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## Letter to the editor

### Bullous pemphigoid induced by the AstraZeneca COVID-19 vaccine



#### ARTICLE INFO

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Bullous pemphigoid (BP) is the most common autoimmune bullous disease, commonly encountered in the elderly and characterized by autoantibodies toward basement membrane zone antigens BP180 and BP230. Many trigger factors for BP have been identified such as ultraviolet (UV), radiation, drugs, trauma and burns. Post-vaccination bullous pemphigoid is rare. A few cases have been reported following diphtheria, tetanus, whooping cough, poliomyelitis, influenza, pneumococcal, meningococcal, hepatitis B, BCG and rabies vaccinations [1–15]. We report the first case of BP triggered by a COVID-19 vaccine.

A 77-year-old patient, with no significant pathological history, consulted for a diffuse itchy bullous eruption that appeared 24 hours after the first injection of AstraZeneca COVID-19 vaccine. The clinical examination found tense bullous lesions, with clear or hemorrhagic content, over erythematous skin, on the trunk, scalp and limbs (Fig. 1). No mucosal bullous lesion was noticed. Considering the age and the clinical characteristics, BP was suspected. Skin

biopsy showed a sub-epidermal cleavage and linear deposit of Ig G at the membrane zone using direct and indirect immunofluorescence assays, confirming the diagnosis. The occurrence of the eruption one day after the AstraZeneca COVID-19 vaccine was in favor of a post-vaccine BP that was reported to the pharmacovigilance department. The patient was treated by high potency topical steroids (propionate of clobetasol 0.05% cream) once a day and doxycycline (100 mg/day) with a favorable outcome. The vaccine booster was contraindicated.

BP is an autoimmune bullous disease, the main antigenic target of which is the trans membrane protein BP180 [16]. Various triggering factors have been reported in the literature, mostly radiotherapy, but also different drugs [17]. When a genetically predisposed individual is exposed to a trigger, it may lead to the revelation of the BP180 antigen, generating an immune response mediated by B-cells, leading to the production of auto-antibodies against BP180 antigen. These antibodies lead to the activation of the complement pathway causing protease release resulting in degradation of the basement membrane zone [18].

The skin side effects of vaccines are numerous. In addition to hypersensitivity reactions at the injection sites and drug eruptions, the onset or exacerbation of some skin diseases is possible. The occurrence of autoimmune bullous diseases after vaccination has already been reported in the literature. A few cases of pemphigus vulgaris have been reported after rabies and hepatitis B vaccines [19,20]. Post-vaccination BP is rare; only ten cases have been reported in the literature in adults and in children [1–15]. The period between the vaccine injection and the occurrence of BP varies between 1 day and 1 month [1–15]. The mechanism for induction of BP in response to vaccine is not very well understood. It has been proposed that BP can be triggered in some genetically predisposed individuals after attenuated life exposure vaccine that strengthens the immune response. The main mechanism by which specially inactivated vaccines provide immunity is the humoral pathway, that is, by stimulation of B-lymphocytes, leading to the production of antibodies. Post-vaccine BP has the same clinical, histological and immunological characteristics as other pemphigoids. Scalp involvement as in our observation is rare and could be a characteristic of post-vaccination BP. However, this has been never reported in published cases of post-vaccination BP. A causal relationship between vaccination and BP is difficult to establish, as vaccinations are extremely frequent. Thus, we cannot eliminate a coincidence between the occurrence of this bullous autoimmune skin disease in our patient and the AstraZeneca COVID-19 vaccine but a short delay of one day between the vaccination and BP may be in favor of a causal link.

Recurrence cases after another vaccine booster injection confirms the hypothesis of the role of vaccine and require more vigilance before proposing a vaccine booster to these patients [4]. However, in children it remains difficult to prohibit a vaccine booster, especially for certain serious diseases.



Fig. 1. Diffuse bullous eruption with hemorrhagic or tense blisters on the trunk.

## Disclosure of interest

The authors declare that they have no competing interest.

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