Cutaneous Pseudolymphoma Secondary to Exogenous Pigment in a Polychrome Tattoo

Mendez Diaz Y1, Garcia-Arpa M2, Parra Cubillos A2, De Lara Simón IM3, Garcia Rodriguez R1, Gómez Torrijos E1
1Allergology Section, Hospital General Universitario de Ciudad Real, Spain
2Dermatology Service, Hospital General Universitario de Ciudad Real, Spain
3Histopathology Service, Hospital General Universitario de Ciudad Real, Spain

Key words: Cutaneous pseudolymphoma. Tattoo. Lymphocytes. Contact dermatitis.


Cutaneous lymphoid hyperplasia, or cutaneous pseudolymphoma (CPL), is a benign lymphoproliferative disorder of reactive T or B lymphocytes that clinically and/or histologically mimics cutaneous lymphoma [1]. The most frequent complications related to tattoos include infections, allergic and irritant contact dermatitis, and granulomatous dermatitis, mainly of the foreign body and sarcoid types. Pseudolymphomatous reactions are rarer, although their frequency may be underestimated [2]. Red pigment is the most common cause of reactions and pseudolymphomas due to tattoos [3-5].

We report the case of a patient with CPL associated with pink pigment in specific areas of a tattoo.

Figure. Papulovesicular lesions affecting only some pink areas of the tattoo (arrows). B, Skin tissue with a lymphohistiocytic infiltrate in the superficial dermis associated with an exogenous black-pink pigment deposit (hematoxylin-eosin, ×20); the inset shows the pigment deposit in detail (hematoxylin-eosin, ×40). C, Lymphocytes of the inflammatory infiltrate labeled using the immunohistochemical technique: i, CD4; ii, CD7; iii, CD2; iv, CD8.
A 25-year-old woman reported the appearance of pruritic lesions on her right shoulder at the site where a multicolored tattoo had been placed 2 months earlier. The lesions had been present for 2 weeks before she came to our clinic. The patient had had other black tattoos for 10 years, with no local reaction. Examination revealed a tattoo with green, black, and blue ink, as well as several shades of pink, with grouped papulovesicular lesions affecting only some pink areas of the tattoo (Figure).

No enlarged lymph nodes were palpated in the area of the tattoo. Biopsy of a papule revealed acanthosis with focal parakeratosis in the epidermis and a dense inflammatory infiltrate in the superficial dermis comprising lymphocytes (some with mild atypia), histiocytes, and some eosinophils. A disperse deposit of exogenous dark-pink pigment was observed in the interstitium. The deposit was phagocytosed by histiocytes (Figure). The immunohistochemical study showed that most of the lymphocytes were of the T (CD3⁺) strain, with predominant CD4 expression and a small accompanying population of B lymphocytes (CD20⁺). The lymphocytes maintained expression of CD2 and CD7 (Figure). These findings led us to diagnose T-lymphocyte CPL secondary to exogenous tattoo pigment. The patient underwent patch testing with the European standard and metal series and the pink pigment of the tattoo, which was provided by the patient. All the patch tests were negative except for nickel, for which the patient reported contact eczema induced by jewelry. The lesions disappeared spontaneously after 2 months.

During the last decade, there have been reports of allergic contact dermatitis to tattoos related to p-phenylenediamine [6]. In Spain, this substance, is the most commonly involved allergen in reactions affecting hairdressers (19.5%), which is the occupation with the highest risk of reactions to p-phenylenediamine (26%) [7].

The exact incidence of tattoo reactions is unknown. Tattooing has become very popular in both children and adults [8]. The tattoos are easily identifiable in the biopsy, since the pigment, which is usually distributed in the superficial and middle dermis, is seen in the form of extracellular deposits between the collagen bands and inside the macrophages. In addition to infections and neoplasms, tattoos can trigger nonspecific inflammatory reactions that affect the area of the tattoo or only a part of it. There may be different histological patterns, the lichenoid pattern being the most frequent [1]. The first case of CPL was reported in 1903, and, since then, approximately 40 cases have been reported. As in other inflammatory reactions to tattoos, the cutaneous manifestations are diverse and nonspecific, appearing mainly as macules, papules, or nodules [2,4,5].

The lesions sometimes worsen with exposure to sunlight and excessive sweating. The latency period is variable (from 1 month to several years after, even as much as 30 years) [1-3]. The histological examination is characterized by a diffuse dermal infiltrate consisting mainly of macrophages and small lymphocytes (mainly T lymphocytes, although a predominance of B and mixed lymphocytes has also been reported). Eosinophils, plasma cells, histiocytes, and multinucleated giant cells may also be observed. The presence of macrophages phagocytizing the pigment is a key finding in the diagnosis, as is the polyclonal rearrangement of the lymphocytes [4]. While pathogenesis is unknown, antigenic stimulation by the exogenous pigment is thought to be responsible for sensitization [2,4] and the delayed hypersensitivity reaction, which induces polyclonal proliferation of lymphocytes [2]. Red pigment is the most frequently implicated, although the reaction has also been described with green and blue pigments [1].

Complications of tattoos seem to be less frequent owing to improved techniques, more hygienic conditions, and less frequent use of sensitizing substances, such as mercury sulfide (red pigments), cadmium (yellow pigments), and cobalt (blue pigments), which are being replaced by more modern organic dyes [1,5]. In the case we report, we were unable to identify the components of the pink ink, although we believe that one of its components could be the culprit. Kluger et al found several metallic salts in the composition of the red ink, including zinc, copper, and nickel. In the present case, patch test results to nickel were positive, but we do not know if nickel was responsible for the CPL. Some tattoo inks analyzed contained nickel, although its causal relationship in individuals sensitized to this metal is not clear. In a series of 90 cases with inflammatory reactions to tattoos that were patch tested with standard, textile, and dye series, as well as with tattoo inks, positive results were observed in 29% of cases with the standard series (mainly to nickel), while results were positive for only 7% of the textile dyes and 11% of inks provided by the patient, thus indicating the limited role of contact tests in the diagnosis of tattoo reactions [9].

In the diagnosis of CPL caused by tattoos, the clinical and pathological correlation is fundamental, and the architecture and composition of the infiltrate, the presence of pigment, the immunohistochemistry results, and the absence of clonality have to be taken into account to enable a differential diagnosis with lymphoma [4].

There is no standard treatment for CPL induced by tattoo ink [3]. Topical or intralesional application of corticosteroids shows variable results [2,3,5], with recurrences that can be explained by the persistence of the pigment in the dermis. Other lines of treatment include surgical removal and/or CO2 and YAG laser therapy [5]. Spontaneous remission has also been reported, as occurred in the present case. Progression towards malignancy is exceptional; to date, only 1 case has been reported, with a tattoo that contained mercury, probably as a result of chronic antigenic stimulation of B lymphocytes. Therefore, careful follow-up is recommended [1,10].

We report a case of CPL as rare complication of a tattoo.

Funding
The authors declare that no funding was received for the present study.

Conflicts of Interest
The authors declare that they have no conflicts of interest.

References
Delayed Hypersensitivity Reaction to Oral Dimethyl Fumarate

Antolin-Amerigo D 1, Sánchez-González MJ 1, Barbarroja-Escudero J 1, Ayuso-Peralta L 2, Bellón-Heredia T 3, Ortega-Berruezo MA1, Alvarez-Mon M1, Rodríguez-Rodríguez M1

1 Servicio de Enfermedades del Sistema Inmune-Alergia, Hospital Universitario Hospital Príncipe de Asturias, Alcalá de Henares, Spain
2 Servicio de Neurología, Hospital Universitario Hospital Príncipe de Asturias, Alcalá de Henares, Spain
3 Instituto de Investigación Hospital La Paz, IDIP AZ, Hospital Universitario La Paz, Madrid, Spain

J Investig Allergol Clin Immunol 2018; Vol. 28(3): 201-203
doi: 10.18176/jiaci.0239

Key words: Hypersensitivity. Drug allergy. Multiple sclerosis. Allergy. In vitro technique.


Dimethyl fumarate (DMF) is a molecule derived from fumaric acid that is administered orally for the treatment of relapsing-remitting multiple sclerosis (MS). The mechanism of action of DMF comprises suppression of allergen-induced T-cell proliferation and modulation of the cytokine balance by suppressing only IFN-γ and not IL-5 [1-2], thus restoring the TH1/TH2 balance. It is also involved in interference with the intracellular redox balance and activation of nuclear (erythroid-derived 2)-like 2 related factor–mediated antioxidative response pathways leading to additional cytoprotective effects, and it seems to exert a marked effect on mitochondria [1-3].

We present the case of a 46-year-old woman with relapsing-remitting MS who had received several treatment options. However, given that she was a frequent traveller, she switched to DMF, which is a convenient, practical, efficacious first-line treatment for MS. Several hours after intake of the first dose of 240 mg, she developed generalized confluent and intensely pruritic wheals without angioedema or other systemic symptoms. These were treated at the emergency department with intramuscular methylprednisolone (60 mg) and dexchlorpheniramine (5 mg). She denied having taken NSAIDs, infection, or other possible causes of urticaria and had never experienced urticaria during her lifetime. Following recently published indications [1], we performed a skin prick test (SPT) with DMF at 7 µg/mL and intradermal tests with DMF at 0.7 µg/mL and 0.07 µg/mL. The results were negative. We also carried out a lymphocyte transformation test (LTT) using several concentrations: 70 µg/mL, 35 µg/mL, 15 µg/mL, 7 µg/mL, 5 µg/mL, 3.5 µg/mL, 1 µg/mL, 0.5 µg/mL, 0.1 µg/mL, and 0.05 µg/mL. The results were positive.

Phytohemagglutinin A (PHA) as a control mitogen LTT is considered weakly positive or doubtful if the stimulation index (SI) is between 2 and 3 and definitely positive if the SI is >3.

9. Serup J, Hutton Carlsen K. Patch test study of 90 patients with tattoo reactions: negative outcome of allergy patch test to baseline batteries and culprit inks suggests allergen(s) are generated in the skin through haptenization. Contact Dermatitis. 2014;71:255-63.

Manuscript received December 4, 2017; accepted for publication February 12, 2018.

Elisa Gomez Torrijos
Hospital General Universitario de Ciudad Real
C/ Obispo Rafael Torija, s/n
13005 Ciudad Real, Spain
E-mail: egomezt.cr@gmail.com