

We describe a patient with enterovirus encephalitis who presented with oral ulcers and a rash (Figs 1-3). Although the rash appeared to be non-specific, enterovirus was immediately suspected as the aetiological agent based on its hand, foot, and mouth (HFMD), and body distribution. Enterovirus type 71 (EV71) was promptly identified by reverse transcription-polymerase chain reaction at multiple sites.

The previously healthy 12-month-old girl lives with her family in Shenzhen. She presented in August 2009 during the H1N1 influenza pandemic with a 2-day history of fever (maximum 38.4°C) and rigors, and a 3-day history of 'red spots' on the abdominal wall, which spread to the limbs and soles and became vesicular. The rash faded as the child developed oral ulcers 1 day preceding her admission. She also had some watery stool and vomited once. Her mother had had a rash on her hands 20 days previously, some diarrhoea but no fever. The family had travelled to

Henan in China the week before. The child received oral and intramuscular medications in Henan, and the family flew to Hong Kong the day before presentation.

At a district hospital in Hong Kong, the child was initially afebrile and haemodynamically stable; her Glasgow Coma Scale score was 15. She was treated for gastroenteritis with intravenous fluid. However, she spiked a fever of 39°C and developed generalised tonic-clonic convulsion with upward rolling eyes and tongue biting, which was treated with rectal diazepam and lorazepam. Urgent computed tomographic scan of the brain showed no abnormal cerebral findings. Intravenous antibiotics (ampicillin 50 mg, 6 hourly; cefotaxime 50 mg/kg/dose, 6 hourly) and acyclovir (25 mg/kg/dose, 6 hourly) were started. She had three more episodes of seizures until phenytoin (loading dose; 20 mg/kg) was added. She was transferred to a paediatric intensive care unit (PICU) for management. Cerebrospinal fluid showed white cells $5 \times 10^6/L$, red cells $820 \times 10^6/L$, glucose 3.5 mmol/L (reference range, 2.2-3.9 mmol/L), and protein 0.26 g/L (reference range, 0.15-0.45 g/L). Weakness of the right upper limb became apparent. During her stay in the PICU, there was no evidence of cardiac involvement. Intravenous immunoglobulin infusion (1 g/kg) was given. Computed tomographic scan of the brain showed subtle hypodensity involving the left posterior temporal region with blurring of the grey-white matter differentiation. Cerebral magnetic resonance imaging (MRI) showed abnormal



FIG 1. A tiny papule with erythematous rim on one finger



FIG 2. A tiny papule on the ventral surface of the first metatarsal of the right big toe



FIG 3. Oral mucosal ulcers

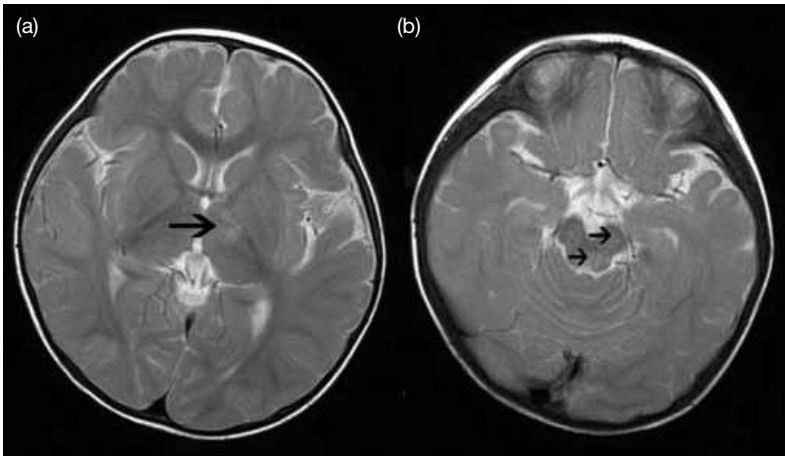


FIG 4. Axial T2-weighted magnetic resonance images showing hyperintensity (arrows) over (a) left thalamus and (b) brainstem, which demonstrate restricted diffusion on diffusion-weighted imaging (not shown) compatible with acute encephalitic changes

hyperintense T2/fluid-attenuated inversion recovery signals at the left thalamus, posterior limb of the internal capsule, brainstem, and cerebellar dentate nuclei (Fig 4). Cerebrospinal fluid, throat swab, and rectal swab subsequently confirmed EV71. The child recovered and was transferred to the general ward 3 days later, and discharged 16 days later. There was residual right upper limb weakness at discharge. When assessed 1 and 2 months after onset, there was no residual weakness and the MRI was normal.

Scattered vesicles with rim of erythema on the palms and soles, and vesicles or ulcers on the oral mucosa are typical of HFM disease.¹⁻⁹ In this patient, the rash involved the abdomen initially, spreading to the limbs, fingers, and feet. Vesicular rash can be confused with herpes and mycoplasma. Nevertheless, the typical HFM distribution may not always be present. The typical imaging finding may also aid diagnosis. Proper isolation and public health measures should be instituted in a timely fashion in enterovirus outbreaks.

Enterovirus type 71 was the major causative agent of HFM disease and encephalomyelitis. The clinical outcome is variable, with complete recovery being possible in some patients despite abnormal radiological findings.²⁻¹⁰ Brainstem and cervical spinal cord involvement are characteristic findings of enteroviral encephalomyelitis. The radiological findings and outcome in this patient are consistent with those reported in the literature.¹⁰

KL Hon, MD, FCCM
 Email: ehon@cuhk.edu.hk
 Department of Paediatrics
Winnie CW Chu, MD, FRCR
 Department of Imaging and Interventional Radiology
 The Chinese University of Hong Kong
 Prince of Wales Hospital
 Shatin
 Hong Kong

References

1. Zhu Z, Xu WB, Xu AQ, et al. Molecular epidemiological analysis of echovirus 19 isolated from an outbreak associated with hand, foot, and mouth disease (HFMD) in Shandong Province of China. *Biomed Environ Sci* 2007;20:321-8.
2. Chang LY, King CC, Hsu KH, et al. Risk factors of enterovirus 71 infection and associated hand, foot, and mouth disease/herpangina in children during an epidemic in Taiwan. *Pediatrics* 2002;109:e88.
3. Chong CY, Chan KP, Shah VA, et al. Hand, foot and mouth disease in Singapore: a comparison of fatal and non-fatal cases. *Acta Paediatr* 2003;92:1163-9.
4. McMinn P, Stratov I, Nagarajan L, Davis S. Neurological manifestations of enterovirus 71 infection in children during an outbreak of hand, foot, and mouth disease in Western Australia. *Clin Infect Dis* 2001;32:236-42.
5. Ooi MH, Wong SC, Mohan A, et al. Identification and validation of clinical predictors for the risk of neurological involvement in children with hand, foot, and mouth disease in Sarawak. *BMC Infect Dis* 2009;9:3.
6. Lee TC, Guo HR, Su HJ, Yang YC, Chang HL, Chen KT. Diseases caused by enterovirus 71 infection. *Pediatr Infect Dis J* 2009;28:904-10.
7. Tsao KC, Chang PY, Ning HC, et al. Use of molecular assay in diagnosis of hand, foot and mouth disease caused by enterovirus 71 or coxsackievirus A 16. *J Virol Methods* 2002;102:9-14.
8. Zhang X, Yan HP, Huang C, et al. The etiology and clinical manifestations of 70 patients with hand-foot-mouth disease [in Chinese]. *Zhonghua Yu Fang Yi Xue Za Zhi* 2009;43:872-4.
9. Yang Y, Wang H, Gong E, et al. Neuropathology in 2 cases of fatal enterovirus type 71 infection from a recent epidemic in the People's Republic of China: a histopathologic, immunohistochemical, and reverse transcription polymerase chain reaction study. *Hum Pathol* 2009;40:1288-95.
10. Shen WC, Chiu HH, Chow KC, Tsai CH. MR imaging findings of enteroviral encephalomyelitis: an outbreak in Taiwan. *AJNR Am J Neuroradiol* 1999;20:1889-95.