# World Journal of Clinical Cases

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#### **ABOUT COVER**

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CASE REPORT

## Epidemic Japanese B encephalitis combined with contactinassociated protein-like 2 antibody-positive autoimmune encephalitis: A case report

#### Pan Huang

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#### **Abstract**

#### **BACKGROUND**

It is not uncommon to develop viral encephalitis. Epidemic Japanese B encephalitis infection combined with contactin-associated protein-like 2 (CASPR-2) antibody-positive autoimmune encephalitis has not been reported at present. In clinical work, we need to consider more options.

#### CASE SUMMARY

A 32-year-old male worker presented with headache, fever and call-unresponsive presentation. Complete cranial magnetic resonance image showed symmetrical abnormal signals in bilateral medial temporal lobe, bilateral thalamus and basal ganglia. Improved lumbar puncture showed that cerebrospinal fluid protein and cell count increased significantly. Viral encephalitis was considered, and the patient's consciousness still increased rapidly after antiviral treatment. Further detection of Cerebrospinal fluid Japanese B encephalitis virus Polymerase Chain Reaction positive, serum autoimmune encephalitis antibody showed CASPR-2 antibody positive (1:320), the patient's condition gradually improved after plasma exchange treatment. 3 mo later, the serum CASPR-2 antibody was negative and the patient's condition was stable.

#### **CONCLUSION**

This article reports the world's first case of Epidemic Japanese B encephalitis infection combined with CASPR-2 antibody-positive autoimmune encephalitis, with a view to raising awareness.

Key Words: Epidemic Japanese B encephalitis; Contactin-associated protein-like 2 antibody; Autoimmune encephalitis

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**Core Tip:** For viral encephalitis in summer, there is no significant improvement after antiviral treatment, so the possibility of Japanese B encephalitis and autoimmune encephalitis should be considered.

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#### INTRODUCTION

Epidemic Japanese B encephalitis is a disease of the central nervous system caused by Japanese encephalitis virus. Due to the birth of a vaccine, this disease is relatively rare. Autoimmune encephalitis is a type of encephalitis caused by an immune response directed against central nervous system antigens mediated by autoimmune mechanisms. contactin-associated protein-like 2 (CASPR-2) antibody-positive autoimmune encephalitis is currently clinically rare and only a few cases have been reported. This article reports the world's first case of Epidemic Japanese B encephalitis infection combined with CASPR-2 antibody-positive autoimmune encephalitis, with a view to raising awareness.

#### CASE PRESENTATION

#### Chief complaints

A male patient, 32 years old, a worker, was admitted to the hospital due to headache, fever for 7 d and unconsciousness for 1 d.

#### History of present illness

7 d before admission, the patient developed headache and fever after cold (the patient was poorly conscious at the time of admission and could not describe the nature and location of the headache), with a maximum body temperature of 42 degree Celsius, accompanied by chills, no nausea and vomiting, abdominal pain, diarrhea, incontinence, and limb The convulsions were treated at the local hospital but no significant improvement was seen. The patient's condition worsened 1 d before admission, and began to grab clothes and sheets, and gradually developed to be unresponsive. The body temperature continued to exceed 39 degree Celsius. Considering the possibility of encephalitis, in order to further confirm the diagnosis "Infection" income department.

#### History of past illness

In the past, he suffered from asthma and improved after hormone therapy.

#### Personal and family history

The patient had no relevant personal or family history.

#### Physical examination

Body temperature 38.8 degree Celsius, pulse 84 times/min, breathing 22 times/min, blood pressure 137/50 mmHg. Sleepy, uncooperative physical examination, no autonomous speech, advanced intelligent activity examination cannot be performed. Double pupils and other large round shapes, about 3mm in diameter, light reflection is dull, conjunctival congestion and edema. No facial tongue paralysis, pharyngeal reflex exists. The muscle tension of the extremities increased, and there was avoidance during pain stimulation. The tendon reflexes of the extremities were normal (++), and the bilateral pathological signs were negative. Neck resistance was positive, 5 transverse fingers, and Glasgow Coma Scale was 8 points.

#### Laboratory examinations

At admission, C reactive protein was 52.68 mg/L, and the white blood cells were normal. Admitted to consider viral encephalitis, given acyclovir antiviral, dehydration and intracranial pressure treatment, the next day after admission, oxygen saturation decreased, consciousness deepened to coma, weak spontaneous breathing, bilateral pupils 4mm, light reflection disappeared, limb The cyanosis is cold and immediately transferred to intensive care unit (ICU) for tracheal intubation. A complete lumbar puncture examination during ICU treatment revealed cerebrospinal fluid protein 0.87 g/L, white blood cell counts 0.071 x 10°/L, glucose 4.31 mmol/L, chloride 117.9 mmol/L, negative test for herpes simplex virus DNA and fungus, and deliver Cerebrospinal fluid and plasma autoimmune encephalitis antibody

profile. During the period of waiting for the results of autoimmune encephalitis, after receiving communication with the patient, he was given methylprednisolone 1000 mg qd intravenous drip shock combined with gamma globulin 25 g qd treatment. On the 8th day of admission, the patient's level of consciousness was reduced to lethargy, spontaneous eye-opening, and body temperature gradually decreased too normal. On the same day, the patient's delivery of autoimmune encephalitis antibody showed positive plasma CASPR-2 antibody (1:320), The cerebrospinal fluid was negative for CASPR-2 antibody (Figure 1A and B). At this point, the patient was considered to have CASPR-2 antibody encephalitis, and then plasma exchange was performed (the plasma volume was calculated according to 40 m/kg). A total of 4 plasma exchanges were performed on the next 7 d. After the above treatment, the patient's condition gradually improved and the ventilator treatment was discontinued.

#### Imaging examinations

After the patient's condition was stable, a head magnetic resonance image (MRI) examination revealed mild brain atrophy and abnormal symmetry signals in the medial temporal lobe, bilateral thalamus, and basal ganglia (Figure 2A-D).

#### FINAL DIAGNOSIS

Epidemic Japanese B encephalitis combined with CASPR-2 antibody-positive autoimmune encephalitis.

#### TREATMENT

Combined with the patient's head MRI image results and the onset time is summer, it is highly suspected to be Epidemic Japanese B encephalitis, Cerebrospinal fluid is sent to the disease control center, and after 5 d, the report is Japanese encephalitis. At this point, the diagnosis of Epidemic Japanese B encephalitis combined with CASPR-2 antibody-positive autoimmune encephalitis was clear. After the above comprehensive treatment, the patient's condition improved 42 d after admission and was discharged. The patient's consciousness and speech were clear when discharged, but the senior Knowing functional decline, limb strength is 5 grades, bilateral pathological signs are negative, and meningeal irritation signs are negative.

#### OUTCOME AND FOLLOW-UP

After discharge from the hospital, he was followed up in the outpatient clinic every month, and there were no fluctuations in his condition. After 3 mo, the serum and cerebrospinal fluid autoimmune encephalitis antibody CASPR-2 antibody turned negative (Figure 1C and D).

#### DISCUSSION

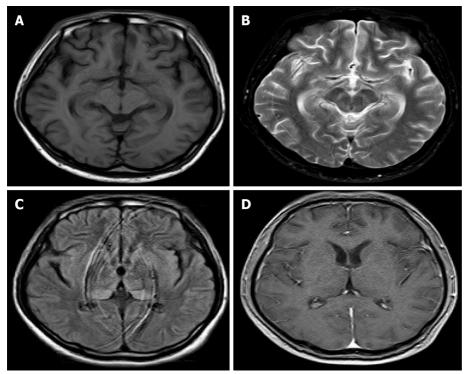
In recent years, autoimmune encephalitis has gradually attracted people's attention[1]. CASPR-2 is the main target antigen for the autoantibody of the voltage-gated potassium channel complex of neurons in and around the central nervous system The nervous system is expressed, so its clinical manifestations are central and peripheral nerve symptoms[2,3]. Peripheral nerve symptoms are due to neurological and muscle rigidity, muscle convulsions caused by overexcitation of peripheral nerves, such as Morvan syndrome and Isaacs syndrome[4,5], The central symptoms are often due to cognitive decline and seizures caused by the limbic system. The imaging findings of CASPR-2 antibody-positive autoimmune encephalitis are mainly manifested as abnormalities of the limbic system, accounting for about one-third of patients with positive findings on the head MRI. Compared with other types of encephalitis, the incidence of CASPR-2 antibody-positive autoimmune encephalitis is not high, about 3%, but the treatment effect is relatively significant[6,7]. The diagnostic criteria for CASPR-2 antibody-positive autoimmune encephalitis are as follows[8]: (1) Have anti-CASPR2 antibody positive; and (2) Rule out other possible causes such as: Morvan syndrome, primary central nervous system vasculitis, Rasmussen encephalitis, etc. In treatment, anti-CASPR2 antibody-positive autoimmunity is treated with high-dose corticosteroids (oral or intravenous), immunoglobulin, and/or plasma exchange as first-line treatment, while cyclophosphamide and rituximab are refractory cases's choice[9].

Epidemic Japanese B encephalitis is an acute zoonotic disease caused by Japanese encephalitis virus (JE virus). Epidemiology shows that there are currently 67900 new cases worldwide each year, of which about 50% occur in China, and the mortality rate is 25%-30%[10,11]. JE virus infection expands in peripheral tissues and enters the central nervous system, causing extensive central nervous system

Α	Antibody Project (CSF)	Result	Antibody Project (Serum)	Result
	Anti-AMPA1 receptor antibody	Negative	Anti-AMPA1 receptor antibody	Negative
	Anti-AMPA2 receptor antibody	Negative	Anti-AMPA2 receptor antibody	Negative
	Anti-CASPR-2 antibody	Negative	Anti-CASPR-2 antibody	Positive(1:320)
	Anti-GABAB receptor antibody	Negative	Anti-GABAB receptor antibody	Negative
	Anti-LGI1 antibody	Negative	Anti-LGI1 antibody	Negative
	Anti-NMDA receptor antibody	Negative	Anti-NMDA receptor antibody	Negative
	Anti-LgLON5 antibody	Negative	Anti-LgLON5 antibody	Negative
	Anti-DPPX antibody	Negative	Anti-DPPX antibody	Negative
	Anti-Gly1 receptor antibody	Negative	Anti-Gly1 receptor antibody	Negative
	Anti-DRD2 receptor antibody	Negative	Anti-DRD2 receptor antibody	Negative
	Anti-GAD65 antibody	Negative	Anti-GAD65 antibody	Negative
C	Antibody Project (CSF)	Result	Antibody Project (Serum)	Result
С	Antibody Project (CSF) Anti-AMPA1 receptor antibody	Result Negative	Antibody Project (Serum)  Anti-AMPA1 receptor antibody	Result Negative
С				111111111111111111111111111111111111111
С	Anti-AMPA1 receptor antibody	Negative	Anti-AMPA1 receptor antibody	Negative
С	Anti-AMPA1 receptor antibody Anti-AMPA2 receptor antibody	Negative Negative	Anti-AMPA1 receptor antibody Anti-AMPA2 receptor antibody	Negative Negative
С	Anti-AMPA1 receptor antibody Anti-AMPA2 receptor antibody Anti-CASPR-2 antibody	Negative Negative Negative	Anti-AMPA1 receptor antibody Anti-AMPA2 receptor antibody Anti-CASPR-2 antibody	Negative Negative Negative
С	Anti-AMPA1 receptor antibody Anti-AMPA2 receptor antibody Anti-CASPR-2 antibody Anti-GABAB receptor antibody	Negative Negative Negative Negative	Anti-AMPA1 receptor antibody Anti-AMPA2 receptor antibody Anti-CASPR-2 antibody Anti-GABAB receptor antibody	Negative Negative Negative Negative
С	Anti-AMPA1 receptor antibody Anti-AMPA2 receptor antibody Anti-CASPR-2 antibody Anti-GABAB receptor antibody Anti-LGI1 antibody	Negative Negative Negative Negative Negative	Anti-AMPA1 receptor antibody Anti-AMPA2 receptor antibody Anti-CASPR-2 antibody Anti-GABAB receptor antibody Anti-LGI1 antibody	Negative Negative Negative Negative Negative
С	Anti-AMPA1 receptor antibody Anti-AMPA2 receptor antibody Anti-CASPR-2 antibody Anti-GABAB receptor antibody Anti-LGI1 antibody Anti-NMDA receptor antibody	Negative Negative Negative Negative Negative Negative	Anti-AMPA1 receptor antibody Anti-AMPA2 receptor antibody Anti-CASPR-2 antibody Anti-GABAB receptor antibody Anti-LGI1 antibody Anti-NMDA receptor antibody	Negative Negative Negative Negative Negative Negative Negative
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Figure 1 Serum and cerebrospinal fluid antibodies to autoimmune encephalitis. A: Cerebrospinal fluid autoimmune encephalitis antibodies; B: Serum autoimmune encephalitis antibodies; C: Cerebrospinal fluid autoimmune encephalitis antibodies recheck after 3 mo; D: Serum autoimmune encephalitis antibodies recheck after 3 mo.



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Figure 2 Head magnetic resonance image. A: T1 image: No obvious abnormalities; B: T2 image: Bilateral medial temporal lobe high signal; C: T2 Flair image: High signal in both basal ganglia and thalamus; D: T1 enhanced image.

inflammation and destruction of the blood-brain barrier, thereby triggering central nervous system symptoms. Its typical clinical manifestations include recurrent seizures, paralysis and Knowledge barrier[12]. The imaging findings are mainly manifested as abnormal signals involving the thalamus and limbic system. The magnetic resonance T2-Flair and diffusion weighted imaging images are more likely to detect the lesion. In terms of treatment, vaccine immunoprevention is the main measure against Japanese encephalitis. Intravenous administration of methylprednisolone combined with human immunoglobulin can also improve the clinical symptoms and long-term recovery of some Japanese encephalitis patients. The diagnosis of Japanese B encephalitis in this case was based on the following:

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The patient's onset of illness was in the summer, the favourable season for Japanese B encephalitis, with the typical clinical presentation of high fever/impaired consciousness/convulsions and, most importantly, a polymerase chain reaction test of the patient's cerebrospinal fluid by the Centre for Disease Control that found positive for Japanese B encephalitis virus.

There have been reports of Herpes simplex encephalitis combined with autoimmune encephalitis, and Epidemic Japanese B encephalitis combined with autoimmune encephalitis has only been reported in a few cases, but Epidemic Japanese B encephalitis combined with CASPR-2 antibody-positive autoimmunity No related reports of encephalitis. From the diagnostic criteria, the patient not only met the diagnostic criteria for Epidemic Japanese B encephalitis, but also met the diagnostic criteria for CASPR-2 antibody-positive autoimmune encephalitis, However, the relationship between the two in this patient needs further study. By consulting the literature, we preliminarily speculated that this patient may be secondary to CASPR-2 antibody-positive autoimmune encephalitis after Japanese encephalitis infection: The first is a molecular simulation mechanism, that is, the JE virus protein that infects the central nervous system has the same or similar antigen structure cluster as CASPR-2[13]. The second is that after infection of the central nervous system JE virus leads to destruction of brain tissue, a large number of CASPR-2 antigen structure clusters are exposed, thereby inducing the body to produce autoantibodies against CASPR-2 and secondary autoimmune reactions[14].

#### CONCLUSION

In summary, this article reports a case of Epidemic Japanese B encephalitis combined with CASPR-2 antibody-positive autoimmune encephalitis.

#### **FOOTNOTES**

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