Hypercoagulable studies were sent including Protein C & S, factor V leiden, prothrombin G20210A gene mutation, lupus anticoagulant, anticardiolipin antibody, homocysteine, and antithrombin III. Systemic anticoagulation (unfractionated heparin drip) was started immediately. Prednisone was continued for Crohn disease. For the next 2 days, her aphasia improved. At the time of discharge, she was transitioned to warfarin with low molecular weight (LMW) heparin bridge. After 1 month, the aphasia and headache completely resolved. Hypercoagulable workup revealed that she was heterozygous for prothrombin G20210A gene mutation, whereas other laboratory tests were unremarkable. Repeat MRV at 3 months showed partial recanalization of the left transverse sinus.

Cerebral venous and dural sinuses thrombosis (CVT) is an uncommon and frequently unrecognized, representing ≈0.5% to 1% of all strokes.1 CVT is potentially serious and life threatening.

Clinical Findings
Symptoms in CVT are usually secondary to increased intracranial pressure or focal brain injury from venous infarction or hemorrhage. Headache is often localized and is the most frequent and usually the first symptom. Isolated headache without focal neurological findings or papilledema occurs in 25% of patients with CVT and poses a significant diagnostic challenge.2,3 Other clinical symptoms are encephalopathy, focal neurological signs, and seizures. Clinical presentation depends on several factors, including location of the thrombosis, the presence of venous infarction or hemorrhage, patient’s age, and duration of CVT (acute versus chronic). Superior sagittal sinus thrombosis is the most common form and usually leads to bilateral deficits. Focal or generalized seizures are frequent, occurring in ≈40% of patients with CVT.3

Risk Factors
Risk factors for CVT are those that affect blood stasis, changes in the vessel wall, and changes to the composition of blood (Virchow triad). Both genetic and acquired prothrombotic conditions can contribute to CVT. American Heart Association guidelines define antithrombin III, protein C and protein S deficiency, homozygosity for either factor V leiden or prothrombin G20210A mutations as high-risk thrombophilias.3,4 Antithrombin III, protein C and S testing acutely is of limited use in the acute setting and should be tested 2 to 4 weeks after anticoagulation has been stopped.1 Heterozygosity for factor V leiden or prothrombin G20210A mutation is considered a relatively mild thrombophilia but still carries an increased risk.3 More recently, an association between the JAK 2 V617F gene and an increased incidence of CVT has been reported.5 In addition to genetic prothrombotic states, acquired prothrombotic states should be investigated. The more common of these include oral contraceptives, pregnancy, puerperium, malignancy, infection, and trauma.6 In developed countries, the most common cause of CVT is pregnancy/puerperium, whereas parameningeal infections (ear, sinus, mouth, face, and neck) remain relatively common causes in developing countries. The relationship between CVT and inflammatory bowel disease is well established, although the mechanism is poorly understood. The association between the 2 entities may be secondary to dehydration, or the inflammatory state may predispose the patient to clot formation. Only ≈3% of CVT occurs in patients with inflammatory bowel disease,7 and it is usually associated with other risk factors. Oral contraceptives are associated with a ≈22-fold risk of CVT. The risk of CVT with oral contraceptive

Received October 31, 2013; accepted November 11, 2013.
From the Department of Neurology, University Hospitals of Cleveland/Case Medical Center, OH (D.G., S.S.); and Departments of Neurology and Stroke Units, Helsinki University Central Hospital, Helsinki, Finland (D.S.).
Correspondence to Sophia Sundararajan, MD, PhD, Department of Neurology, University Hospitals of Cleveland/Case Medical Center, 11100 Euclid Ave, Cleveland, OH 44106; E-mail sophia.sundararajan@UHhospitals.org

(Stroke. 2014;45:e16-e18.)

© 2013 American Heart Association, Inc.

Stroke is available at http://stroke.ahajournals.org

DOI: 10.1161/STROKEAHA.113.003964
use in women is greater among those with a hereditary prothrombotic factor. Although the risk of venous thrombosis is thought to be less with vaginal rings that release low progestin/estrogen, a recent study found that users of vaginal rings with ethinylestradiol and etonogestrel have a 6.5-fold increase in venous thrombosis when compared with nonusers.

Importantly, CVT is often multifactorial and, therefore, every patient requires a full workup. Our patient is an excellent example of this. She has a previously unrecognized prothrombin gene mutation and had a flare of her inflammatory bowel disease. In addition, she was using a vaginal ring for contraception. Other potential triggers for CVT in patients with an underlying procoagulable predisposition include head trauma, lumbar puncture, jugular catheter placement, pregnancy, surgery, infection, or drugs.

**Neuroimaging**

CT of the head is often the first investigation performed. The primary sign of acute CVT on noncontrast CT is hyperdensity of a cortical vein or dural sinus. Thrombosis of the posterior portion of the superior sagittal sinus may appear as a dense triangle, the dense or filled delta sign. CT venography is most useful in subacute or chronic situations because of varied density in a thrombosed sinus. MRV is usually preferred to CT venography because of bone artifact and radiation exposure. MRI is more sensitive for CVT than CT at every stage of thrombosis. The characteristics of the MRI signal depend on the age of the thrombus. The principal early signs of CVT on non–contrast-enhanced MRI are the combination of absent flow voids with altered signal intensity within the dural sinus. Invasive cerebral angiographic procedures are rarely needed to establish the diagnosis of CVT, given the availability of MRV and CT venography (CTV). Catheter angiography is usually considered only when the diagnosis is not clear, despite MRV, or if an endovascular procedure is being considered.

**Management**

Treatment should be started as soon as the diagnosis is confirmed. Management of CVT consists of (1) identification and treatment, of the underlying cause, (2) symptomatic management, and (3) antithrombotic therapy. Underlying infection, dehydration, or other correctable precipitants should be investigated and aggressively treated to reduce the prothrombotic state. In our patient, intravenous fluids were given to treat dehydration, and her prednisone was continued to treat the underlying inflammation associated with Crohn’s disease. Seizures are a common complication of venous thrombosis and there should be a low threshold for electroencephalogram in patients with unusual behavior or reduced level of consciousness. American Heart Association guidelines recommend the use of anticonvulsants for seizures but not for prophylactic treatment. Patients may have increased intracranial pressure and treatment using standard measures may be life saving. Patients frequently have severe headache and may require analgesics, but care should be taken not to sedate the patient to the point that the examination becomes unreliable. Anticoagulation is the mainstay of acute and subacute treatment for CVT. Because heparin can influence the

---

**Figure.** Cerebral venous thrombosis (CVT). CTV indicates CT venography; ICH, intracranial hemorrhage; IV, intravenous; MRV, magnetic resonance venography; and SC, subcutaneous.
interpretation of hypercoagulable testing, blood work should be
drawn for hypercoagulable studies before heparin. Initial
anticoagulation is generally with adjusted-dose unfractionated
heparin or weight-based low molecular weight heparin in full
anticoagulant doses (Figure). Several randomized controlled
clinical trials support the use of anticoagulation in CVT, even
in the setting of intracranial hemorrhage. Although no trial
showed clear statistical benefit, it is unlikely that additional
trials will be conducted because of a lack of equipoise. A
recent study suggests that low molecular weight heparin may
be superior to unfractionated heparin. Patients receiving
low molecular weight had a greater likelihood of full recov-
ery and less bleeding complications, including intracerebral
hemorrhage. There are little data from controlled trials to
support endovascular thrombolysis, and the American Heart
Association recommends it be restricted to patients with a
diseases. 

Prognosis

CVT can result in death or permanent disability but usually
has a favorable prognosis. Currently 3% to 15% of patients
die in the acute phase of the disorder, and many patients
make a complete recovery. Predictors of mortality at 30 days are
depressed consciousness, altered mental status, thrombo-
sis of the deep venous system, right hemisphere hemorrhage,
posterior fossa lesions. Available data suggest that cerebral
vein and sinus recanalization occurs in 40% to 90% of patients
after CVT, usually within the first 4 months.

Disclosures
None.

References
2. Crassard I, Bousser MG. Headache in patients with cerebral venous
Cucchiara B, Cushman M, et al; American Heart Association Stroke
Council and the Council on Epidemiology and Prevention. Diagnosis
and management of cerebral venous thrombosis: a statement for health-
care professionals from the American Heart Association/American
4. Einhäuser K, Stam J, Bousser MG, De Bruijn SF, Ferro JM, Martielli I,
et al; European Federation of Neurological Societies. EFNS guideline on
the treatment of cerebral venous and sinus thrombosis in adult patients.
5. Passamonti SM, Biguzzi E, Cazzola M, Franchi F, Giannelli F,
Bucciarelli P, et al. The JAK2 V617F mutation and cerebral venous
thrombosis in inflammatory bowel diseases: eight cases and literature
6. Ferro JM, Canhão P, Stam J, Bousser MG, Barinagarrementeria F;
ISCVT Investigators. Prognosis of cerebral vein and dural sinus throm-
bosis: results of the International Study on Cerebral Vein and Dural Sinus
7. Cognat E, Crassard I, Denier C, Vahedi K, Bousser MG. Cerebral venous
thrombosis in users of non-oral hormonal contraception: follow-up study,
9. Coutinho JM, Stam J. How to treat cerebral venous and sinus thrombosis.
10. Coutinho JM, Ferro JM, Canhão P, Barinagarrementeria F, Bousser
MG, Stam J; ISCVT Investigators. Unfractionated or low-molecular
2010;41:2575–2580.
11. Coutinho JM, Ferro JM, Zuurveld SM, Mink MS, Canhão P, Crassard I,
et al. Thrombolysis or anticoagulation for cerebral venous thrombosis:
12. Dentali F, Crowther M, Ageno W. Thrombophilic abnormalities, oral
contraceptives, and risk of cerebral vein thrombosis: a meta-analysis.
H, et al. Cerebral venous sinus thrombosis: Incidence of venous throm-
14. Canhão P, Ferro JM, Lindgren AG, Bousser MG, Stam J,
Barinagarrementeria F; ISCVT Investigators. Causes and predictors of
15. Baungartner RW, Studer A, Arnold M, Georgiadis D. Recanalisation of

Key Words: cerebral hemorrhage | stroke | venous thrombosis
Cerebral Venous Thrombosis: Diagnosis and Management
Deepak Gulati, Daniel Strbian and Sophia Sundararajan

Stroke. 2014;45:e16-e18; originally published online December 19, 2013;
doi: 10.1161/STROKEAHA.113.003964
Stroke is published by the American Heart Association, 7272 Greenville Avenue, Dallas, TX 75231
Copyright © 2013 American Heart Association, Inc. All rights reserved.
Print ISSN: 0039-2499. Online ISSN: 1524-4628

The online version of this article, along with updated information and services, is located on the
World Wide Web at:
http://stroke.ahajournals.org/content/45/2/e16

Permissions: Requests for permissions to reproduce figures, tables, or portions of articles originally published in Stroke can be obtained via RightsLink, a service of the Copyright Clearance Center, not the Editorial Office. Once the online version of the published article for which permission is being requested is located, click Request Permissions in the middle column of the Web page under Services. Further information about this process is available in the Permissions and Rights Question and Answer document.

Reprints: Information about reprints can be found online at:
http://www.lww.com/reprints

Subscriptions: Information about subscribing to Stroke is online at:
http://stroke.ahajournals.org/subscriptions/