A case of cerebral venous thrombosis following the use of contraceptive medication

Abstract
Cerebral venous thrombosis is an uncommon and serious complication of contraceptive medication, which often masquerades as a simple headache. The clinical picture is often confusing and imaging is critical to the diagnosis of this disorder. This case report illustrates a case of dural venous thrombosis and a review of the radiological features of cerebral venous thrombosis.

Introduction
Venous sinus thrombosis is a rare event and may have an insidious onset and late sequelae in contradistinction to cavernous sinus thrombosis associated with infection and florid signs. Thrombosis involving the dural sinuses is due to three causes: those from changes in blood flow, changes in the vessel wall, or blood abnormalities. Blood flow changes are most commonly seen in dehydration.

Changes in flow and vessel wall are seen in depressed fractures about the midline of the vault. Operative interhemispheric (e.g. transcallosal) procedures near the superior sagittal sinus may produce local trauma and stasis. Another entity is the vasculitis of Behçet's disease. Common, non-infectious causes of cerebral venous thrombosis include oral contraceptives, pregnancy and the puerperium. Abnormal changes in blood constituents are seen in malnutrition, and blood deficiencies antithrombin III, protein S & C, disseminated intravascular coagulation, iron deficiency anaemia.

Occlusion of the anterior third of the superior sagittal sinus does not produce symptoms or signs whereas thrombosis of the middle third produces upper motor neurone signs, hemi- or quadriaparesis. Visual field disturbances or blindness may occur with occlusion of the posterior third of the sinus rapidly followed by decreasing level of consciousness.

Thrombosis of a transverse sinus is less significant than if the opposite sinus is also occluded. Any involvement of the posterior two-thirds is associated with cerebral oedema indicated by engorgement of retinal veins,
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meningism and decreased level of consciousness. In subdural parasalx empyema, venous thrombosis followed by infarction may be seen in neglected cases. As the empyema may be restricted to one side or another of the falx, so the neurological deficit may be unilateral before rapid deterioration as the whole superior sagittal sinus becomes involved.

**Case report**

The patient P.M., a healthy 23-year-old woman, had no previous medical history. She had a child of 5 years of age. A year after delivery, she began with three-monthly injections of norethisterone enantate 200 mg. Soon after starting on this treatment, she developed headaches and amenorrhea. Three months prior to admission, owing to the headaches, she was changed to levonorgestrel 150 mg, ethinyl oestradiol 30 mg oral contraceptive. The change relieved the headaches and there were scanty periods. Approximately two days prior to admission, the patient presented at a peripheral hospital with severe headache, nausea and vomiting, decreased vision and confusion (GCS 7/15) followed by deepening coma. On recovering consciousness, she had a right hemiparesis. Her mental recovery was such as to provide the foregoing history. Computerized tomography (CT) done at the peripheral hospital (Figure 1 and 2) was reported as “a large subdural haematoma (possible venous sinus) occipitally; blood in the quadrigeminal cisterns, possible subarachnoid haemorrhage (SAH), blood in the region of the straight sinus. Post contrast filling defect in the sagittal sinus – possible dural sinus thrombosis”. Because of the reported SAH, the patient was treated with nimodipine. Four-vessel angiography, performed 11 days later (Figure 3), confirmed the diagnosis of venous sinus thrombosis.
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A second CT with contrast (Figure 4) was done 12 days later which showed a clear delta sign and clot within straight sinus. Magnetic resonance (MR) imaging was requested at 18 days and showed typical high signal thrombus (Figure 5) in the dural sinuses. It also showed a left cortical venous infarct.

**Discussion**

The cerebral side effects of the contraceptive medication include non-specific headaches or migraine, withdrawal headaches, benign intracranial hypertension, and cerebral venous thrombosis (CVT). The evaluation of a patient on contraceptive medication, who complains of worsening headaches, requires exclusion of CVT by computed tomography or magnetic resonance imaging.

The manufacturers state that the occurrence for the first time of a migraine type of headache, the more frequent occurrence of an unusually severe headache, or sudden perception disorders is sufficient reason for immediate discontinuation. A mere change of the drug is probably hazardous. This case shows that CVT can masquerade for many months as a simple headache or even a benign intracranial hypertension.

Extensions to the deep cerebral veins are associated with a sudden neurological deterioration and poor outcome. Imaging is critical to the diagnosis of this disorder, which can be made by non-invasive modalities such as CT or MR.

Non-enhanced CT scans may show hyperdense thrombus in the dural sinus (dense sinus sign), deep cerebral veins, or cortical veins (cord sign), cortical and subcortical haemorrhagic infarctions, and diffuse cerebral swelling. After contrast the thrombosed sinus remains unopacified with enhancement of the
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collaterals in the dura leaves (empty delta sign), the falx appears thickened, there is tentorial (shaggy tentorium) and gyral enhancement, and the transcortical medullary veins may enhance strongly. Deep cerebral vein thrombosis appears as a high density thrombus in the deep veins, vein of Galen, or straight sinus, with or without basal ganglia infarctions and petechial haemorrhages. The differential diagnosis of dural sinus thrombosis on CT scans includes normal neonates with unmyelinated brain and dense sinus, high-splitting tentorium and pseudodelta sign seen in subarachnoid haemorrhages.

MR findings vary with clot age. An acute thrombus is iso-intense with the cortex on T1-weighted images, older haematomas are hyper-intense on T1-weighted scans and hypo-intense on T2-weighted images, while sub-acute thrombi are typically hyper-intense on all pulse sequences. In the chronic phase, prominent collateral venous channels can be seen around and within the thrombosed sinus (i.e. recanalisation), and intense enhancement of the thrombus after gadolinium injection (i.e. conversion to vascularized connective tissue). Other causes of increased signal within a sinus on spin-echo images must be excluded, such as: turbulent or slow flow, flow entry phenomenon, even echo rephasing, and flow compensation techniques. Flow enhancing gradient-echo sequences along with spin-echo sequence should allow the differentiation. In acute thrombosis, high field strengths can give a low signal which can be confused with a patent dural sinus. MR Angiography or Venography provides conclusive evidence of flow in the sinus rather than relying on flow-related enhancement effects in standard imaging.

Angiographic signs of a thrombosed sinus include non-filling of the dural sinus, filling defects, enlarged medullary veins and other collaterals. Thrombosed cortical veins are seen as contrast collections which appear to hang in space with contrast persisting well into the very late venous phase. Deep cerebral vein thrombosis is seen as the non-filling of the vein of Galen and internal cerebral veins, with or without collaterals.

Cerebral CT Venography, possible with spiral CT and three-dimensional reconstructions, allows an accurate evaluation of the flow in the cerebral venous system. Venous transcranial Doppler ultrasonography can be used as a monitoring tool in the evaluation of the collateral venous flow in superior sagittal sinus thrombosis, however it needs further evaluation. Apart from anticoagulation and control of raised intracranial pressure, selective venous administrations of fibrinolytic agents by an interventional neuroradiologist is possible. It is conjectural as to whether or not the use of nimodipine contributed to the patient’s eventual excellent recovery.

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