

- of two rapid latex agglutination tests for detection of cryptococcal capsular polysaccharide. *Am J Clin Pathol* 1998; 109: 634–641.
3. Gatti M, Di Silverio A, Cespa M, Mosca M. Primary unusual cutaneous cryptococcosis in an HIV former drug-abuser. *Mycoses* 1997; 40: 101–102.
 4. Hunger RE, Parades BE, Quattroppani C, Krähenbuhl S, Braathen LR. Primary cutaneous cryptococcosis in a patient with systemic immunosuppression after liver transplantation. *Dermatology* 2000; 200: 352–355.
 5. Yu RY. Cutaneous cryptococcosis. *Mycoses* 1996; 39: 207–210.
 6. Bohne T, Sander A, Pfister-Wartha A, Schöpf E. Primary cutaneous cryptococcosis following trauma of the right forearm. *Mycoses* 1996; 39: 457–459.
 7. Antti I, Jeskanen L, Rantanen T, Stubb S, Kariniemi AL. Cryptococcosis during systemic glucocorticosteroid treatment. *Dermatology* 1999; 199: 180–182.
 8. Patel P, Ramanathan J, Kayser M, Baran J. Primary cutaneous cryptococcosis of the nose in an immunocompetent woman. *J Am Acad Dermatol* 2000; 43: 344–345.
 9. Naka W, Masuda M, Konohana A, Shinoda T, Nishikawa T. Primary cutaneous cryptococcosis and *Cryptococcus neoformans* serotype D. *Clin Exp Dermatol* 1995; 20: 221–222.
 10. Hamman ID, Gillespie RJ, Ferguson JK. Primary cryptococcal cellulitis caused by *Cryptococcus neoformans* var. *gattii* in an immunocompetent host. *Aust J Dermatol* 1997; 38: 29–32.
 11. Antony SA, Antony SJ. Primary cutaneous cryptococcus in non-immunocompromised patient. *Cutis* 1995; 56: 96–98.
 12. Salm R, Groth D, Kappe R, Muller J. Primary cutaneous cryptococcosis after microtrauma of the index finger. *Mycoses* 1988; 1: 88–92.
 13. Bellosta M, Gaviglio MR, Mosconi M, Cavanna C, Viglio A, Rabbiosi G. Primary cutaneous cryptococcosis in an HIV-negative patient. *Eur J Dermatol* 1999; 9: 224–226.
 14. Boisnic S, Frances C, De Lassalle EM, Le Charpentier Y. Cutaneous papular cryptococcosis in AIDS. *Ann Pathol* 1989; 9: 71–72.
 15. Shuttleworth D, Philpot CM, Knight AG. Cutaneous cryptococcosis: treatment with oral fluconazole. *Br J Dermatol* 1989; 120: 683–687.
 16. Blanco P, Viallard JF, Beylot Barry M, Faure I, Mercie P, Vergier B, et al. Cutaneous cryptococcosis resembling molluscum contagiosum in a patient with non-Hodgkin's lymphoma. *Clin Infect Dis* 1999; 29: 683–684.
 17. Nicolas C, Truchetet F, Christian B, Dorvaux V, Cuny JF. Large crusted ulceration of the scalp: first manifestation of cryptococcosis in AIDS patient. *Ann Dermatol Venereol* 2000; 127: 188–190.
 18. Hall JC, Brewer JH, Crouch TT, Watson KR. Cryptococcal cellulitis with multiple sites of involvement. *J Am Acad Dermatol* 1987; 17: 329–332.
 19. Singh N, Rihs JD, Gayowski T, Yu VL. Cutaneous cryptococcosis mimicking bacterial cellulitis in a transplant recipient: case report and review in solid organ transplant recipient. *Clin Transplant* 1994; 8: 365–368.

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Crude Coal Tar Treatment Every Day Versus Every Other Day for Plaque Psoriasis

Sir,

Coal tar has been used in the topical treatment of psoriasis for more than a century and is assumed to have keratolytic, anti-pruritic, anti-mitotic and anti-inflammatory effects (1–3). Tar has, mostly due to its smell and staining properties, to some extent been replaced by other local treatments.

Coal tar is a complex mixture of thousands of compounds produced by condensation during the carbonization of coal (1, 4); however, the active substances in coal tar have never been identified (1, 4). It is well known that exposure to UV irradiation in the days following tar treatment can result in sunburn, indicating that the effect of tar in the skin persists for > 24 h.

In Denmark, crude coal tar (CCT) is still used daily for the treatment of plaque psoriasis. The procedure is very time-consuming and is limited by the number of nurses and bathrooms available. To evaluate the procedure and perhaps observe an extended effect of the tar in the skin, we investigated the effect of treatment with CCT every other day as compared with every day.

MATERIAL AND METHODS

The trial was conducted as a prospective, investigator-blinded, right/left randomized comparison of CCT treatment every day versus every other day. The study was approved by the Ethics Committee of Copenhagen.

A total of 15 adults (six males, nine females; mean age 54 years;

range 23–90 years) volunteered for the study and gave their informed consent. Patients were recruited among those referred to the Department. All patients suffered from chronic plaque psoriasis and none of them received any other treatment for their skin disease during the study. Exclusion criteria were allergy to coal tar and pregnancy. All patients were > 18 years old.

Treatment with CCT every weekday was randomly assigned to one side of the body by drawing lots. On the opposite side of the body CCT was applied every other weekday. The application of CCT was followed by 20 min in a bathtub (37°C). The patients used tar cream 5% during weekends.

The psoriasis was assessed by a second doctor who was unaware of the treatment. Psoriasis severity was assessed with respect to erythema, infiltration and desquamation by means of a modified Psoriasis Area and Severity Index (PASI) (5). The severity of the psoriatic lesions was recorded on a five-point scale (0 = absent, 1 = slight, 2 = moderate, 3 = severe and 4 = very severe). Patients were assessed before treatment and once a week during the treatment period. The maximum treatment period was 4 weeks.

The efficacy was evaluated by comparing PASI score at the end of treatment to PASI score at baseline using the Wilcoxon matched-pairs signed-rank test. PASI scores at baseline and 1, 2, 3 and 4 weeks after the start of treatment were compared using the non-parametric Friedman two-way analysis of variance. $p < 0.05$ was regarded as statistically significant.

RESULTS

Patients participated in the study until their psoriasis was markedly improved or cleared or they withdrew from the study. On average, patients participated for 3.3 weeks (range 1–4 weeks).

Of the 15 participating patients, the mean baseline score for the body site receiving treatment every day was 8.5 ± 2.7 and that for the site treated every other day was also 8.5 ± 2.7 . The PASI score decreased significantly on both sides of the body during the treatment ($p < 0.001$) (Fig. 1). At each assessment there was no statistically significant difference in PASI score between the two body sides during the 4-week treatment period.

DISCUSSION

Our study confirms the effectiveness of CCT for treatment of chronic psoriasis and indicates that treatment every other day has the same effect as daily treatment. To improve the skin penetration of coal tar, tar treatment was followed by a bath. Without doubt a significant amount of the tar is thereby removed from the skin surface. CCT-treated patients are UVA light-sensitive in the days following tar treatment; together with our data this provides clear evidence that tar has a prolonged effect in the skin. In our study patients had a bath every day; having a bath every other day might have produced an even better effect on the psoriasis, by removing less tar and thus extending the effectiveness of the treatment. Further studies need to be done to determine exactly the duration of the effect of tar in the skin.

CCT treatment three times a week allows the patient to be treated in an outpatient clinic every other weekday. This increases the possibility of patients receiving CCT treatment still being able to continue with their jobs if a little odor is tolerated. This may encourage more patients to accept CCT as an alternative to other local/systemic treatments.

The safety of CCT treatment has been regarded as one of its great virtues but other side-effects besides smell and staining of clothes (4) must be considered. Coal tar contains a large number of polycyclic aromatic hydrocarbons (PAH), which have been found in blood (6) and urine (7), but their systemic toxicity is still unknown. Absorbed PAH can be metabolized to reactive derivatives that bind to DNA (8) and these PAH-DNA binding products are thought to be involved in PAH-induced carcinogenesis (8). However, no clear evidence of an increased incidence of skin cancer has been reported in patients who have been exposed to therapeutic doses of coal tar (3, 9, 10).

In conclusion, replacing the traditional daily CCT treatment with treatment every other day in an outpatient clinic will without doubt be an attractive alternative for patients suffering from chronic plaque psoriasis.

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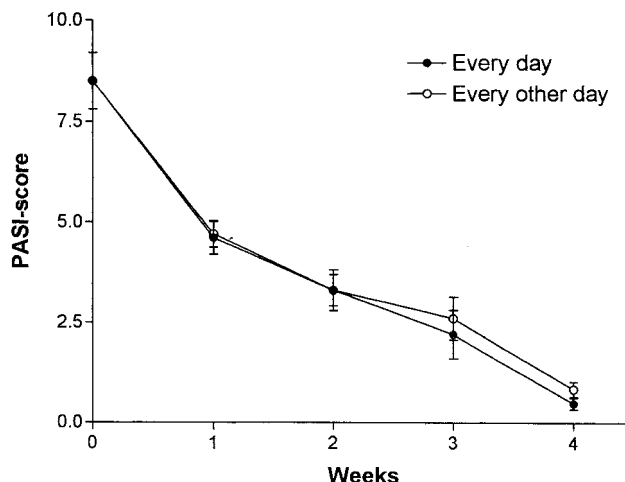


Fig. 1. Change in the modified PASI score during the 4-week treatment period. Bars represent standard deviations.

REFERENCES

- Schubert B, Juillard J. Tars. In: Dubertret L, ed. Psoriasis. Brescia: ISED, 1994: 116–122.
- Fisher LB, Maibach HI. Topical antipsoriatic agents and epidermal mitosis in man. *Arch Dermatol* 1973; 108: 374–377.
- Schmid MH, Korting HC. Coal tar, pine tar and sulfonated shale oil preparations: Comparative activity, efficacy and safety. *Dermatology* 1996; 193: 1–5.
- Griffiths WAD, Wilkinson JD. Topical therapy: Tars. In: Champion RH, Burton JL, Burns DA and Breathnach SM, eds. *Textbook of dermatology*. Oxford: Blackwell Science, 1998: 3535–3536.
- Fredriksson T, Pettersson U. Severe psoriasis—oral therapy with a new retinoid. *Dermatologica* 1978; 157: 238–244.
- Storer JS, DeLeon I, Millikan LE, Laseter JL, Griffing C. Human absorption of crude coal tar products. *Arch Dermatol* 1984; 120: 874–877.
- Hansen AM, Poulsen OM, Menne T. Longitudinal study of excretion of metabolites of polycyclic aromatic hydrocarbons in urine from two psoriatic patients. *Acta Derm Venereol* 1993; 73: 188–190.
- Schocket B, Horkay I, Kosa A, Paldeak L, Hower A, Grover PL, et al. Formation of DNA adducts in the skin of psoriasis patients, in human skin in organ culture, and in mouse skin and lung following topical application of coal-tar and juniper tar. *J Invest Dermatol* 1990; 94: 241–246.
- Van Schooten F-J, Godschalk R. Coal tar therapy. Is it carcinogenic? *Drug Saf* 1996; 15: 374–377.
- Jones SK, Mackie RM, Hole DJ, Gillis CR. Further evidence of the safety of tar in the management of psoriasis. *Br J Dermatol* 1985; 113: 97–101.

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