INTRODUCTION

Kératoacanthoma (KA) is a common cutaneous neoplasm, probably derived from hair follicle cells. KA is a keratin-plugged, crater-shaped nodule that arises spontaneously, grows fast, and then typically regresses. It usually presents as a solitary lesion. However, multiple lesions in a sporadic form or an inherited manner are possible. The most common form of multiple KA is Ferguson-Smith type. We reported herein the first Tunisian case of Ferguson-Smith type keratoacanthoma that occurred in a 41-year-old man, with no family history of this variant.

CASE REPORT

A 41-year-old man, with no medical history, presented with multiple tumors on his forearms and trunk. Lesions appeared in the second decade of his life. Tumors grow slowly and resolve within months, leaving disfiguring scars. Physical examination revealed 23 erythematous, dome-shaped, cutaneous nodules of 0.5–3 cm diameter, with a central keratotic plug, covered with crusts and blood spots, and located on the trunk and limbs (Figure 1). Some of the lesions were clustered leaving Blaschko-linear pitted scars. The conjunctiva, palms, soles, genital, and oral mucosa were not affected. Dermoscopic examination showed a central structureless red-purple keratotic crater surrounded by a structureless white area with white circles surrounding hair follicles and linear, serpentine, and looped vessels (Figure 1). Histopathological examination of an excised tumor revealed a hyperkeratotic acanthotic epidermis with well-differentiated glassy keratinocytes and basal mitoses, surrounded by a polymorphonuclear inflammatory infiltrate (Figure 2). Based on clinical and histopathological findings, the diagnosis of multiple KA of Ferguson-Smith type was made. Treatment with acitretin showed a marked response. Recognizing this syndrome is crucial. Early treatment helps avoid scar formation.

DISCUSSION

Clinical forms of multiple KAs include Ferguson-Smith type, generalized eruptive Grzybowski type, centrifugum marginatum type, multiple persistent non-familial type,
agglomerate type, KAs in Muir-Torre syndrome, and KAs in xeroderma pigmentosum.4

Kératoacanthoma of Ferguson-Smith type, also known as multiple self-healing squamous epitheliomas (MSHSE), is the most common form of multiple KAs. It was first described in 1934 in Scottish families. However, sporadic cases have been reported in various countries.2,4

Haplotypes for polymorphic markers segregating with Ferguson-Smith syndrome in non-Scottish and Scottish families differ, suggesting that Ferguson-Smith syndrome is not caused by a founder mutation, thus considered now as a digenic/multilocus disease5,6 caused by loss of function mutations of Transforming growth factor, beta receptor I (TGFBRI) gene interacting with permissive variants at a second linked locus on the long arm of chromosome 9.7 Inheritance is autosomal dominant with incomplete penetrance. De novo mutations are possible which may explain the absence of a family history of KA in our patient.

FIGURE 1  Multiple self-healing lesions on the trunk (A) and limbs (B–G). Tumors are dome-shaped with a central keratotic plug, covered with crusts and blood spots (C). Some of the lesions were clustered leaving Blaschko-linear pitted scars (B). Dermoscopic examination showed a central structureless red-purple keratotic crater surrounded by a structureless white area with white circles surrounding hair follicles and linear, serpentine, and looped vessels (H).
Multiple self-healing lesions usually appear during childhood, adolescence, or early adulthood. Men and women are equally affected. Although lesions may arise in any part of the skin, KAs are mainly located on the face and extremities. The trunk is rarely affected. Palms and soles are usually spared. Each lesion starts as a reddish macule, becomes papular, and then grows rapidly into an ordinary solitary KA. The number of KAs varies from a few to hundreds. They evolve and then disappear rapidly within a few months, leaving atrophic and shallower scars and new KAs continue to emerge. The diagnosis of KA of Ferguson-Smith type can be made based on clinical aspect and distribution of the lesions, histopathological features, age at the onset of the disease, and self-healing course of the cutaneous tumors. It can be further confirmed by genetic tests (mutations in the gene TGFBR1).

An established association with internal malignancies is reported in cases of multiple KA associated with Muir-Torre syndrome and in other skin conditions such as xeroderma pigmentosum, but not in Ferguson-Smith syndrome. Only a recently published article on the occurrence of pancreatic cancer in Ferguson-Smith syndrome, yet the authors suggest that this association may be fortuitous.

Retinoids are the first-line treatment option for Ferguson-Smith syndrome. They are associated with a good clinical response. However, they have only a suspensive action. Therefore, a long-term regimen is necessary to sustain the clinical response. In our case, treatment with a relatively low dose of acitretin was efficient and well tolerated. Limited lesions can be treated with intralesional methotrexate. Cyclophosphamide was also used with good results in retinoid- and methotrexate-resistant cases of multiple KAs.

AUTHOR CONTRIBUTIONS
N. Litaiem conceived the idea for the document and contributed to the writing and editing of the manuscript. F. Khammouma contributed to the writing and editing of the manuscript. M. Mrad reviewed and edited the manuscript. The other authors discussed the results by revising critically for important intellectual content and have given final approval of the version to be published. All authors approved the final draft of the manuscript.

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CONFLICT OF INTEREST
The authors declare that they have no conflict of interest or sources of funding for this particular study.

DATA AVAILABILITY STATEMENT
Personal data of the patient were respected. No data are available for this submission.

ETHICAL APPROVAL
Personal data have been respected.

CONSENT
Written informed consent was obtained from the patient to publish this report in accordance with the journal’s patient consent policy.

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