Table 1. The causes of non-compliance with treatment of 16 patients.

	Time constraints (n)	Being tired of the treatment (n)	Problems with vascular access (n)
Month 0	3	7	6
Month 6	3	4	2
Month 12	3	4	0

non-compliant were monitored for 1 year, and it was determined that the rate of compliance increased only to 56.25% in our prospective cohort study.

Adolescent patients are more resistant to comply with recommended treatment plans. In this age group, the patients go through several biological, social, and emotional changes that influence their approach to the disorder [4]. Due to these factors, the non-compliance problem has a complicated nature that cannot be resolved through advising only. Treatment nonadherence is a chronic process in life-long chronic diseases such as hemophilia. As each patient is affected by different factors,

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it might be useful to conduct individual meetings with each patient instead of group trainings.

Keywords: Hemophilia, Compliance, Adolescent

Anahtar Sözcükler: Hemofili, Tedavi uyumu, Adölesan

**Conflict of Interest:** The authors of this paper have no conflicts of interest, including specific financial interests, relationships, and/or affiliations relevant to the subject matter or materials included.

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## **Bleomycin-Induced Flagellate Dermatitis**

Bleomisin ile İliskili Flagella Dermatit

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## To the Editor,

Bleomycin is a cytostatic, antineoplastic antibiotic that is used in both of the first-line treatments of Hodgkin lymphoma: ABVD (doxorubicin, bleomycin, vinblastine, dacarbazine) BEACOPP (doxorubicin, and bleomycin, vincristine, cyclophosphamide, etoposide, prednisone, procarbazine). The bleomycin hydrolase enzyme metabolizes bleomycin. This enzyme is not found in the skin or lung tissues; therefore, bleomycin accumulates in those areas and causes side effects [1]. The dermatologic side effects of bleomycin may vary from onycholysis, pruritus, and scleroderma-like skin changes to Stevens-Johnson syndrome. Flagellate dermatitis, resulting after bleomycin therapy, was originally described

by Moulin et al. [2] in 1970 as "bleomycin-induced linear hyperpigmentation" [3]. Although the term "flagellate dermatitis" was described for bleomycin-induced dermatitis, other causes of this symptom have been defined over time (Table 1) [4]. The characteristic symptoms are pruritic linear hyperpigmentations, arranged in a flagellate pattern and developing, in particular, on the trunk. Even though the exact mechanism is not clear, minor skin traumas are thought to be responsible since they increase blood flow to the affected area and cause drug accumulation [1].

We present a 24-year-old female patient who was diagnosed in August 2016 with stage IIA Hodgkin lymphoma (right cervical, submandibular, and bilateral palatine tonsil involvement was observed in positron emission tomography/computed tomography). A BEACOPP chemotherapy regimen was chosen for first-line therapy. After the second cycle of BEACOPP, the patient developed generalized and intense pruritus along with the appearance of papules and plagues on her back, shoulders, and trunk, with a remarkable whip-like mark formation (Figures 1 and 2), which evolved into hyperpigmentation. There was no evidence of mucosal or systemic involvement. Contrary to expectations, there was no evidence of dermatographia. Flagellate dermatitis was diagnosed by the clinical features. The patient did not have a history of dermatomyositis, Still's disease, hypereosinophilic syndrome, or shiitake mushroom intake. The BEACOPP regimen was interrupted after three cycles of chemotherapy were completed. The skin lesions started to resolve two weeks after the bleomycin-inducing therapy was suspended.

Bleomycin-induced flagellate dermatitis is a dose-dependent reaction that usually occurs with total doses above 100 U [5,6]. In contrast with these results, some patients develop

skin symptoms after low doses. The incidence of developing flagellate dermatitis and consequent hyperpigmentation after receiving bleomycin treatment is reported between 8% and 22% [7]. The lesions usually diminish 3-4 months after the interruption of the bleomycin treatment. Other than the suspension of the bleomycin treatment, no effective treatment has been reported for bleomycin-induced flagellate dermatitis. In the literature, there are some cases that report the use of topical or systemic corticosteroid treatments, as well as oral antihistamine treatments. However, it is stated that those treatments provide only symptomatic relief. The cessation of bleomycin is necessary to prevent further relapse [8]. We found it worthwhile to present our case since the development of this condition is rarely seen after a low dosage, the lesions disappear shortly after the suspension of the medication, and flagellate dermatitis is not observed with the other medications that our patient was receiving. Clinicians must be aware of this uncommon complication and act immediately to interrupt the causative agent.

Table 1. Causes of flagellate dermatitis.			
True flagellation/mechanical	Religious punishment, torture, abuse, sadomasochism, dermatitis artefacta		
Chemotherapy-induced	Bleomycin, peplomycin, docetaxel, bendamustine		
Rheumatologic disorders	Dermatomyositis, adult-onset Still's disease		
Toxin-induced	Shiitake mushroom ingestion, cnidarian stings, Paederus and other insects		
Other pruritic dermatitis	Dermatographism, excoriations from pruritic conditions, phytophotodermatitis, poison ivy dermatitis		
Hypereosinophilic syndrome	-		
Chikungunya fever-induced	-		
Idiopathic	-		



Figure 1. Flagellate dermatitis on trunk.



Figure 2. Flagellate dermatitis on extremity.

Keywords: Bleomycin, Hodgkin lymphoma, Flagellate dermatitis

Anahtar Sözcükler: Bleomisin, Hodgkin Lenfoma, Flagella dermatit

Informed Consent: Received.

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