

## Contact Dermatitis after Calcipotriol and Patch Test Evaluation

Sir,

The vitamin D3 analogue calcipotriol is more and more used topically as replacement of corticosteroids in the treatment of psoriasis. It is usually well tolerated and safe. The adverse events are mainly lesional/perilesional irritations, reported in about 15%, and facial irritations, reported in about 10% of treated patients, mostly mild and in only a few cases the reason for withdrawal of calcipotriol treatment (1–5). The nature of calcipotriol-induced dermatitis is usually considered to be of irritant nature, since it is a general experience that patients may tolerate the drug later if reinstated.

### CASE REPORT

In a study of treatment of psoriasis with UVB plus calcipotriol in an ointment formulation, 50 µg/g, compared with UVB plus the ointment base, one patient developed a severe dermatitis causing withdrawal of the treatment.

The patient was a 73-year-old man of good general health with the exception of rheumatic complaints, in particular regarding the back, since many years. He had had psoriasis of limited nummular to plaque type for 13 years. He had used calcipotriol ointment for 4 treatment periods of 4–5 weeks each during the last 2 years without any problems. No additional UVB was given during these periods of calcipotriol treatment. He had previously tolerated sunlight as well as UVB treatment.

At start of the actual treatment he had symmetrical psoriasis plaques on arms and legs. He accepted to participate in a double-blind study treatment of UVB plus calcipotriol ointment on one side and UVB plus ointment base on the other side. There was a slight but notable improvement after 2 weeks' treatment. Three weeks after start of treatment the patient noted some itching on one side just after application on the test ointment. A dermatitis involving the lesional as well as the perilesional skin gradually increased in severity, finally causing withdrawal of the topical as well as the UVB treatment. The trial code was broken, revealing that the ointment used on the dermatitis side contained calcipotriol.

An open test was instituted some weeks later on the inner part of the lower arm on skin not involved by psoriasis or adjacent to psoriasis lesions and previously not treated with calcipotriol. The calcipotriol ointment was applied twice daily within two indicated areas of the skin. After two applications there was an erythema, after four applications a clear dermatitis and after six applications a severe vesicular reaction. No additional UVB was given. The dermatitis at the test areas was treated with corticosteroid cream and healed within a week, but a dark pigmented colour remained several weeks after the test.

The patient was then treated with topical betamethasone valerate ointment and cream. He also exposed himself to the sun and suffered no harm.

### RESULTS

Three months after the treatment, an open test was performed on the inner part of the lower arm, with application of the same ointment, active and base, used at treatment as well as a patch test including the European standard allergy series used at the clinic, the marketed calcipotriol ointment and cream, the ointment base, and a dilution series of calcipotriol in isopropanol at the concentrations 0.4, 2, 10 and 50 µg/ml. All tests were negative. No additional UVB was given at test areas.

### DISCUSSION

The severe dermatitis induced by the calcipotriol ointment application both at treatment and at the first open test indicated that an allergic contact reaction had to be considered. The patient had used calcipotriol ointment during some treatment periods before, without any problems. He had also previously exposed himself to sunlight as well as UVB treatment without any sign of increased light sensitivity. The closed patch tests could not verify any allergic reaction against calcipotriol itself or the formulations used at treatment. Calcipotriol itself has irritating properties (6). The case presented illustrates that in single patients a severe non-allergic contact dermatitis might be provoked with simultaneously strongly positive open application test. The second open test indicated, however, that the lowered threshold was of short duration.

To what extent the UVB irradiation had contributed to the irritancy might be questioned. However, the open test during the active phase of the dermatitis, which turned out to be positive, was not combined with light exposure, and a photo-toxicity or photoallergy mechanism was, thus, not likely.

In two recent studies of the efficiency of calcipotriol combined with phototherapy, UVB, including 101 and 77 patients, respectively, no increase of lesional/perilesional or facial dermatitis was recorded as compared to calcipotriol alone (unpublished).

Altogether this case illustrates that endogenous variation in the threshold of irritation may render selected patients more vulnerable to irritative events in limited periods of time. To establish a diagnosis of allergic sensation, repetition of testing on some later occasion should therefore be considered mandatory (7, 8). The special risk of false-positive test reading to calcipotriol has been described in recent studies (6, 9). It is likely that some of the previously reported cases of allergy/possible allergy to calcipotriol were not valid, since repeated testing during a non-dermatitis phase was not always performed (10).

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*Accepted September 6, 1995.*

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