Atypical Presentation of Syphilis in an HTLV-I Infected Patient

Dimas Carnaúba Jr., Achiléa Bittencourt and Carlos Brites

We report the case of a 44 year-old female, who presented a long-lasting, clinically atypical, secondary syphilis (“malignant syphilis”) in the right foot, which started six months before medical evaluation. The patient had a serological diagnosis of HTLV-I infection and syphilis two years before the onset of the skin lesions, following a blood donation. As she believed she was allergic to penicillin, she initially received sulfamethoxazole + trimethoprim, without any improvement of the clinical picture. After failure of this first treatment regimen, she was given penicillin, which promoted complete healing of the lesion. We found evidence that infection by HTLV-I is capable of modifying the clinical course of secondary syphilis.

Key Words: Secondary syphilis, HTLV-I infection; syphilis/HTLV-I coinfection; malignant syphilis.

Clinical manifestations of infection by human lymphotropic virus type I (HTLV-I) often include skin involvement, as seen in patients with infective dermatitis, severe forms of scabies, and adult T-cell leukemia/lymphoma [1-3]. It is also known that this agent can cause a variable degree of immune dysfunction, and this may modify the clinical presentation of infections [1,4-6].

Syphilis is a treponemal infection that, if not treated, can progress through four stages: primary (which usually is oligosymptomatic), secondary, and late, which includes central nervous system and cardiovascular involvement. Secondary syphilis is generally latent, but can present as a generalized and symmetrically distributed maculopapular rash, which typically involves palm and sole surfaces. [7].

Although there is experimental evidence showing that the clinical course of syphilis can be altered in animals infected by HTLV-I [8], we did not find any previous report of such interaction in human beings.

We report a case of a woman infected by HTLV-I, who developed an unusual presentation of secondary syphilis.

Case Report

A 44 year-old female teacher who lives in Maceió, the capital of Alagoas (a northeastern Brazilian state), asked for medical assistance for skin lesions in the upper face of her right foot, characterized by confluent, erythematous papules and plaques which started about six months before medical evaluation. She initially used a topical cream containing steroids, without any improvement. She had a serological diagnosis of HTLV-I two years before, following a blood donation. In addition, she had a positive test for syphilis (VDRL=1:64), which was initially treated with sulfamethoxazole plus trimethoprim, since she reported allergy to penicillin. She had never received blood products, and had no previous history of sexually transmitted diseases.

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Address for correspondence: Dr. Carlos Brites. Rua João das Botas, SN, 6o. Andar. Laboratório de Retrovírus, Canela, Salvador, Bahia, Brazil. Zip code: 40.155-020. Telephone: 55-71-2354866/235-4901. Fax: 55-71-247-2756. E-mail: crbrites@ufba.br

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Six months before the onset of the skin lesion, the patient presented with a vesicular eruption in the trunk, diagnosed as herpesvirus infection, which was complicated by secondary bacterial infection. At that time, a CD$_{4/8}$ measurement showed 990 CD$_{4^+}$ cells/mm$^3$, and 656 CD$_{8^+}$ cells/mm$^3$.

Dermatological examination: The patient presented confluent and erythematous papules and plaques, with small ulcers, on the dorsum of the right foot (Figure 1A), involving an area of 10 X 12 cm. Papules displaying these same characteristics were seen at the edges of the major lesions. She was initially diagnosed as having larvae migrans infection, and was treated with topical Thiabendazole. The lesion did not change with this treatment, and she was submitted to a biopsy, which revealed a pattern suggestive of secondary syphilis. The patient received 2,400,000 IU Benzathine Penicillin, intramuscularly, with a progressive improvement of the lesion, and complete healing after two months (Figure 1B).

Pathological report - The epidermis presented with hyperkeratosis and mild atrophy. Throughout the dermis, surrounding blood vessels, there was a perivascular and periadnexal infiltration of lymphocytes and plasma cells, sometimes associated with granulomatous reactions (Figures 2A and 2B).

Discussion

Infection by HTLV-I causes neurological (HAM/TSP), hematological (ATLL), and other types of diseases (infectious dermatitis, uveitis, pneumonitis, etc.) [9]. However, over than 90% of patients infected by HTLV-I do not present any clinical symptoms. Nevertheless, some of these individuals show subclinical evidence of immune dysfunction, which can modify the course of some diseases, such as Strongyloides stercoralis infection, and others [4,10].

Syphilis is an ancient, well-known, sexually transmitted disease [7]. Although treatment of syphilis is simple and efficacious, in patients with severe forms of immunodeficiency (such as AIDS) it can be harder to control, and can cause more severe forms of disease, with frequent CNS involvement [11]. A severe form of secondary syphilis, called “malignant syphilis” can occur in HIV patients and in other immunodeficient conditions, presenting as papulo-vesicular, pustular and necrotizing lesions [12,13]. HTLV-I infection can cause some degree of immune dysfunction, and thus could modify the clinical course of syphilis.

The long lasting, localized and atypical eruption that we found in our patient is not a usual finding in secondary syphilis. However, the observed pattern of perivascular granulomatous reaction is typical of long-lasting secondary syphilis [14]. In an animal model, two out four rabbits infected with HTLV-I and Treponema pallidum presented with a protracted course of syphilis, when compared with those that were not infected with HTLV-I, which showed the usual course of infection, with rapid healing of skin lesions. One animal needed to be treated with penicillin after eight weeks of the challenge with T. pallidum, suggesting defective cell-mediated immunity [8].

The unusual manifestation of syphilis in our patient probably was a consequence of co-infection with HTLV-I. The only immunological evaluation she was submitted to showed a normal CD$_{4/8}$ cell count, and a negative PPD test. These findings do not preclude immune dysfunction, since CD$_{4^+}$ values are often normal, or even increased, in patients infected by HTLV-I, and a negative PPD test cannot be taken as conclusive evidence of an altered immune response.

In conclusion, we found evidence that infection by HTLV-I can modify the natural course of syphilis. The usual syphilis treatment regimen seems to work as well in co-infected patients.

References

**Figure 1A.** Confluent and erythematous papules and plaques on the dorsum of the right foot, some of them with shallow ulcers.

![Image of confluent and erythematous papules and plaques on the dorsum of the right foot](image1.jpg)

**Figure 1B.** Aspect of skin lesion in the right foot, after treatment.

![Image of skin lesion in the right foot, after treatment](image2.jpg)
Figure 2A. Granulomatous reaction with epithelioid cells. HE, A 200.

Figure 2B. A moderate perivascular and periannexal infiltrate of lymphocytes and plasma cells in the deep dermis. HE, A 160.


