TITLE:

Refractory cutaneous lichenoid sarcoidosis treated with tranilast.

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Title:

A case of refractory cutaneous lichenoid sarcoidosis successfully treated with tranilast

Keywords: lichenoid sarcoidosis, sarcoidosis, granuloma annulare, tranilast,

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To the Editor: We would like to report a case of cutaneous lichenoid sarcoidosis, which was refractory to the conventional therapy but treated successfully with tranilast.

A 76-year-old Japanese man was referred to our clinic complaining of asymptomatic papules and erythematous plaques on his extremities and trunk (Fig. 1a-c). His health was good in general and he didn't take any oral medication. Superficial lymph nodes were not enlarged, and there was no abnormality on urinalysis, peripheral blood count or blood chemistry including blood sugar, hemoglobinA1c, angiotensin converting enzyme, and calcium. No abnormal finding was observed on chest radiography, chest and abdominal CT scans, electrocardiogram, and ophthalmologic examinations. QuantiFERON-TB Gold, an indirect diagnostic test for latent tuberculosis (TB) and active TB infections, was negative, either. The histology showed non-caseating necrotizing granulomatous infiltrates of epithelioid cells and giant cells throughout the upper to middle dermis. No atypical cell or histiocytic palisade surrounding mucin and degenerated collagen was found (Fig. 1d). Stains of Periodic acid-Schiff, Grocott methanamine silver, and Ziel-Nielson were all negative.

Based upon the clinical and histological findings above, the patient was diagnosed as cutaneous lichenoid sarcoidosis. Treatment with topical very strong class steroid ointment (0.1% mometasone furoate) in accordance with psoralen-ultraviolet A

(PUVA) were started, however, the eruptions were refractory to these therapies for over one year and extended progressively for the last several months. Since the manifestations were limited to the skin, oral administration of tranilast 300 mg/day, instead of systemic steroid treatment, was initiated. In response to tranilast, the eruptions became smaller and subsided, leaving only pigmentation after the 5-month course of this therapy (Fig. 2a, b).

Initially, cutaneous sarcoidosis, generalized granuloma annulare (GA), or infectious granulomatous diseases was considered for differential diagnosis. Generalized GA is occasionally complicated with diabetes mellitus and often shows palisades of histiocytes surrounding prominent mucin histologically^{1,2}, however, these findings were not seen in our case. Infectious granulomas were also excluded by a series of examinations on infection. Therefore, we diagnosed this case as the lichenoid type of cutaneous sarcoidosis. Topical corticosteroid can be effective to cutaneous sarcoidosis, but large or disfiguring lesions may require systemic corticosteroid therapy. Since sarcoidosis can relapse after cessation of steroid treatment³, indications for systemic corticosteroids should be limited to the patients with vital organ involvements. In our case, the skin eruptions, refractory to topical steroids and PUVA therapy, were attenuated by the treatment with tranilast. Tranilast, N-3,4-dimethoxycinnamoyl

antheranilic acid, is an anti-allergic drug used for the treatment of atopic dermatitis or keloid since 1980s. The precise underlying mechanism is unknown, but it has been demonstrated that tranilast inhibited the release of chemical mediators and the multinucleated giant cell formation of monocytes *in vitro*.⁴ There has been no report of diffusely-scattered lichenoid sarcoidosis treated successfully with tranilast. Although limited to a single case, it is considered that tranilast can be a choice for cutaneous lichenoid sarcoidosis refractory to conventional regimen.

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Figure legends

Fig 1. Clinical manifestations and histology. (a) Erythematous plaques and maculopapules were seen over the whole body (explanatoy; the darky-red lesion (*) is hemangioma simplex). (b) Individual lesions were 3- to 5-mm in a diameter, pinkish, follicular papules on the abdomen. (c) The clusters of the papules looked like erythematous plaques on the upper extremity. (d) Histological examination showed a dermal granulomatous infiltrate of epithelioid cells without caseating necrosis. A few giant cells and a sparce infiltration of lymphocytes were noted. (hematoxylin and eosin staining; magnification 200×.)

Fig 2. Clinical findings after oral tranilast therapy. After the treatment with tranilast, the skin lesions on the abdomen (a) and on the upper extremity (b) were attenuated.

Fig. 1

(a)

(b)

Fig. 2