

Imatinib-Induced Psoriasis

İmatinibe Bağlı Psoriasis Olgusu

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To the Editor,

Imatinib is a signal transduction regulator that selectively inhibits the tyrosine kinase family, including bcr-abl and c-kit, and the platelet-derived growth factor (PDGF) receptor. It is currently the first-line therapy for newly diagnosed chronic myeloid leukemia (CML) patients [1]. We report the case of a patient who had no previous history of psoriasis but developed psoriasis after starting imatinib.

A 21-year-old woman was diagnosed with CML in the chronic phase. Imatinib mesylate was started at a daily dose of 400 mg. The patient achieved a complete hematological response within 3 months. Five months after her CML diagnosis and imatinib usage, she developed an erythematous scaly eruption with plaques of various sizes on her trunk and extremities (Figures 1, 2, and 3). She had no previous history of psoriasis and had not taken any drugs except for imatinib, nor did she have any relatives with a history of psoriasis. The patient underwent a skin biopsy, which revealed a neutrophilic scale crust and loss of the granular cell layer, which are most consistent with psoriasis (Figure 4). The discontinuation of imatinib treatment and subsequent introduction of narrowband ultraviolet B therapy improved the skin condition, and her psoriatic skin lesions had almost disappeared within 3 weeks. Since that time, nilotinib has been started. So far, the patient has not complained of any cutaneous side effects, and she achieved a complete cytogenetic response at 6 months and remains clinically well, currently receiving nilotinib at a dose of 200 mg twice daily.

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Received/*Geliş tarihi* : October 09, 2013 Accepted/Kabul tarihi : January 14, 2013 Cutaneous reactions to imatinib are common and may occur in 7% to 88.9% of patients in different series. Maculopapular eruptions, erythematous eruptions, edema, and periorbital edema are the most common adverse events seen [2]. In 2002, Miyagawa et al. reported a patient who had intractable psoriasis but experienced significant improvement while being treated with imatinib for



Figure 1: Scaly erythematous papules and plaques on the right arm.

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Figure 2: Erythematous papulosquamous lesions on the lower extremities.



Figure 3: Scaly erythematous papules and plaques on the neck, upper extremities, and and trunk.

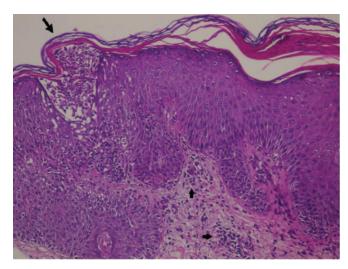


Figure 4: Parakeratosis, leukocyte abscesses in the keratin layer (big arrow), psoriasiform hyperplasia of the epithelium, loss of the granular layer areas of parakeratosis, leukocyte abscesses in the surface epithelium, superficial perivascular lymphocytes (small arrows), extravasated erythrocytes.

concomitant metastatic gastrointestinal stromal tumors [3]. Valeyrie et al. also reported psoriatic dermatological changes in 4 out of 54 patients who were using imatinib. Two of these 4 patients had no history of psoriasis [2,4]. Psoriasis has long been identified as an immune disorder in which T lymphocytes play a primary role in the pathogenesis. Imatinib affects cytokine production and the proliferation of T cells and inhibits the secretion of interferon-c by T effector cells. These effects, together with imatinib's suppression of c-kit and PDGF receptors, may help to explain the exacerbation of psoriasis in some patients [5,6,7,8,9]. The cause of imatinibrelated nonpsoriatic forms of skin lesions is not clear. The most probable cause is the fact that imatinib affects mast cells. Because mast cells express a functional c-kit, which is susceptible to imatinib, this drug causes mast cells to proliferate. Another mechanism involves chemoattractant substances, such as cytokines and growth factors, which can lead to an accumulation of dermal mast cells. Imatinibrelated skin toxicities are usually dose-dependent, and skin biopsies have shown a mixed cellular infiltrate [10]. In our case, we replaced imatinib with nilotinib therapy, and the psoriasis has not recurred over the course of about 1 year. The patient still maintains a complete molecular and hematological response. Imatinib-induced skin reactions can be self-limiting, but occasionally drug withdrawal is required. This case demonstrates that imatinib can cause psoriasis to occur or can exacerbate the condition. In cases where patients using imatinib develop psoriasis, nilotinib can be a safe alternative.

Key words: Chronic Myelogenous Leukemia, Imatinib, Psoriasis

References

- 1. Scheinfeld N. Imatinib mesylate and dermatology part 2: a review of the cutaneous side effects of imatinib mesylate. J Drugs Dermatol 2006;5:228-231.
- 2. Heidary N, Naik H, Burgin S. Chemotherapeutic agents and the skin: an update. J Am Acad Dermatol 2008;58:545-570.
- 3. Miyagawa S, Fujimoto H, Ko S, Hirota S, Kitamura Y. Improvement of psoriasis during imatinib therapy in a patient with a metastatic gastrointestinal stromal tumour. Br J Dermatol 2002;147:406-407.
- 4. Valeyrie L, Bastuji-Garin S, Revuz J, Bachot N, Wechsler J, Berthaud P, Tulliez M, Giraudier S. Adverse cutaneous reactions to imatinib (STI-571) in Philadelphia chromosome-positive leukemias: a prospective study of 54 patients. J Am Acad Dermatol 2003:48:201-206.
- Amitay-Laish I, Stemmer SM, Lacouture ME. Adverse cutaneous reactions secondary to tyrosine kinase inhibitors including imatinib mesylate, nilotinib, and dasatinib. Dermatol Ther 2011;24:386-395.

- 6. Dickens E, Lewis F, Bienz N. Imatinib: a designer drug, another cutaneous complication. Clin Exp Dermatol 2009;34:603-604.
- 7. Krane JF, Murphy DP, Gottlieb AB, Carter DM, Hart CE, Krueger JG. Increased dermal expression of platelet-derived growth factor receptors in growth-activated skin wounds and psoriasis. J Invest Dermatol 1991;96:983-986.
- 8. Yamamoto T, Katayama I, Nishioka K. Possible contribution of stem cell factor in psoriasis vulgaris. J Dermatol Sci 2000:24:171-176.
- 9. Leder C, Ortler S, Seggewiss R, Einsele H, Wiendl H. Modulation of T-effector function by imatinib at the level of cytokine secretion. Exp Hematol 2007;35:1266-1271.
- 10. Thachil J. T-regulatory cell response in psoriasis and changes with imatinib therapy. Clin Exp Dermatol 2009;34:e1022.