

# Pityriasis Rosea and Pityriasis Rosea-like Eruption Following COVID-19 Vaccinations: Case Series from Oman

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## ABSTRACT

To mitigate the impact of the COVID-19 pandemic, several vaccines have been developed and administered to the public since 2021. A spectrum of cutaneous reactions has been reported among some of the vaccinated individuals. In this case series, we describe three cases of pityriasis rosea and pityriasis rosea-like eruption that manifested after COVID-19 vaccinations, which might suggest the vaccines as a possible trigger.

A spectrum of cutaneous reactions has been reported worldwide by recipients of COVID-19 vaccines. Regarding reactions to mRNA COVID-19 vaccines, a registry-based study of 414 cases with cutaneous reactions to vaccines showed that delayed large local reactions were most common, followed by local injection site reactions, urticarial eruptions, and morbilliform eruptions.<sup>1</sup> Less commonly reported were pernio/chilblains, cosmetic filler reactions, zoster, herpes simplex flares, and pityriasis rosea (PR)-like reactions.<sup>1</sup>

Rare cutaneous reactions to viral vector COVID-19 vaccines have also been reported.<sup>2,3</sup> In Brazil, where the viral vector-based Oxford-AstraZeneca vaccine was the most widely used vaccine, the most common cutaneous reactions were injection site reactions, acute urticaria, and morbilliform rash. More rarely reported were lichen planus, purpura/vasculitis, erythroderma, and fixed drug eruption.<sup>2</sup>

We describe three cases of PR and PR-like eruption (PR-LE) that emerged after COVID-19 vaccinations. Two of the patients received Pfizer-BioNTech, an mRNA vaccine, while the third was administered Oxford-AstraZeneca, a viral vector vaccine.

## CASE REPORTS

### Case One

A 19-year-old young man with no previously known health issues presented with a two-month history of pruritic, painless erythematous skin eruption over the trunk and the proximal extremities. The eruption was progressive and persistent. There was no associated systemic symptom or involvement of the scalp, nails, or mucus membranes. The patient also denied a history of recent dietary changes or new medication intake, but said that the symptoms began one week after receiving the first dose of Pfizer-BioNTech, an mRNA COVID-19 vaccine. Prior to the current presentation, he was treated with oral antihistamine and topical steroids, with no significant benefit.

Skin examination [Figure 1] showed erythematous, non-tender scaly papules and plaques that followed the skin tension lines over the lateral sides of the chest and coalesced in a reticular pattern over the lower back and abdomen. There was no petechial rash or wheals, and clinical examination revealed no evidence of a herald patch.

Skin biopsy microscopy [Figure 2] revealed spongiotic dermatitis with mild parakeratosis. There



**Figure 1:** (a) On presentation, erythematous papules and plaques were seen coalescing over the trunk. (b) Two weeks into the treatment, the lesions are clearing, albeit with persistent reticular post-inflammatory hyperpigmentation.

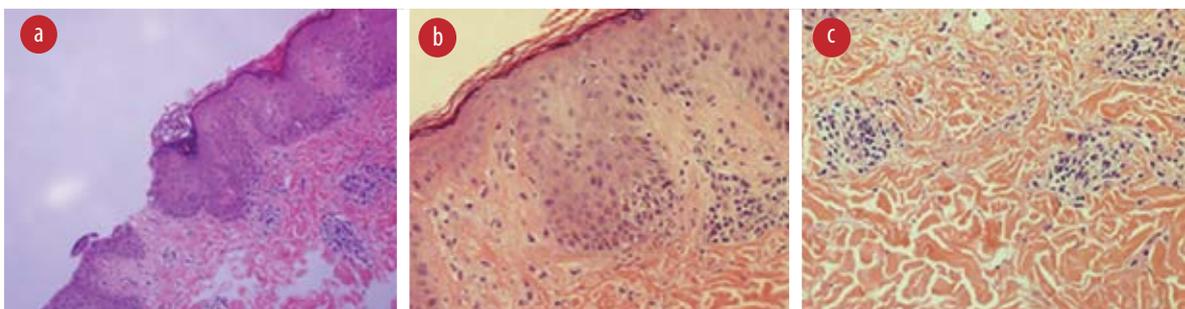
were few collections of inflammatory cells in the epidermis. The dermis showed perivascular and periadnexal inflammation, mostly lymphocytes. The features were suggestive of PR. Immunofluorescence was negative for the immunoglobulins IgA, IgM, IgG, and complement component C3. Serology for human herpesviruses 6 and 7 (HHV6/7) was not performed.

A course of oral azithromycin 500 mg once daily for three days and topical corticosteroids was prescribed. Progressive clearance of the eruption was noted after two weeks [Figure 1].

### Case Two

A woman in her thirties with no previously known health issues presented with one week history of an erythematous pruritic skin eruption on the trunk, arms, and axillae. There were no associated constitutional symptoms. There was no mucosal or palmoplantar involvement. The patient denied any new medication intake. The eruptions had begun two weeks after receiving the first dose of Pfizer-BioNTech mRNA COVID-19 vaccine.

Skin examination showed erythematous confluent targetoid lesions with collarette scales on



**Figure 2:** Hematoxylin and eosin stained skin biopsy showing (a) spongiosis and parakeratosis, magnification = 44 ×; (b) minimal inflammation in the epidermis, magnification = 10 ×; and (c) periadnexal and perivascular lymphocytic infiltrate, magnification = 10 ×.



**Figure 3:** (a) On presentation, erythematous confluent targetoid lesions with collarette scales are seen consistent with herald patch formation. (b) After three weeks of treatment with azithromycin and corticosteroid, the rashes have cleared.

both forearms, arms, and trunk. There was a large annular erythematous plaque with central clearing and collarette scaling located on the left axilla, consistent with the herald patch. No petechial rash or wheal was noted [Figure 3].

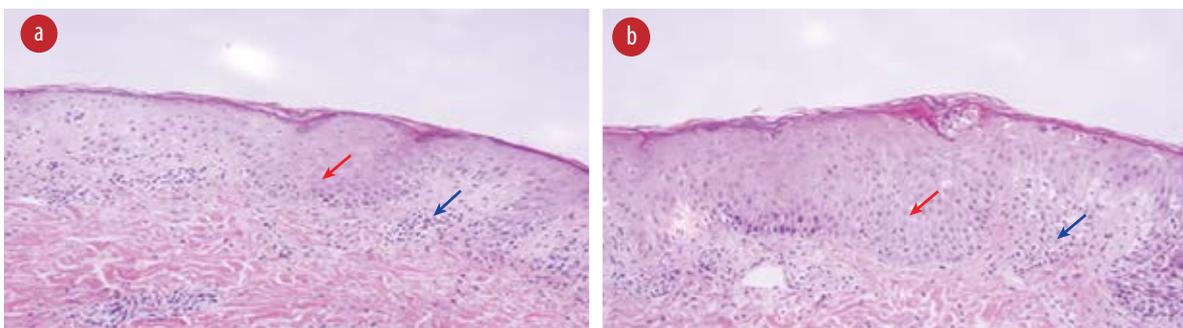
A course of oral azithromycin 500 mg for three days and topical corticosteroids were prescribed, and the eruption cleared in three weeks [Figure 4]. The patient subsequently took the second dose of the same vaccine, and a milder eruption occurred, which cleared spontaneously.

### Case Three

A man in his seventies with a history of hand dermatitis (though well-controlled by topical therapy) presented with two months history of

new-onset pruritic eruption. It started on the trunk and then spread to the extremities. There were no preceding viral illness symptoms or recent exposure to new medications. Three weeks before the eruption, he had received his second dose of Oxford-AstraZeneca COVID-19 vaccine (a viral vector vaccine that encodes SARS-CoV-2 spike protein). The patient reported that the first dose had been uneventful.

Skin examination revealed eczematous oval-shaped plaques following cleavage lines, especially on the back, forming a Christmas tree pattern [Figure 5]. Some of the plaques showed collarette scales. There were multiple annular plaques with a clearing center on the left shoulder, similar to a herald patch. Other eczematous plaques were scattered over the forehead,



**Figure 4:** Hematoxylin and eosin stained skin biopsy showing spongiosis (red arrows) with perivascular lymphocytic infiltrate (blue arrows), magnification = 4 ×.



**Figure 5:** (a) On presentation, erythematous oval plaques are visible on the trunk along the cleavage lines. (b) After two weeks of treatment with azithromycin and antihistamine, improvement is noticeable.

neck, and abdomen. No mucosal or palmoplantar involvement was noted.

Skin biopsy showed focal parakeratosis and spongiosis with perivascular lymphocytic infiltrate with occasional neutrophils and red blood cell extravasation in the dermis [Figure 4].

Complete blood count showed no evidence of eosinophilia. HHV6/7 testing was not performed. The patient was treated with azithromycin and antihistamine, and a good response was noted in the two-week follow-up visit [Figure 5].

## DISCUSSION

PR is a papulosquamous disease, typically preceded by a primary solitary herald patch followed within days to weeks by the emergence of smaller scaly lesions along the skin tension lines.<sup>4</sup> Possible triggers are bacterial or viral infections, medications, and vaccination. PR is mainly associated with endogenous systemic reactivation of HHV6/7 infections.<sup>5,6</sup>

A wide spectrum of cutaneous reactions was observed after COVID-19 vaccinations worldwide. The reactions included hypersensitivity reactions of type I and type IV, functional angiopathies,

autoimmune-mediated skin conditions, PR, and PR-LE.<sup>7</sup>

In a registry-based study conducted in the USA between December 2020 and February 2021, 414 skin reactions to two mRNA-based COVID-19 vaccines—Moderna (83%) and Pfizer (17%)—were identified. Delayed local reactions were the most common, followed by urticarial and morbilliform eruptions. Among the patients who had reactions to the first dose, 43% experienced recurrence after the second dose, as did our patient in Case Two, who had also received an mRNA vaccine. Less commonly reported reactions include manifestations of herpes zoster, herpes simplex, and PR-LE. The occurrence of PR-LE was mentioned in one case after the first dose of Moderna, two cases after the first dose of Pfizer, and one case after the second dose of Pfizer.<sup>1</sup> A middle-aged woman was reported to have developed PR 15 days after the second dose of the Oxford-AstraZeneca vaccine (viral vector-based), similar to our Case Three.<sup>8</sup>

A four-month-long nationwide study in Spain identified 405 instances of reaction after various types of COVID-19 vaccinations with the following break up: BNT162b2 (Pfizer-BioNTech; mRNA): 40.2%; mRNA-1273 (Moderna; mRNA): 36.3%,

and AZD1222 (AstraZeneca; viral vector): 23.5%. PR-LE constituted 4.9% of all cutaneous reactions.<sup>9</sup>

A systematic review of 31 studies by Khan et al,<sup>3</sup> included 111 patients (55.4% female) who developed PR or PR-LE after COVID-19 vaccination. Around 62% developed symptoms after the first dose and 38% after the second dose. Most had received the mRNA vaccines, Pfizer (35%) and Moderna (25%).<sup>3</sup> The reported mean time of symptoms onset was 8.6 days, and recovery was 6.4 weeks.

Rare cases of PR and PR-LE have been reported after vaccinations against smallpox, tuberculosis, influenza, papillomaviruses, poliomyelitis, tetanus, diphtheria, pneumococcus, diphtheria-pertussis-tetanus, hepatitis B, and yellow fever. In such cases, the average time lapse between vaccination and eruption onset was 5–17 days, and the exanthema lasted 2–6 weeks. Differentiating between PR and PR-LE is difficult and virological investigations for HHV-6/7 reactivation have been performed only in a minority of cases.<sup>10–12</sup> Cutaneous reactions post-COVID-19 vaccinations have been observed more in mRNA vaccines than in non-mRNA vaccines.<sup>13,14</sup>

Based on the criteria proposed by Drago et al,<sup>15,16</sup> PR-LE is better viewed as a drug/vaccine hypersensitivity reaction that morphologically resembles PR, than as classic PR. PR-LE usually presents with severe itching and atypical morphology of the skin lesions, such as papules, vesicles, urticated plaques, purpura, and target lesions (erythema multiform-like). They present as large-sized lesions or as confluent plaques. Unusual distribution patterns of the skin lesions are more common in PR-LE, like an inverse pattern, prominent involvement of the skin folds, face, or greater involvement of limbs than the trunk. In addition, mucosal involvement is more prominent with PR-LE than with classic PR. The lack of herald patch is more prevalent in PR-LE. Regarding histopathology, PR-LE presents with a picture of interface dermatitis with eosinophils, while PR usually presents with parakeratosis, spongiosis, and lymphocytic infiltrate. PR-LE usually resolves faster if the culprit drug is stopped compared to PR.<sup>16</sup>

The pathogenetic mechanisms of post-vaccination PR and PR-LE are still unknown. One hypothesis is that the occurrence of high cytokine response to the vaccine leading to immune dysregulation and reactivation of latent viral infections such as HHV-6/7.<sup>10</sup> PR-LE post-

vaccination can be considered more of a delayed hypersensitivity reaction that might be explained by molecular mimicry mechanism between the host proteins and viral vector or a vaccine ingredient, resulting in skin eruption mediated by T cells.<sup>5,7</sup>

Case One in this series resembled PR but had atypical morphology and prominent itch. Hence, we diagnosed it as PR-LE. Case Two was more representative of classic PR, given its typical morphology with collarette scales and obvious herald patch, although the involvement of proximal upper limbs was not usual. Case Three had an atypical morphology of PR with typical pattern, possible herald patch and supporting histopathological features so we diagnosed it as PR. All three cases responded well to azithromycin, which was administered for its anti-inflammatory effect.

As per Drago et al,<sup>15</sup> recurrences with the booster dose might not occur with PR. However, there was a recurrence in Case Two, which might point to more of a causal relationship. Our assumption is supported by a case from Turkey where a 45-year-old woman developed PR after receiving Sinovac-CoronaVac, a non-mRNA vaccine based on inactivated SARS-CoV-2. Her rash developed four days after receiving the first dose of the vaccine and was treated symptomatically with topical steroids and an oral antihistamine, and she had a recurrence after the second dose.<sup>17</sup> In addition, although unusual, multiple recurrences of PR have been documented in the literature.<sup>3,18,19</sup>

Large-scale epidemiological studies are called for to further elucidate whether there is a relationship between vaccination regimens and the reactivation of latent viruses, and to assess whether such reactivation could be a coincidence or a consequence of the SARS-CoV-2 vaccines.

## CONCLUSION

The three cases of PR and PR-LE reported here were possible cutaneous reactions to COVID-19 vaccination. Different types of COVID-19 vaccines—both mRNA and traditional vaccines—can produce such reactions. These reactions are rare, self-limiting, and only occasionally require treatment. We emphasize the necessity of vaccinating against COVID-19 while keeping close observation of any skin eruptions.

**Disclosure**

The authors declared no conflicts of interest. Written consent was obtained from all three patients.

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