Does Addition of Topical Calcipotriol to UVB Increase the Risk of Irritant Reactions in Psoriasis?

Sir

The efficacy of both topical application of calcipotriol and UVB in the treatment of psoriasis is well documented. The combination of the two treatments seems to be more effective and to give a higher clearance rate (1, 2). In some cases irritant reactions related to the combined treatment have been reported (3).

The aim of this study was to evaluate irritant reactions related to calcipotriol ointment alone and UVB alone and to the combination of calcipotriol and UVB in the treatment of psoriasis.

MATERIAL AND METHODS

Data from two randomised right/left studies were analysed regarding irritant reactions. Patients included in the studies had chronic stable plaque-type psoriasis with symmetrical lesions on the arms, the legs and/or the trunk.

Series A: One hundred and one patients were treated with calcipotriol on one side and calcipotriol + UVB on the other side of the body (open study) (2).

Series B: Seventy-six patients were treated with calcipotriol+UVB on one side and vehicle+UVB on the other side of the body (double-blind study).

Calcipotriol ointment, 50 μ g/g, was applied twice daily and UVB was used 3 times weekly for 8 weeks. The UVB dose was increased from $0.7 \times MED$ prior to start of treatment in rapid steps up to the erythema threshold. Calcipotriol was applied at least 2 h prior to or after UVB exposure.

RESULTS

Irritation, with itching and erythema within the psoriasis lesions, was recorded in 17% of the sites treated with calcipotriol+UVB, in 14% of the sites treated with calcipotriol alone and in 13% of the sites treated with vehicle+UVB. The differences were not of statistical significance.

Seven patients in series B applied calcipotriol immediately after UVB; 3 of these 7 patients experienced UV burning, compared to 7 out of 170 patients in total applying calcipotriol at least 2 h prior to UVB.

The irritant skin reactions were mild to moderate and of the same kind and severity on the side treated with the combination of calcipotriol+UVB, with calcipotriol alone, or vehicle+UVB, except for one case treated with calcipotriol+UVB, who got a reaction with smarting pain and blisters starting 3 weeks after onset of treatment. Following reduction of UVB, treatment was continued in all except for the case with the reaction mentioned, in whom treatment was withdrawn.

There was no difference as regards skin type, PASI or thickness of the lesions prior to the treatment in patients with irritant reactions as compared to patients not irritated by the treatment.

DISCUSSION

Calcipotriol treatment may cause irritation in psoriatic skin, as calcipotriol itself has irritating properties (4, 5). In two

studies of calcipotriol ointment alone lesional/perilesional irritation was reported in 19% and 20%, respectively (6, 7). The main side-effect of UVB is irritation and burning. Standard volunteer photocontact allergy and phototoxicity studies conducted in the USA concluded that calcipotriol is unlikely to be associated with photoallergic or phototoxic reactions in clinical use (Data on file, Leo Pharmaceutical).

Four cases of burn reactions were reported recently in patients who were established on UVB phototherapy and subsequently began treatment with calcipotriol ointment (3). Two of the patients were able to continue the combination therapy after reduction of the UVB dosage.

In our two analysed treatment series presented, there was no difference in the rate of irritation related to the treatment of calcipotriol, of UVB or of the combination of calcipotriol and UVB. However, there was an increased risk of irritation when calcipotriol was applied just after UVB irradiation.

UVB enhanced the effect of calcipotriol in both the series. The efficacy of the treatment was greater on this side than on sides treated with calcipotriol alone (series A) or vehicle + UVB (series B). The skin on the calcipotriol + UVB-treated sides was thinner and less scaling than on the other sides, allowing more absorption of UVB. The UVB dosage used was relatively aggressive. One likely explanation for the somewhat more irritant events occurring on the calcipotriol + UVB-treated sides is that the UV-related irritant events were mainly caused by the erythemogenic UVB dose itself but enhanced when calcipotriol was applied immediately after exposure to the relatively high UVB doses.

CONCLUSION

Treatment with the combination of calcipotriol ointment and UVB for psoriasis generally does not cause irritation, lesional/perilesional erythema or itching more often than calcipotriol alone or UVB alone. The risk may be increased, however, when calcipotriol is applied immediately after exposure to relatively high doses of UVB. It is therefore recommended that calcipotriol ointment or cream is not applied immediately after UVB irradiation but preferably several hours later.

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REFERENCES

- Kragballe K. Combination of topical calcipotriol (MC903) and UVB radiation for psoriasis vulgaris. Dermatologica 1990; 181: 211–214.
- Molin L, Liberton H, van Weelen H, Austad J, McFadden N, Thune P, et al. Efficacy and safety of calcipotriol ointment in combination with UVB therapy. Third Congress of EADV, Copenhagen, September 26–30, 1993.
- McKenna KE, Stern RS. Photosensitivity associated with combined UV-B and calcipotriene therapy. Arch Dermatol 1995; 131: 1305–1307.

4. Serup J. Calcipotriol irritation: mechanism, diagnosis and clinical implication. Acta Derm Venereol (Stockh) 1994; Suppl 186: 42. 5. Molin L. Contact dermatitis after calcipotriol and patch test

evaluation. Acta Derm Venereol (Stockh) 1996: 76: 163-164. 6. Cunliffe WJ, Berth-Jones J, Claudy A, Fairiss G, Goldin D, Gratton D, et al. Comparative study of calcipotriol (MC903)

ointment and betamethasone 17-valerate ointment in patients with psoriasis vulgaris. J Am Acad Dermatol 1992; 26: 736-743. 7. Berth-Jones J, Chu AC, Dodd WAH, Ganpule M, Griffiths WAD,

Haydey RP, et al. A multicentre, parallel-group comparison of calcipotriol ointment and short-contact dithranol therapy in chronic plague psoriasis. Br J Dermatol 1992; 127: 266-271.

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