CASE REPORT

Bleomycin-induced flagellate hyperpigmentation

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SUMMARY
Bleomycin is a chemotherapeutic agent used in the treatment of different tumours. It has several side effects, including flagellate hyperpigmentation, which is a unique and a well-documented side effect of bleomycin therapy. We report a case of a 23-year-old woman with a personal history of ovarian dysgerminoma, who developed flagellate hyperpigmentation on trunk after bleomycin therapy.

BACKGROUND
Bleomycin is a chemotherapeutic agent, which has been used in the treatment of lymphoproliferative syndromes, squamous cell carcinomas, ovarian germ cell tumours and testicular tumours.1 2 Most organs contain a cysteine proteinase, capable of hydrolysing and inactivating bleomycin, except skin and lungs.2–4 The cutaneous-mucous side effects of bleomycin are diverse, including alopecia, stomatitis, nail changes and hyperpigmentation.2 Hyperpigmentation associated with bleomycin may be diffuse, patchy or linear.3 5 Flagellate hyperpigmentation is a linear or streaked pigmentation, which is a well-documented side effect of bleomycin therapy with an incidence as high as 20%.1 4

CASE PRESENTATION
A 23-year-old woman presented with flagellate hyperpigmentation on her trunk. She had a history of ovarian dysgerminoma, treated with unilateral salpingo-oophorectomy and adjuvant chemotherapy. Chemotherapy included 20 mg/m²/day of cisplatin and 100 mg/m²/day of etoposide intravenously, on days 1–5. She also received bleomycin, 30 mg intravenously on days 1, 8 and 15. The cycle was repeated every 3 weeks until three complete cycles. Within 4 weeks after the treatment, painless, pruritic, flagellate hyperpigmentation developed on her trunk. At presentation to our department, 5 months after the last cycle of chemotherapy, the skin examination revealed multiple curvilinear brownish hyperpigmented streaks, scattered profusely throughout her trunk (figures 1 and 2). There was no erythema, scaling or atrophy.

She had no alopecia and no changes in the appearance of nails or mucosa. She reported having no respiratory symptoms. On the basis of the history and examination, the diagnosis of bleomycin-induced flagellate hyperpigmentation was performed.

TREATMENT
Low-potency topical steroid was given to be applied to the pigmentation once a day, during 3 months. The patient was counselled that the flagellate hyperpigmentation usually fades over a period of several months after the cessation of bleomycin.

OUTCOME AND FOLLOW-UP
During 8 months of the follow-up, the hyperpigmentation in this patient persisted.

DISCUSSION
Bleomycin is a glycopeptide antibiotic commonly used for chemical pleurodesis, the treatment of cutaneous warts and the treatment of several human cancers, in particular lymphomas, ovarian and testicular germ cell tumours and squamous cell carcinomas.1 15 Flagellate hyperpigmentation can appear from 24 h to 9 weeks after the administration of bleomycin.1 6 It usually occurs after cumulative doses of 100–300 mg; however, it has been reported after using lower doses, and there is a case described in the literature of flagellate hyperpigmentation after intralesional injection of 14 units of bleomycin for plantar wart.5 Several hypotheses regarding the cause of hyperpigmentation have been proposed, including induced lesions by scratching the skin and by irritation from clothing, which permit the drug to leak out of blood vessels,6 increased levels of melanin because of various intercellular interactions and toxic effects of the drug itself.3 6

There is no specific treatment available for flagellate hyperpigmentation.3 5 Pruritus may be treated with topical steroids and oral antihistamines.3 4

The course of bleomycin-induced flagellate hyperpigmentation is variable.1 The pigmentation usually fades over 4 months, once the course of

Figure 1 Multiple curvilinear brownish hyperpigmented streaks, scattered throughout her upper trunk.
chemotherapy is complete and bleomycine is discontinued. Nevertheless, persistence of hyperpigmented streaks for up to 1 year after the treatment has been reported, like this case.34

Learning points

▸ Flagellate hyperpigmentation of the skin is a linear or streaked pigmentation, which is a well-documented side effect of bleomycin therapy with an incidence as high as 20%.
▸ There is no specific treatment available for flagellate hyperpigmentation, but pruritus may be treated with topical steroids and oral antihistamines.
▸ The pigmentation usually fades over 4 months, after bleomycin is discontinued. Nevertheless, persistence of hyperpigmented streaks for up to 1 year after treatment has been reported, like this case.

REFERENCES