

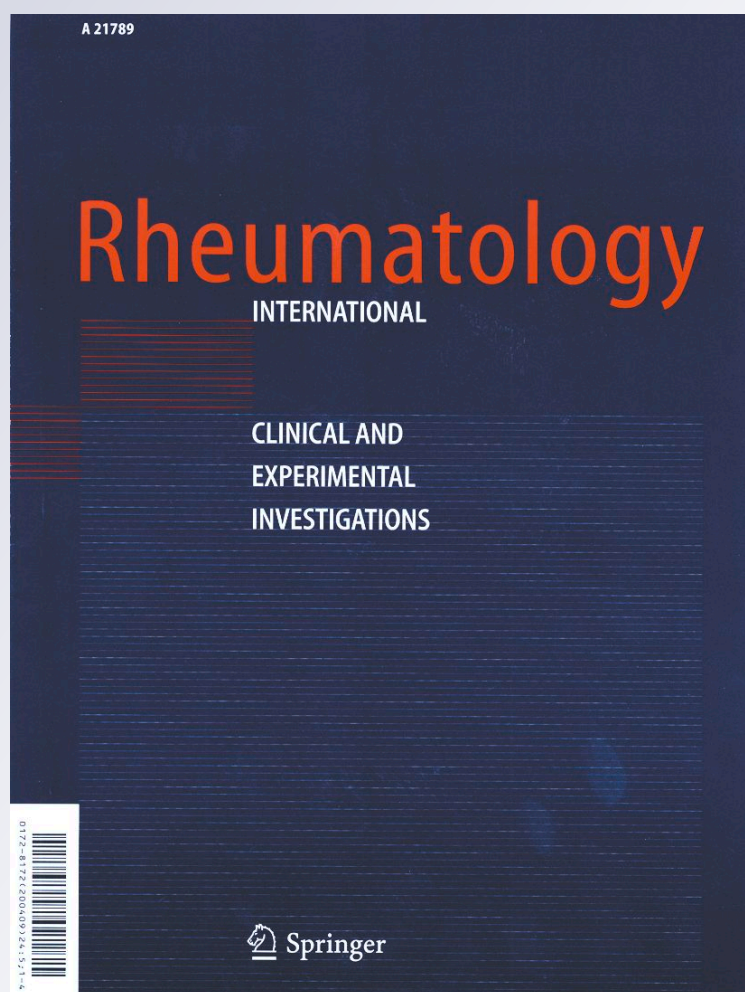
# *The impact of HSV for inflammatory arthropathy patients*

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**Rheumatology International**  
Clinical and Experimental Investigations

ISSN 0172-8172  
Volume 32  
Number 2

Rheumatol Int (2012) 32:489-490  
DOI 10.1007/s00296-010-1758-x



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## The impact of HSV for inflammatory arthropathy patients

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Received: 18 August 2010 / Accepted: 30 December 2010 / Published online: 18 January 2011  
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**Abstract** Herpes simplex virus type 1 (HSV-1), also known as herpes labialis, is the etiologic agent of vesicular lesions of the oral mucosa commonly referred to as “cold sores”. HSV-1 can also cause clinical disease in a wide variety of other anatomic locations including the genitalia, liver, lung, eye, and central nervous system. These infections can be severe, particularly in the setting of immunosuppression, such as inflammatory arthropathy patients on Methotrexate  $\pm$  biological therapies. Here, we highlight the importance of physician awareness of HSV due to its potential impact for rheumatology patients.

**Keywords** HSV · Inflammatory arthropathy

Dear Editor,

Herpes simplex virus type 1 (HSV-1), also known as herpes labialis, is the etiologic agent of vesicular lesions of the oral mucosa commonly referred to as “cold sores”. HSV-1 can also cause clinical disease in a wide variety of other anatomic locations including the genitalia, liver, lung, eye, and central nervous system. These infections can be severe, particularly in the setting of immunosuppression, such as inflammatory arthropathy patients on Methotrexate  $\pm$  biological therapies.

Once HSV infection has occurred, the virus lives in a latent state in nerve cell bodies in ganglion neurons and can reactivate. The frequency and severity of reactivation is determined by many factors, including immunodeficiency or stress [1]. In contrast to primary HSV-1, recurrent

HSV-1 is rarely associated with systemic signs or symptoms except for local lymphadenopathy. The majority of patients are aware of prodromal symptoms that herald the onset of a reactivation episode, such as pain, burning, tingling, and pruritus [2]. These symptoms may last from 6 to 53 h prior to the appearance of the first vesicles [2]. Recurrence patterns can demonstrate great variability from person to person. However, the specific triggers can be quite predictable for each individual patient and lesions tend to recur at the same site. Subclinical shedding is common in both immunocompetent and immunocompromised patients and is probably a key factor in transmitting virus to others.

The initial containment of HSV infection requires intact cellular immunity. Thus, immunocompromised hosts are at risk for increased frequency and severity of recurrent HSV infections. They are also at risk for dissemination of infection, which may include sites that are rarely involved in immunocompetent hosts, such as the lungs or gastrointestinal tract. Visceral dissemination is associated with high mortality. Risk factors for increased severity of HSV infection include history of HIV infection, malignancy, organ transplantation [3], malnutrition [4], pregnancy [5], and advanced age.

In immunocompromised patients, such as inflammatory arthropathy patients on Methotrexate  $\pm$  biological agents, HSV infection can lead to chronic mucocutaneous herpes simplex infection that may extend into deeper cutaneous layers, leading to easy friability and tissue necrosis. This can be associated with severe pain and atypical-appearing lesions on examination [6]. Mucocutaneous dissemination of HSV-1 should also raise suspicion of acyclovir resistance [7]. Also of note, Kaposi's varicelliform eruption characterized by severe oral–facial HSV can occur in immunocompromised patients, with vesicles and pustules often leading to extensive erosions that complicate recovery.

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Outside of oral complications, immunocompromised patients who have a HSV infection can develop gastrointestinal pathology (HSV esophagitis, which usually presents with odynophagia, dysphagia, or retrosternal chest pain), respiratory pathology (HSV pneumonitis), hepatic pathology (fulminant hepatitis), neurological pathology (HSV encephalitis, aseptic meningitis, autonomic dysfunction, transverse myelitis, benign recurrent lymphocytic meningitis, and Bell's palsy), and ocular pathology (conjunctivitis and/or blepharitis, chorioretinitis, acute retinal necrosis).

Due to the potential serious complications from HSV infection in immunocompromised patients, such as inflammatory arthropathy patients receiving Methotrexate  $\pm$  biological treatments, physicians need to be aware of the signs and symptoms of HSV infections and the need for quick/appropriate treatment.

## References

1. Freeman ML, Sheridan BS, Bonneau RH, Hendricks RL (2007) Psychological stress compromises CD8+ T cell control of latent herpes simplex Virus type 1 infections. *J Immunol* 179(1):322–328
2. Spruance SL, Overall JC Jr, Kern ER, Krueger GG, Pliam V, Miller W (1977) The natural history of recurrent herpes simplex labialis: implications for antiviral therapy. *N Engl J Med* 297(2): 69–75
3. Kusne S, Schwartz M, Breinig MK, Dummer JS, Lee RE, Selby R, Starzl TE, Simmons RL, Ho M (1991) Herpes simplex virus hepatitis after solid organ transplantation in adults. *J Infect Dis* 163(5):1001–1007
4. Becker W, Naude WD, Kipps A, McKenzie D (1963) Virus studies in disseminated herpes simplex infections: association with malnutrition in children. *S Afr Med J*, 37–74
5. Hillard P, Seeds J, Cefalo R (1982) Disseminated herpes simplex in pregnancy: two cases and a review. *Obstet Gynecol Surv* 37(7): 449–453
6. Westheim AI, Tenser RB, Marks JG Jr (1987) Acyclovir resistance in a patient with chronic mucocutaneous herpes simplex infection. *J Am Acad Dermatol* 17(5 Pt 2):875–880
7. Hardy WD (1992) Foscarnet treatment of acyclovir-resistant herpes simplex virus infection in patients with acquired immunodeficiency syndrome: preliminary results of a controlled, randomized, regimen-comparative trial. *Am J Med* 92(2A):30S–35S