



## LETTER - DERMATOPATHOLOGY

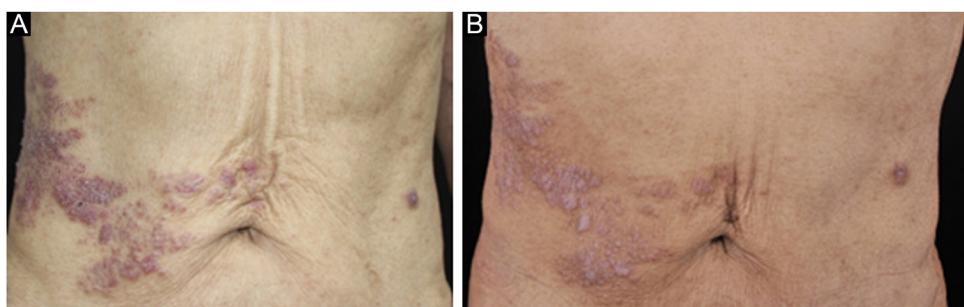
### An unusual Wolf's isotopic response: myelodysplastic syndrome with acute myeloid leukemia developed on herpes zoster lesions<sup>☆</sup>



Dear Editor,

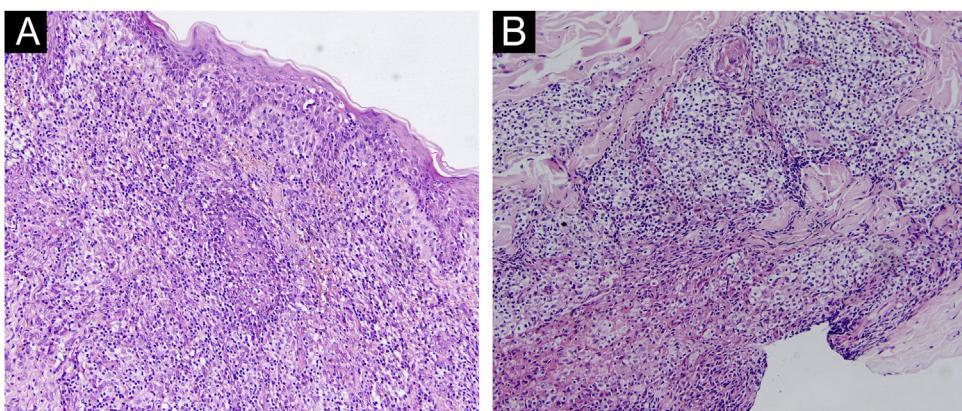
A 67-year-old male visited our clinic, complaining of severely itchy and tufted papules on his right waist and abdomen for more than 6 months. More than 6 months ago, cluster erythema blisters appeared on the right waist and abdomen which were diagnosed as "herpes zoster" clinically. Treated with traditional Chinese medicine for 10 days, erythema and blisters disappeared. Later on metameric papules, and plaques appeared on the healing herpes zoster with intense pruritus. His medical history was otherwise unremarkable. Physical examination: vital signs are stable; cardiopulmonary and abdominal examination was normal. Enlarged lymph nodes were palpable in the cervical and axillary regions, bilaterally. Metameric violaceous papules and plaques with infiltration were seen on the right waist and abdomen, corresponding to the right dermatomes of T8–T10 (Fig. 1 A). Histopathology (Fig. 2 A–B): lymphoid cells infiltrate in the epidermis and hair follicles, diffuse mild atypia lymphocytes, and epithelioid cells infiltrate in the whole dermis and subcutaneous fat layer. Immunohistochemistry: CD20, Myeloperoxidase (MPO), and CD117 were negative (Fig. 3 A–C). CD3, CD4, and CD8 were positive (CD4+ are more than twice CD8+ cells), and above 30% cells showed a positive staining with Ki67 (Fig. 4 A–D). Axillary lymph node puncture : lymph node inflammation. The patient refused lymph node biopsy. Cutaneous T-cell lymphoma was considered initially. However, he also had a persistent decrease in leukocyte count and moderate anemia. Further tests were performed to clear if hematological disorders could be associated. Electrophoresis of serum proteins revealed a rise in gamma globulin and no abnormal globulin M band. Bence Jones protein in urine was negative. Eventually, myelodysplastic syndrome with Acute Myeloid Leukemia (AML) was diagnosed by the results of bone marrow and gene tests (mutation of DNMT3A, IDH2, STAG2, and ASXL1). He was then transferred to the Department of Hematology to receive chemotherapy with azacitidine, daunorubicin and cytarabine. After one week of chemotherapy, the patient's pruritus was significantly relieved and the lesions were significantly improved. After four months of chemotherapy, his rash had improved. (Fig. 1 B). In addition, his blood changes and bone marrow images returned to normal, and the enlarged

lymph nodes disappeared. The patient is currently in complete remission.

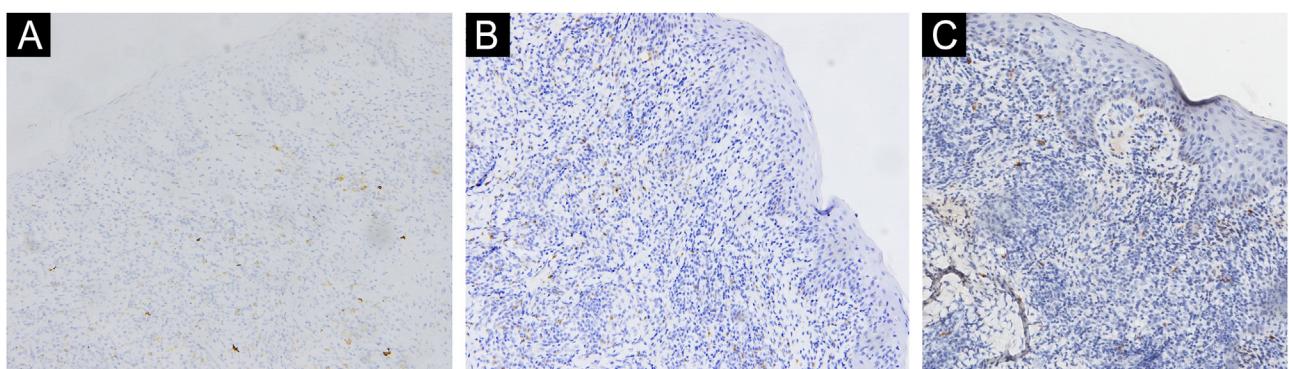


**Figure 1** (A) Metameric dark red papules, plaques with infiltration on the right waist and abdomen, corresponding to the right dermatomes of T8–T10. (B) After 4-months of chemotherapy: infiltration becomes flatter and lighter than before.

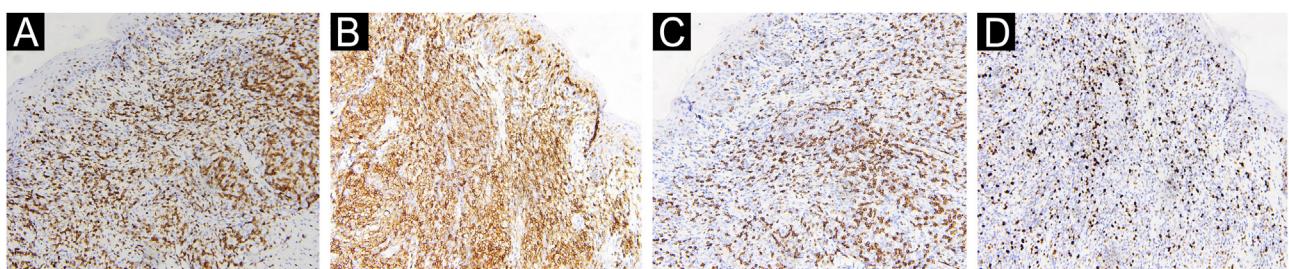
<sup>☆</sup> Study conducted at the Department of Dermatovenereology, Affiliated Hospital of Guangdong Medical University, Zhanjiang, Guang Dong, China.



**Figure 2** (A) Lymphoid cell infiltration in the epidermis and hair follicles, (B) Diffuse lymphocytes and epithelioid cell infiltration in the whole dermis. (Hematoxylin & eosin,  $\times 200$ ).



**Figure 3** Negative immunohistochemical staining for CD20 (A), Myeloperoxidase (MPO) (B) and CD117 (C) ( $\times 200$ ).



**Figure 4** Positive immunohistochemical staining for CD3 (A), CD4 (B), and CD8 (C), Ki 67 was positive in more than 30% of the cells (D) ( $\times 200$ ).

lymph nodes disappeared. The infiltration of tumor cells in the epidermis and the positive for markers of T-lymphocyte, especially the TH4 markers are dominant, with the absence of myeloid antigens observed in skin biopsies and immunohistochemistry respectively, led the misdiagnose of Mycosis Fungoides (MF) initially. Paradoxically, his rash is in accord with the plaque stage of MF, but the atypia lymphocytes infiltrate in the whole dermis and subcutaneous fat layer observed in the histopathology could be a tumor stage of MF. Moreover, the disease course is only 6 months, so the diagnosis of MF was ruled out. Given the positive effects of chemotherapy directed at AML, we consider that the metameric skin manifestation is cutaneous metastasis of myelodysplastic syndrome with acute myeloid leukemia in

healed herpes zoster, corresponding Wolf's Isotopic Response (WIR).

WIR describes the occurrence of a new skin disorder at the site of another unrelated and already healed skin disease. Herpes infection is the most common first disease, especially herpes zoster. The second diseases at the same site include granulomatous and lichenoid reactions, leukemia cutis, skin tumors, and infections.<sup>1</sup> Cutaneous metastasis of myeloid leukemia following herpes infection is pretty infrequent, compared with lymphoid leukemia, only one case has been reported until now.<sup>2</sup> The previous report is leukemia cutis developing as a WIR in a case of treatment-refractory AML, but the case we reported is WIR preceding the discovery of myelodysplastic syndrome

with acute myeloid leukemia. Moreover, myeloid leukemia cutis at the site of striae distensae<sup>3</sup> and sarcoid tissue reaction on herpes zoster scars in myelodysplastic syndrome patients<sup>4</sup> have also been reported. When new skin lesions appear at the site of healed herpes zoster accompanied by peripheral blood abnormality, a biopsy of the skin and bone marrow is required to rule out potential hematological tumors. The case we reported is an unusual WIR, with cutaneous metastasis of myelodysplastic syndrome with acute myeloid leukemia s. We are unable to find any similar previous report.

## Financial support

None declared.

## Authors' contributions

Yu Zhu: Contribution with the preparation and writing of the manuscript.

Wei Wu: Contribution to manuscript critical review, and approval of the final version of the manuscript.

## Conflicts of interest

None declared.

## Acknowledgment

We are very grateful to Professor Yeqiang Liu from Shanghai Dermatology Hospital for providing us with academic assistance and the immunohistochemistry of the patient.

## Eccrine syringofibroadenoma: report of two exuberant cases<sup>☆</sup>



Dear Editor,

Eccrine syringofibroadenoma (ESFA) is a rare benign adnexal tumor that arises from the excretory portion of the eccrine sweat glands.<sup>1</sup> It usually involves distal extremities in middle-aged to elderly patients,<sup>1</sup> presenting as solitary or multiple, coalescent, firm, skin-colored verrucous nodules, of variable sizes.<sup>1</sup> It is currently classified into five types: solitary lesions, multiple lesions associated with ectodermal dysplasia, lesions without additional cutaneous pathology, nevoid lesions and reactive lesions.<sup>2</sup> Clinical diagnosis is very difficult and, therefore, histopathological evaluation is essential.<sup>1</sup> Complete excision seems to be the definitive treatment.<sup>2</sup> The present report describes two cases of ESFA with an exuberant clinical presentation and diagnostic dif-

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Received 13 December 2022; accepted 10 February 2023

Available online 22 February 2024

<https://doi.org/10.1016/j.abd.2023.02.011>

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ficulty due to limited access to specialized services, albeit with excellent final results after excision by shaving.

## Case 1

A 76-year-old hypertensive diabetic male patient was referred to the dermatology service due to a vegetative lesion around a painful ulcer on the right lateral malleolus that had been noticed in 2018. He denied triggering factors and reported a previous biopsy at an external service diagnosed as verruca vulgaris. Associated with the condition, he had lymphedema of the ipsilateral lower limb. He underwent treatment for the ulcer but showed progression of the vegetative lesion. On examination, there was a hypertrophic scar on the lateral region of the dorsum of the right foot and multiple hardened verrucous pink nodules measuring 9 cm in their largest diameter, which coalesced on the lower edge of the scar (Fig. 1A). Dermoscopy showed no findings suggestive of malignancy (Fig. 1B). A new biopsy was performed, followed by complete removal of the lesion, by shaving, after histopathological evaluation. The image (Fig. 1C) shows the four-month postoperative period, without recurrence. Histopathology disclosed thin cords of cuboidal epithelial cells anastomosed in a network-like fashion in connection

☆ Study conducted at the Department of Dermatology, Hospital Universitário Pedro Ernesto, Universidade do Estado do Rio de Janeiro, Rio de Janeiro, RJ, Brazil.