positive results. The results of a chest, abdomen, and pelvis CT scan and a serum tumour marker test ruled out a paraneoplastic origin. The patient was diagnosed with nonparaneoplastic anti-NMDA receptor encephalitis and treated with corticosteroids, immunoglobulins, and rituximab. Clinical progression was unstable; language impairment and neuropsychiatric manifestations persisted, motor symptoms (myoclonus and rigidity, with no clear epileptogenic focus) worsened, and the patient developed symptoms of dysautonomia. Neurological symptoms improved progressively with cyclophosphamide. A follow-up brain SPECT scan performed 5 months after the initial symptoms showed nearcomplete resolution of the alterations (Fig. 1B and D). Other successfully treated patients with this type of encephalitis have also shown disappearance or improvement of abnormal findings in brain perfusion SPECT<sup>3,4</sup> or <sup>18</sup>F-FDG-PET.<sup>7</sup> Brain perfusion SPECT played a crucial role in the diagnosis of this type of encephalitis given the atypical initial symptoms; language impairment of subacute onset suggested other diagnoses.

#### References

- Dalmau J, Lancaster E, Martínez-Hernández E, Rosenfeld MR, Balice-Gordon R. Clinical experience and laboratory investigations in patients with anti-NMDAR encephalitis. Lancet Neurol. 2011;10:63-74.
- Titulaer MJ, McCracken L, Gabilondo I, Armangué T, Glaser C, lizuka T, et al. Treatment and prognostic factors for long-term outcome in patients with anti-NMDA receptor encephalitis: an observational cohort study. Lancet Neurol. 2013;12:157–65.
- Llorens V, Gabilondo I, Gómez-Esteban JC, Agúndez M, Mendibe M, Bergara JC, et al. Abnormal multifocal cerebral blood flow

# Varicella-zoster virus meningitis in an immunocompetent paediatric patient<sup>\*,\*\*</sup>

### Meningitis causada por el virus varicela-zóster en un niño inmunocompetente

#### Dear Editor:

Herpes zoster is the localised manifestation of varicellazoster virus (VZV) infection. It occurs as a result of reactivation of the virus, which remains latent in sensory ganglia of cranial and spinal nerves once the infection resolves. Neurological complications of herpes zoster are on Tc-99m HMPAO SPECT in a patient with anti-NMDA-receptor encephalitis. J Neurol. 2010;257:1568–9.

- Yoshino A, Kimura Y, Miyazaki M, Ogawa T, Matsumoto A, Nomura S, et al. Limbic encephalitis with autoantibodies against the glutamate receptor epsilon 2 mimicking temporal lobe epilepsy. Psychiatry Clin Neurosci. 2007;61:335.
- Miyazaki M, Yoshino A, Teraishi T, Nomura S, Nemoto H, Takahashi Y. Encephalitis of unknown etiology with anti-GluR epsilon2 autoantibody, showing divergent neuroradiologic and clinical findings. Eur Neurol. 2007;57:111–3.
- Iizuka T, Sakai F, Ide T, Monzen T, Yoshii S, Iigaya M, et al. Anti-NMDA receptor encephalitis in Japan: long-term outcome without tumor removal. Neurology. 2008;70:504–11.
- 7. Probasco JC, Benavides DR, Ciarallo A, Sanin BW, Wabulya A, Bergey GK, et al. Electroencephalographic and fluorodeoxyglucose-positron emission tomography correlates in anti-N-methyl-D-aspartate receptor autoimmune encephalitis. Epilepsy Behav Case Rep. 2014;2:174–8.

J.P. Suárez<sup>a,\*</sup>, M.L. Domínguez<sup>a</sup>, M.A. Gómez<sup>a</sup>, J.C. Portilla<sup>b</sup>, M. Gómez<sup>b</sup>, I. Casado<sup>b</sup>

<sup>a</sup> Servicio de Medicina Nuclear, Hospital San Pedro de Alcántara, Cáceres, Spain
<sup>b</sup> Servicio do Neurología, Hospital San Pedro do Alcánte

<sup>b</sup> Servicio de Neurología, Hospital San Pedro de Alcántara, Cáceres, Spain

\* Corresponding author.

E-mail address: juanpablosuarez@yahoo.es

(J.P. Suárez). 2173-5808/

© 2016 Sociedad Española de Neurología. Published by Elsevier España, S.L.U. This is an open access article under the CC BY-NC-ND license (http://creativecommons.org/licenses/by-ncnd/4.0/).

relatively infrequent, especially in immunocompetent children.

We present the case of a 7-year-old boy who was admitted due to headache, vomiting, photophobia, and somnolence. The day before admission, the child's parents noticed that the patient had an erythematous exanthem on the left scapula, which was initially micropapular and subsequently vesicular; the exanthem was not pruriginous or painful. Two weeks previously, the patient had experienced upper respiratory tract infection. He had no recent history of foreign travel, contact with sick people, or insect bites. The patient had been vaccinated according to the Portuguese vaccination calendar; he had not received the varicella vaccine but had developed the condition at the age of 2 years. His parents were non-consanguineous.

During examination, the child was afebrile, haemodynamically stable, and drowsy (but easily awakened); he displayed no meningeal or focal neurological signs. Eye fundus examination revealed no abnormalities. The left scapular region showed a vesicular exanthem on an erythematous base, suggestive of herpes zoster. A complete blood count yielded 6000 leukocytes/mm<sup>3</sup>, 90.2% neutrophils, and normal C-reactive protein levels. A CSF analysis revealed 480 leukocytes/mm<sup>3</sup> (96.7% mononuclear cells), a protein level of 90 mg/dL, and low glucose levels (41 mg/dL in the



<sup>☆</sup> Please cite this article as: Oliveira K, Fonseca J, Moreira D, Vila Real M. Meningitis causada por el virus varicela-zóster en un niño inmunocompetente. Neurología. 2018;33:623–624.

<sup>\*\*</sup> This paper was presented in poster format at the 30th Annual Meeting of the European Society for Paediatric Infectious Diseases.

CSF and 85 mg/dL in the serum). Results from Gram staining and a CSF culture were negative for pathogenic bacteria. A polymerase chain reaction (PCR) study detected VZV in the CSF. No alterations were detected in cell-mediated and humoural immunity. The patient received IV aciclovir (10 mg/kg every 6 hours) for 14 days, progressing favourably. After 4 years of follow-up, our patient remains asymptomatic.

Two factors may increase the risk of herpes zoster in paediatric patients: immunosuppression<sup>1,2</sup> and varicella in early childhood.<sup>3-5</sup> Although the underlying mechanisms are yet to be understood, the inability of children below the age of 2 years to develop a specific cell-mediated and humoural immune response to VZV may explain why healthy paediatric patients with a primary VZV infection occurring in early childhood may develop herpes zoster and aseptic meningitis.<sup>6,7</sup> Herpes zoster lesions in children are less frequently associated with localised pain, paraesthesia, pruritis, or fever.<sup>8,9</sup> VZV is a neurotropic human herpesvirus<sup>8</sup> that may cause a wide range of neurological disorders if it reactivates.<sup>1-4</sup> Neurological involvement is usually accompanied by the exanthem that characterises herpes zoster,<sup>6,10</sup> except in immunosuppressed patients.<sup>1,3</sup> However, the literature also includes cases of meningitis secondary to VZV infection and no exanthem in immunocompetent patients, 3,5,11,12 since the reactivated virus can travel directly to the central nervous system without travelling to the epidermis.<sup>2,5</sup> CSF profiles of patients with meningitis secondary to VZV infection are clinically undistinguishable from those of patients with other types of viral meningitis.<sup>3,5,6,13</sup> Low CSF glucose levels have been observed in immunocompetent children and adults with meningitis due to VZV infection.<sup>3,12</sup> PCR analysis of the CSF for VZV DNA is recommended in cases of unexplained aseptic meningitis, especially in children who develop varicella in early childhood.<sup>4</sup> PCR analysis, especially in the first week after onset of acute symptoms of meningitis (with or without exanthem), is essential for diagnosis and treatment. Negative PCR results do not rule out meningitis due to VZV infection, however.<sup>12</sup> In these cases, anti-VZV IgM antibodies may be detected in the CSF. Presence of IgM antibodies in the CSF may point to CNS diseases since these antibodies do not easily cross the blood-brain barrier.<sup>1</sup> During the second week after disease onset, viral DNA disappears and anti-VZV IgG antibodies can be found in the CSF.<sup>12</sup>

IV aciclovir dosed at 10-15 mg/kg every 8 hours for 10-14 days is the treatment of choice for encephalitis due to VZV infection.<sup>1</sup> Most paediatric patients recover fully from viral CNS infections.<sup>8</sup> VZV should be considered as a possible cause of meningitis in children with vesicular exanthem and a history of varicella in early childhood.

## Funding

The authors have received no funding for this study.

#### References

- Tunkel A, Glaser C, Bloch K, Sejvar J, Marra C, Roos K, et al. The management of encephalitis: clinical practice guidelines by the Infectious Diseases Society of America. Clin Infect Dis. 2008;47:303–27.
- Gilden D, Kleinshmidt-DeMasters B, LaGuardia J, Mahalingam R, Cohrs R. Neurologic complications of the reactivation of varicella-zoster virus. N Engl J Med. 2000;342:635–45.
- **3.** Jhaveri R, Sankar R, Yazdani S, Cherry J. Varicella-zoster virus: an overlooked cause of aseptic meningitis. Pediatr Infect Dis J. 2003;22:96–7.
- Chiappini E, de Martino M, Jhaveri R. Varicella-zoster virus reactivation in the central nervous system. Pediatr Infect Dis J. 2004;23:185–6.
- Leahy T, Webb D, Hoey H, Butler K. Varicella zoster virus associated acute aseptic meningitis without exanthem in an immunocompetent 14-year-old boy. Pediatr Infect Dis J. 2008;27:362–3.
- Peña J, Pirics M, DiCaprio H, Julapalli M, Phelps B, Castagnini L, et al. Varicella reactivation presenting as shingles and aseptic meningitis in an immunocompetent 11-year-old boy. Clin Pediatr. 2009;48:435–7.
- Terada K, Kawano S, Yoshihiro K, Morita T. Varicella-zoster virus (VZV) reactivation is related to the low response of VZVspecific immunity after chickenpox in infancy. J Infect Dis. 1994;169:650–2.
- Myers M, Stanberry L, Seward J. Varicella-zoster virus. In: Behrman R, Kliegman R, Jenson H, editors. Nelson textbook of pediatrics. Philadelphia: Saunders Elsevier; 2007. p. 1057–61.
- Kangath RV, Lindeman TE, Brust K. Herpes zoster as a cause of viral meningitis in immunocompetent patients. BMJ Case Rep. 2013, http://dx.doi.org/10.1136/bcr-2012-007575.
- Han J-Y, Hanson D, Way S. Herpes zoster and meningitis due to reactivation of varicella vaccine virus in an immunocompetent child. Pediatr Infect Dis J. 2011;30:266–8.
- Big C, Reineck L, Aronoff D. Viral infections of the central nervous system: a case-based review. Clin Med Res. 2009;7: 142-6.
- Habib A, Gilden D, Schmid D, Safdieh J. Varicella zoster virus meningitis with hypoglycorrhachia in the absence of rash in an immunocompetent woman. J Neurovirol. 2009;15: 206–8.
- Cunha B, Warren-Favorito H, Mickail N. Unusually severe varicella zoster (VZV) virus viral (aseptic) meningitis in an unimmunized, immunocompetent host with chickenpox. Heart Lung. 2011;40:349–51.

K. Oliveira\*, J. Fonseca, D. Moreira, M. Vila Real

Departamento de Pediatría, Centro Hospitalar de Vila Nova de Gaia/Espinho, Vila Nova de Gaia, Oporto, Portugal

\* Corresponding author.

*E-mail address*: karina.valente.oliveira@gmail.com (K. Oliveira).

2173-5808/

© 2016 Sociedad Española de Neurología. Published by Elsevier España, S.L.U. This is an open access article under the CC BY-NC-ND license (http://creativecommons.org/licenses/by-nc-nd/4.0/).