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# Acral papulovesicular eruption to smallpox vaccination

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#### Introduction

This is a case of a unique reaction to the ACAM2000° Small Pox vaccination previously reported in one case series. We find the morphology and locations of this reaction to be very interesting and appeared clinically consistent with a cutaneous hypersensitivity versus vasculitic reaction on patient presentation.

### Report of a case

A 27 year-old man with an unremarkable medical history received pre-deployment anthrax, MMR, Typhoid, Hepatitis B, and ACAM2000° Small Pox vaccinations on February 08, 2013. He reported an expected course with a vaccination site blister on his left upper arm with associated lymphadenopathy and malaise on post vaccination day (PVD) 3 which resolved by PVD 7. This was followed by the appearance of erythematous tender papules on his dorsal hands and fingers. By PVD14 multiple erythematous blanching papules measuring 2-5 mm in diameter were present on the dorsal hands, lateral fingers, extensor elbows, and knees (Figure 1). He had no atypical targetoid lesions and no oral or ocular involvement was present. The vaccination site on his left upper arm was well healed with no residual crust or drainage. There was no preceding viral respiratory infection, myalgia, pharyngitis, gastrointestinal upset or difficulty breathing and he had no reported contact with ill individuals following immunizations. Laboratory analysis showed a normal ESR, CRP, CBC, Chem 18, and ANA. Biopsy specimens were obtained from papular lesions on his right elbow and right dorsal hand.

## Diagnosis

 $A cral \, Papulove sicular \, Eruption \, Secondary \, to \, Small pox \, Vaccination$ 

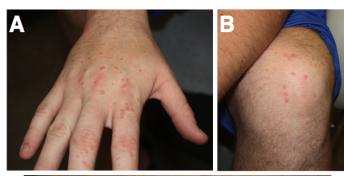
## Microscopic findings

Biopsy showed mild spongiosis with focal vacuolar degeneration along the dermal-epidermal junction. In the dermis there was papillary dermal edema and a moderately dense superficial to mid-dermal perivascular and perieccrine lymphocytic infiltrate (Figure 2). There was one focal area of epidermal necrosis and no leukocytoclastic vasculitis. PCR was not performed on the biopsy specimen.

#### Discussion

Small pox is a life threatening disease caused by an orthopoxvirus with a mortality rate reported as high as 30% [1]. In December 2002, the United States implemented a program of smallpox vaccinations for approximately 500,000 military personnel which came as part of a national program of preparedness against biological attack [2]. Because of the virus' long incubation period, the vaccine can be given up to 3 days after exposure and still prevent disease, however, preventative vaccination was determined to be the best way to avoid mass casualty

and loss of mission completion. Dryvax, a calf lymph derived vaccine which contains the NYCBOH vaccinia strain, was initially used in 2002 and yielded similar adverse event statistics as those reported in the 1968 national surveillance data, including incidence of an exanthematous reaction pattern of .054% [3]. In March 2008, ACAM2000° replaced Dryvax° as the vaccine used and stockpiled in the United States to protect against the threat of smallpox. ACAM 2000 contains the same vaccinia strain as Dryvax but is grown in African green monkey cells and the vehicle used was thought to be less reactogenic in comparison



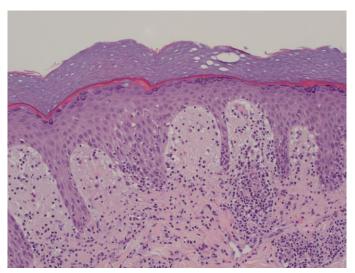


**Figure 1.** Multiple erythematous blanching papules measuring 2-5mm in diameter were present on the A) dorsal hands, lateral fingers, and B) knees. C) Note diascopy demonstrating blanching nature of papules.

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**Figure 2.** Mild spongiosis with focal vacuolar degeneration along dermal-epidermal junction. Noted papillary dermal edema and a moderately dense superficial to mid-dermal perivascular and perieccrine lymphocytic infiltrate.

to first and second generation vaccines [3,4]. Initial data suggests very similar rates, with approximately 11% of ACAM2000° smallpox vaccination recipients reporting cutaneous reactions in locations away from the vaccination site, compared to 10% with those vaccinated using Dryvax [5].

The induction of protective immunogenicity has been shown in animal trials to result from both humoral and cellular immune responses. Adverse reactions range in severity and typically involve skin; including urticaria, autoinoculation, erythema multiforme and nonspecific dermatidites. Potentially more serious adverse effects include but are not limited to eczema vaccinatum, generalized vaccinia, Stevens Johnson Syndrome, myocarditis and encephalitis [6]. This particular case appears to represent a unique response pattern that has been previously referred to as a benign, acral papulovesicular eruption. These specific symptoms thought to be associated with ACAM2000° administration have been reported in one previous case series [5]. Our current case, as well as the aforementioned case series, show striking clinical and histologic similarity but are confounded by the fact that multiple vaccinations were received along with ACAM2000°. Though a causal relationship between smallpox vaccination and the appearance of acral vesiculopustular eruptions cannot be definitively proven, the coadministered vaccines have not been reported to cause the cutaneous reaction pattern described above. Our patient did not require treatment and symptoms resolved by PVD 20.

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