Leukocytoclastic Vasculitis in a Patient with HHV-6 Infection

Sir,

Cutaneous involvement in adults infected with human herpes virus 6 (HHV-6) is rare. Exanthematous reactions in immunosuppressed transplant patients and 1 case of reactive haemophagocytic syndrome have been reported (1, 2). Recently we described a case of erythema elevatum diutinum, a rare form of chronic cutaneous vasculitis, envisaging the aetiological role of HHV-6 (3). We describe here the case of an immunocompetent adult with leukocytoclastic vasculitis and propose that it is caused by HHV-6.

A 54-year-old woman presented with 1-month history of non-itchy papulo-purpuric lesions on her legs that had undergone several episodes of remission and exacerbation. The patient complained of fatigue, diffuse myalgia and arthralgia on her knees, ankles, pelvis and scapulae. Her medical history was unremarkable and she denied taking any drug before the eruption.

On examination she exhibited papular-purpuric lesions with sparse petechiae on the lower legs (Fig. 1). There were neither hepato-splenomegaly nor palpable lymph nodes. A biopsy specimen showed endothelial swelling with deposits of fibrin within and around the blood vessels and a cellular infiltrate consisting mainly of neutrophils with nuclear dust compatible with a diagnosis of leukocytoclastic vasculitis. Direct immunofluorescence revealed deposits of IgM and C3 in the blood vessels of the papillary dermis. Laboratory investigations showed normal values, except for the blood level of alanine aminotransferase 87 U/l (normal values 0-40 U/l) and aspartate aminotransferase 56 U/l (normal values 0-37 U/l). Chest X-rays and abdominal ecography were normal. Immunological investigations were normal and paraproteinemia and cryoglobulins were absent. Serology disclosed anti-HHV-6 IgG 1/160 and anti-HHV-6 IgM 1/160. Serology for the other viruses was negative or indicative of immunity.

The patient was given 30 mg/day deflazacort and heparin gel in occlusion.

The disease slowly improved and the lesions gradually faded over a period of 2 months, leaving pigmented macules. Aminotransferase levels became normal within 2 months.



Fig. 1. Papular and purpuric lesions on the legs.

Deflazacort was tapered and stopped after 45 days. One month later, the anti-HHV-6 IgG titre was 1/320 and anti-HHV-6 IgM 1/40. At present, the patient is in good health and free from lesions.

In early childhood, HHV-6 causes exanthem subitum (4). In adults, instead, the clinical spectrum of diseases associated with HHV-6 is still largely undefined (5). Like other herpes viruses, HHV-6 can produce latent infection and occasionally reactivation in immunocompromised hosts, causing bone marrow suppression, interstitial pneumonitis and encephalitis (1). In immunocompetent adults, its aetiological role has been suggested in several conditions, including mononucleosis-like syndrome, hepatitis and Kikuchi's lymphadenitis (5). Anti-HHV-6 IgM can be detected only in primary infections and reactivation states (6). On the contrary, anti-HHV-6 IgG are present in 80–90% of adults, but tend to disappear over time. In adults, therefore, their increase in titres may be considered indicative of reactivation (7).

Our patient had a rising titre of anti-HHV-6 IgG and a decreasing titre of anti-HHV-6 IgM. In the absence of active Epstein-Barr and cytomegalovirus infections causing serological cross-reactions, her cutaneous vasculitis (and possibly her liver involvement) may be ascribed to either a primary HHV-6 infection or an endogenous reactivation.

The rarity of HHV-6 primary infections in adulthood and its usual severe course both suggest that an endogenous reactivation is the most likely possibility. In conclusion, we recommend examining for HHV-6 infection in any patient presenting with otherwise unexplainable leukocytoclastic vasculitis.

REFERENCES

- 1. Singh N, Carrigan DR. Human herpesvirus-6 in transplantation: an emerging pathogen. Ann Intern Med 1996; 124: 1065–1071.
- Descamps V, Bouscarat F, Langlenne S, Arlangul E, Veber B, Deschamps D, et al. Human herpesvirus-6 infection associated with anticonvulsant hypersensivity syndrome and reactive haemophagocytic syndrome. Br J Dermatol 1997; 137: 605-608.
- 3. Drago F, Semino M, Rampini P, Lugani C, Rebora A. Erythema elevatum diutinum in a patient with human herpesvirus-6 infection. Acta Derm Venereol 1999; 79: 91–92.
- Yamanishi K, Okuno T, Shiraki K, et al. Identification of human herpesvirus-6 as a casual agent for exanthem subitum. Lancet 1999; i: 1065-1067.
- 5. Drago F, Rebora A. The new herpesviruses. Arch Dermatol 1999; 135: 71-75
- Fox JD, Ward P, Briggs M, Irving W, Stammers TG, Tedder RS. Production of IgM antibody to HHV-6 in reactivation and primary infection. Epidemiol Infect 1990; 104: 289 – 296.
- Levy JA, Ferro F, Greespan D, Lennette ET. Frequent isolation of HHV-6 from saliva and high seroprevalence of the virus in the population. Lancet 1990; i: 1047-1050.

Accepted August 10, 1999.

Francesco Drago, Paolo Rampini, Carlo Brusati and Alfredo Rebora Department of Endocrinology and Metabolic Sciences, Section of Dermatology, University of Genoa and Division of Dermatology, Gaslini Institute, Genoa, Italy.