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# Recurrent palmar blister

## Case study

A woman aged 34 years, who had no notable medical history, presented with a 2-day history of a painful and tender blister on her right hand but no associated symptoms. She had a history of recurrent episodes in the past 6 years that were preceded by localised pain, tenderness and burning, and resolved spontaneously. There had been no local trauma, recent exercise or use of medicines. Physical examination revealed a tense palmar blister with discrete perilesional erythema (Figure 1) and painful lymphangitis of the right upper limb (Figure 2). She was afebrile. Laboratory tests for inflammatory markers (C-reactive protein and erythrocyte sedimentation rate) showed no abnormal findings.

# **Question 1**

What is the most likely diagnosis?

# Question 2

What is the differential diagnosis?

## **Question 3**

What is the most appropriate management?

## **Question 4**

What is the most useful investigation to confirm the diagnosis?

## **Answer 1**

Given the absence of medication use, the most likely diagnosis is recurrent *Herpes simplex* infection. *H. simplex* viruses produce primary and recurrent vesicular eruptions that favour the

orolabial and genital regions. The palmar area is involved less commonly. Viral lymphangitis can complicate the clinical presentation, as in this case, and it is frequently misdiagnosed as a bacterial infection. 1.2 It should be noted that the infection can be transmitted to others through contact with skin vesicles.

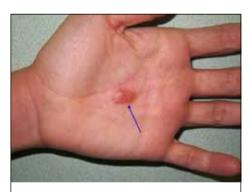


Figure 1. Palmar blister

## Answer 2

Recurrent *H. simplex* needs to be differentiated from other causes of bullous skin lesions:

- In a fixed drug eruption, one round sharply demarcated erythematous plaque is seen, sometimes with a central blister and violaceous hue. However, previous pharmacological exposure is a prerequisite and was not a feature of this case.
- Culicosis bullosa are bullous reactions to insect bites. These reactions are common and are often multiple, pruritic, excoriated papules or plaques. Although the cutaneous manifestations could be compatible with an insect bite, the recurrent course and painful lesions make this option improbable.

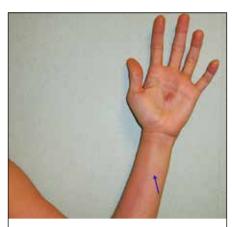


Figure 2. Lymphangitis

- Localised bullous pemphigoid could present
  as a tense blister with perilesional erythema,
  but lymphangitis is not a frequent finding.<sup>3</sup>
  Local factors, such as trauma, ultraviolet
  light or topical therapy, seem to be frequently
  implicated in the production of lesions.<sup>4</sup> The
  disease has a benign course and generally
  responds to cessation of exacerbating factors
  and initiation of topical steroids.
- H. zoster infection begins with a prodrome
  of pruritus, hyperaesthesia and intense pain.
   Most patients develop a painful eruption of
  grouped vesicles on an erythematous base in a
  dermatomal distribution. It is rarely recurrent.
- Bullous erysipelas represents a superficial cellulitis with lymphatic involvement. It is usually caused by group A-haemolytic streptococci. It typically appears as a sharply demarcated, tender area of erythema and oedema with an indurated border. The clinical course may be complicated by the local appearance of bullae. Accompanying signs and symptoms include lymphangitis and chills. The absence of fever and resolution with symptomatic treatment make this an unlikely diagnosis here.

#### Answer 3

Although valaciclovir, famciclovir and aciclovir have all shown high clinical efficacy in the treatment of herpesvirus infections, valaciclovir is preferable for long-term suppressive therapy from a treatment adherence perspective as it requires only daily dosing (500 mg/day). Valaciclovir is an oral pro-drug that is converted to aciclovir after administration. Chronic suppressive therapy is

usually reserved for patients with six or more recurrences per year. In our patient, the acute episode was not treated with valacyclovir but prophylactic treatment was instituted and no recurrence was reported at 1-year follow-up. It was not neccesary to add oral broad spectrum antibiotics in this case because the lymphangitis was due to viral infection.<sup>1,2</sup>

## **Answer 4**

Detection of herpesvirus DNA using real-time polymerase chain reaction (RT-PCR) is the best test to confirm the diagnosis. In addition to being the preferred method for identifying *H. simplex* in cerebrospinal fluid, RT-PCR is increasingly being used as a rapid, sensitive and specific method to detect *H. simplex* DNA in specimens from the skin and other organs. To ensure an adequate sample and avoid false-negative results, the vesicle should be unroofed and the base of the ulcer scraped with a flocked swab in universal transport media (Copan, Italy). H. simplex type 1 was isolated from this patient.

Tzanck smear test is rarely used now for diagnosis. It reveals multinucleated epithelial giant cells but it does not distinguish between *H. simplex* and *Varicella zoster* virus. <sup>6</sup> Viral culture was the traditional gold standard for detection of *H. simplex* and was the reference method against which all other tests were measured. A cell culture positive for *H. simplex* suggests probable active infection. However, a negative cell culture result does not rule out *H. simplex* infection, particularly if the specimen is from cerebrospinal fluid or nonvesicular lesions. <sup>8</sup> Stained biopsy specimens show intra-epidermal vesiculation associated with balloning degeneration of keratinocytes, which often fuse to form multinucleated giant cells. <sup>9</sup>

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#### References

- Cendras J, Sparsa A, Soria P, et al. Herpetic recurrent upper limb lymphangitis. Rev Med Interne 2008;29:158–60.
- Sands M, Brown R. Herpes simplex lymphangitis. Two cases and a review of the literature. Arch Intern Med 1988;148:2066–67.
- Lecluse AL, Bruijnzeel-Koomen CA. Herpes simplex virus infection mimicking bullous disease in an immunocompromised patient. Case Rep Dermatol 2010;2:99–102.
- Salomon RJ, Briggaman RA, Wernikoff SY, Kayne AL. Localized bullous pemphigoid. A mimic of acute contact dermatitis. Arch Dermatol 1987;123:389–92.
- Lebrun-Vignes B, Bouzamondo A, Dupuy A, Guillaume JC, Lechat P, Chosidow O. A meta-analysis to assess the efficacy of oral antiviral treatment to prevent genital herpes outbreaks. J Am Acad Dermatol 2007;57:238–46.
- Schmutzhard J, Merete Riedel H, Zweygberg Wirgart B, Grillner L. Detection of Herpes simplex virus type 1, Herpes simplex virus type 2 and Varicella zoster virus in skin lesions. Comparison of real-time PCR, nested PCR and virus isolation. J Clin Virol 2004;29:120–26.
- Tan TY, Zou H, Ong DC, et al. Development and clinical validation of a multiplex real-time PCR assay for Herpes simplex and Varicella zoster virus. Diagn Mol Pathol 2013;22:245–48.
- Singh A, Preiksaitis J, Ferenczy A, Romanowski B. The laboratory diagnosis of Herpes simplex virus infections. Can J Infect Dis Med Microbiol 2005;16:92–98.
- Fueki H, Sugita K, Sawada Y, Nakamura M, Tokura Y. Atypical large bullae caused by Herpes simplex in a patient with thymoma. Eur J Dermatol 2011;21:435– 36.

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