

Multiple intracranial arteritis and hypothyroidism secondary to *Streptococcus anginosus* infection

A 50-year-old Chinese woman reported a sharp paroxysmal headache and abrupt paralysis of the left leg. She then developed ptosis, blurred vision, diplopia and fever. On admission, a neurological examination revealed right III, IV, VI and left V₁ cranial nerve palsy, bilateral upper eyelid oedema and left leg monoplegia (Medical Research Council grade 2/5). In addition, a left Babinski sign and nuchal rigidity were observed. Blood tests revealed elevated white cell count (WCC) and a majority of the cells were neutrophils. Lumbar puncture revealed that the WCC ($120 \times 10^6/\mu\text{L}$) and protein level (0.79 g/L) of the cerebrospinal fluid (CSF) were slightly elevated, though the intracranial pressure was normal. A cranial MRI showed an infarction in the right corona radiata and base of the skull structures were also involved. MR arteriography indicated that multiple intracranial large arteries were narrowed **Figure 1**. Moreover, the CSF culture indicated *Streptococcus anginosus* infection, which was diagnostically very important. Accordingly, the patient was treated with vancomycin, tinidazole, low-molecular-weight heparin calcium and

dexamethasone for 2 weeks. She achieved remission of the neurological symptoms but her heart rate gradually slowed (45–65 bpm) and blood pressure decreased (75–90/45–50 mm Hg). She became depressed and developed apathy towards food. The Mini-Mental State Examination score (23/30) mainly indicated memory deterioration, disorientation and partial acalculia. The results of timely blood pituitary function tests indicated considerably decreased free T₃, free T₄ and thyroid-stimulating hormone levels, which indicated primary hypothyroidism. After subsequent administration of 12.5 mg/day levothyroxine for 2 months, the patient's heart rate and blood pressure were normalised and mental status returned to normal.

At this point, a CSF bacterial culture established the specific pathogen to be *S. anginosus*, a member of the *Streptococcus milleri* group colonising the human oral cavity, pars pharyngeal pharynges. When a healthy individual's immunity declines, opportunistic infection with *S. anginosus* may occur.^{1 2} Apart from causing a toothache and headache, the infection may spread intracranially to cause meningitis and cerebral venous system thrombophlebitis.^{3 4} In addition, inflammatory involvement of multiple large arteries, that is, the internal carotid artery (ICA) and its branches, the anterior and middle cerebral arteries—is another rare specific feature secondary to *S. anginosus* infection. Because the cavernous segment of the ICA courses through the cavernous sinus, *S. anginosus* infection can also extend to enter the ICA and cause diffuse inflammation of the sinus, which in turn causes thrombosis and narrowing of the ICA and its branches. Inflammation may also spread to the carotid sheath through the parapharyngeal space.³ Monoplegia and hemiplegia

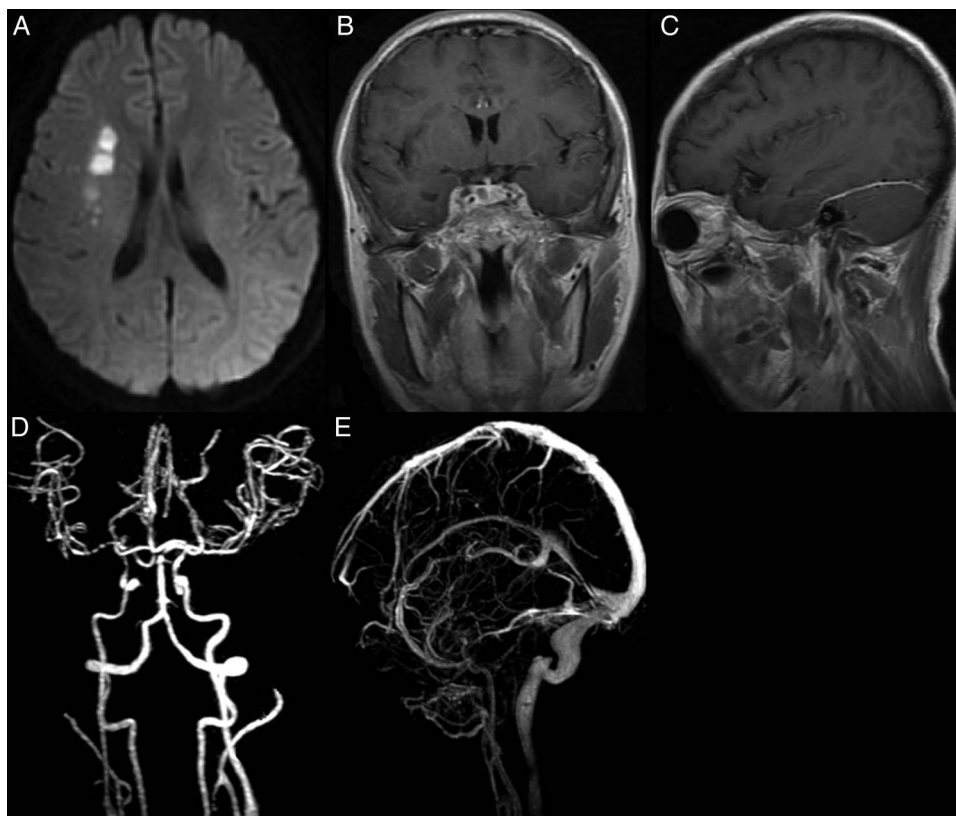


Figure 1 Brain parenchyma, meninges and cerebral vascular angiography studies. (A) Transverse view of diffusion-weighted imaging demonstrates infarction in the right corona radiata. (B and C) MR enhancement showed obvious involvement of the saddle area, pituitary stalk and tentorium of cerebellum. (D and E) MR angiography indicates narrowing of A2 segments of the bilateral anterior cerebral artery, a narrowed M2 segment in the right middle cerebral artery and cavernous segments in the right internal cervical artery; but the venous system was not obviously constricted except for thinness of the left transverse sinus, sigmoid sinus and internal jugular vein.

due to cerebral infarction are possible complications of arterial thrombosis. Considering the inflammatory mechanisms involved in the coagulant system, an early anticoagulation treatment of cerebral infarction is necessary.⁵ In the present case, the enhancement of the pituitary stalks on MRI and the presentation of primary hypothyroidism indicate partial pituitary insufficiency, which is also rarely reported. Theoretically, the cause might have involved the impairment of the hypothalamus–pituitary–thyroid axis through inflammatory damage to the pituitary stalk. The patient's favourable outcome confirmed the validity of the thyroxine replacement therapy.

We found that the initial CSF WCC that resulted from the *S. anginosus* intracranial infection, which is similar to those in viral infections, was considerably lower than that in typical acute bacterial meningitis, in which the count could reach thousands per microlitre in CSF. This may be a unique trait of the bacteria that needs to be investigated. Because the bacteria are facultative aerobes, early use of ceftriaxone, meropenem or vancomycin with tinidazole is necessary.

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Contributors All the authors were responsible for the study concept and design; acquired, analysed and interpreted the data; supervised and coordinated the study; and drafted/revised the manuscript for content.

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Competing interests None.

Patient consent Obtained.

Ethics approval The present study was approved by the ethics committee of Tianjin Medical University General Hospital.

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Image quiz

A 50-year-old woman was brought to medical attention because of fever, toothache, headache and impaired ambulation worsening over the previous week. On examination, she had the right III, IV, VI and left V1 cranial nerve palsy and left leg monoplegia. Head MRI, enhancement, MRA and MRV were done as in the figure below. Lumbar CSF culture revealed gram positive bacteria infection. After anti-biotic treatment, She achieved remission of the headache and fever. But gradually she developed apathy, memory deterioration, disorientation and partial acalculia. Which is the most valuable to find out the cause of the patient's mental status changes?

- A. Serum folate and Vitamin B₁₂ level
- B. Adrenal cortex function (cortisol and ACTH)
- C. Thyroid function (T₃, T₄ and TSH)
- D. A second lumbar puncture
- E. Head positron emission tomography (PET)

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Explanation

Answer C is correct. *Streptococcus* is one of the most common gram positive bacteria colonizing the oral cavity. In this case, CSF culture established opportunistic infection by *Streptococcus Anginosus*.^{1,2}

Apart from causing the toothache and headache, the bacteria may spread intracranially to cause meningitis (figure c) and cerebral venous system thrombophlebitis.^{3,4} But as shown in figure e, the venous system was not obviously constricted except for thinness of the left transverse sinus, sigmoid sinus, and internal jugular vein. Stroke (figure a) is a rare complication because of inflammatory

involvement of multiple large arteries, the internal carotid artery and its branches (figure d) through the parapharyngeal space.³ In the present case, the enhancement of the pituitary stalks on MRI were also seen (figure b). The impairment of the hypothalamus-pituitary-thyroid axis could explain the mental status changes. So the thyroid function should be immediately done. In fact, serum T3, T4 and TSH were considerably decreased in this patient. After two months of the thyroxine replacement therapy, the patient's mental status returned to normal. So C is the correct answer.

Answer A. Decreased serum folate and Vitamin B12 level are more likely seen in alcoholic abuse, malformation, anemia, *et al.*

Answer B. The hypothalamus-pituitary-adrenal axis might be involved in the case but it is difficult to explain the patient's mental status changes.

Answer D. After anti-biotic treatment, the patient's headache and fever remitted and lumbar puncture may reflect the effectiveness of the treatment.

Answer E. PET may find glucose metabolism disorders in the brain but difficult to find out the cause of the patient's mental status changes.

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