LETTER



Psoriasis flare after influenza vaccination in Covid-19 era: A report of four cases from a single center

Dear Editor,

Psoriasis is a chronic inflammatory disease affecting 1% to 3% of the world's population^{1,2} and results from the interaction between genetics and environmental factors such as stress, infections, and drugs, causing a T-cell-mediated response.¹⁻³ Vaccination is an uncommon triggering factor for the flare-ups of several skin diseases,^{2,4-6} and a potential association between vaccination and the onset or exacerbation of psoriasis has been previously documented.^{2,5-7} In this letter, we report four new cases of psoriasis flare-ups after an influenza vaccination (Table 1).

The first patient was a 41-year-old man with chronic plaque psoriasis undergoing adalimumab therapy who developed a severe flare-up that required hospital admission 24 hours after an intramuscular Chiroflu influenza vaccination (Trivalent A/Victoria/2452/2019 H1N1, A/Hong Kong/2671/2019 H3N2, and B/Victoria/705/2018) (Figure 1). The patient clinically improved after treatment with subcutaneous guselkumab, topical corticosteroids, and emollients.

The second patient was a 70-year-old woman with chronic psoriasis who was undergoing treatment with topical corticosteroids and vitamin D analogs. The patient was referred to our department from the emergency room because she had started to develop diffuse erythema and numerous plaques following an intramuscular Chiroflu influenza vaccination 7 days earlier (Figure 2). The patient was started on treatment with oral acitretin, oral prednisone in slow de-escalation, and topical methylprednisolone aceponate, showing marked improvement after 3 weeks.

The third patient was a 55-year-old woman with severe chronic psoriasis treated with subcutaneous secukinumab (previously with

etanercept, adalimumab, and ustekinumab) who developed a facial psoriasis plaque 24 hours after a subcutaneous Chiroflu influenza vaccination. The patient was treated with topical fluticasone, with complete resolution of the skin lesion in 2 weeks.

The fourth patient was a 67-year-old woman with severe chronic psoriasis undergoing guselkumab therapy who developed a guttate psoriasis flare-up following a Chiroflu influenza vaccination 1 month earlier. The patient's biological therapy was changed to brodalumab, with improvement in the cutaneous lesions.

New-onset or severe exacerbations of psoriasis following influenza vaccination are uncommon. Most reported vaccination-related psoriasis flare-ups have been classified as guttate and guttate/plaque variants.^{2,4-7} We report four cases of psoriasis exacerbation following influenza vaccination with H1N1. H3N2. and B influenza strains. In our four patients, the close temporal relationship between the vaccination and the onset of the psoriasis flare-ups suggests a possible causal association.⁵ Although the etiological relationship between psoriasis and vaccination remains uncertain, it is known that the influenza vaccine generates T-helper (Th)1 and Th17 immunologic responses, which could represent a possible mechanism for vaccination-induced psoriasis.² The immunologic reaction to the influenza vaccination might rely on the generation of interleukin (IL)-6 and IL-22, producing Th17 cells that play a key role in the development of the characteristic epidermal changes of psoriasis. 5,6,8 In patients treated with IL-17 inhibitors, Th1 cells might be involved in the development of psoriasis flare-ups instead of Th17 cells. However, we found no differences in the clinical outcomes between the patient treated with secukinumab and the other patients. To date, "psoriasis

TABLE 1 Summary of psoriasis flares following influenza vaccination

Case	Age	Gender	Treatment before psoriasis flare	Vaccine type	Time from vaccination to psoriasis flare	Psoriasis type after vaccination	Treatment for flare
1	41	Male	Adalimumab	Chiroflu (anti-H1N1, H3N2 and B)	24 h	Plaque	Guselkumab Topical corticosteroids
2	70	Female	Topical corticosteroids Vitamin D analogs	Chiroflu (anti-H1N1, H3N2 and B)	7 d	Plaque	Oral acitretin Oral prednisone in slow de-escalation Topical corticosteroids
3	55	Female	Secukinumab	Chiroflu (anti-H1N1, H3N2 and B)	24 h	Plaque	Topical fluticasone
4	67	Female	Guselkumab	Chiroflu (anti-H1N1, H3N2 and B)	30 d	Guttate	Brodalumab



FIGURE 1 Chronic hyperkeratotic plaques on the legs with surrounding erythematous macules



FIGURE 2 Diffuse erythematous scaly plaques on the legs

vaccinalis" has also been described with Bacillus Calmette-Guerin, tetanus-diphtheria, and pneumococcal polysaccharide vaccines, including psoriasis-like eruptions and psoriatic arthropathy. ⁸⁻¹¹ The very low incidence of this condition and the favorable cost-effectiveness of the influenza vaccine should not change the immunization practice, especially for patients with psoriasis undergoing

immunosuppressive and/or biological therapy.^{5,6} Nevertheless, it is important to acknowledge vaccination as a triggering factor of psoriasis flare-ups, particularly in the COVID-19 era, given that we do not yet know whether COVID-19 vaccines will be a triggering factor for preexisting or new dermatological conditions, although it appears possible. Patients should therefore be carefully monitored once the COVID-19 immunization process begins.

CONFLICT OF INTEREST

The authors declare no potential conflict of interest.

AUTHOR CONTRIBUTIONS

All authors have contributed to the work. Marta Drake-Monfort, Susana Armesto, Marcos Antonio González-López, and Pablo Munguía-Calzada reported the four cases. Pablo Munguía-Calzada and Marcos Antonio González-López wrote the article. Leandra Reguero-del Cura and Ana Elisabet López-Sundh made the photographs, reviewed the bibliography, and performed the critical revision of the letter.

DATA AVAILABILITY STATEMENT

The data that support the findings of this study are available from the corresponding author.

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