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# A rare condition: Montelukast allergy

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### Abstract:

Montelukast, selective leukotriene (LT) receptor antagonist specific for cysteinyl LT type 1 receptors, serves as an alternative treatment option for asthma and different allergic clinical conditions. However montelukast, itself, may rarely induce hypersensitivity reactions. Although rare, clinicians, especially those working in pulmonology and allergy clinics, should be aware of the potential for montelukast to cause hypersensitivity reactions. Herein, we present a 61-year-old female patient who was followed up with Samter's syndrome and developed urticarial rashes after montelukast treatment, and montelukast allergy was confirmed by oral drug provocation test.

### Keywords:

Allergy, asthma, hypersensitivity, leukotrienes, montelukast, pulmonary medicine

## Introduction

Leukotrienes (LTs) are mediator molecules that play a pivotal role in the pathogenesis of asthma, leading to mucus secretion and bronchoconstriction. Montelukast is a selective Leukotriene Receptor Antagonist (LTRA) targeting cysteinyl LT type 1 receptors, providing an alternative treatment approach for asthma.<sup>[1]</sup> Apart from its antagonistic effect on LTs, montelukast contributes to bronchodilation by activating beta-2 receptors. It is commonly employed in pulmonary diseases and allergy clinics for various indications. First, according to international

guidelines for asthma and allergic rhinitis, montelukast is recommended for mild, moderate, and severe asthma, as well as for rhinitis that is unresponsive to nasal steroids.<sup>[1,2]</sup> Second, montelukast is sometimes used in conjunction with antihistamines in specific urticaria cases.<sup>[3]</sup> Third, montelukast is suggested as a premedication in desensitization protocols applied for drug allergies.<sup>[4]</sup> Lastly, using LTRA in the Samter triad has shown benefits in alleviating nasal symptoms and increasing asthma control.<sup>[5]</sup> Nonetheless, montelukast, although employed for various allergic conditions, infrequently induces hypersensitivity reactions.

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## Case Report

A 61-year-old female patient diagnosed with Samter syndrome presented to the allergy clinic complaining of month-long redness, swelling, and itching on her body. Her treatment regimen included inhaled budesonide-formoterol fumarate combination, intranasal fluticasone propionate, and oral montelukast and bilastine. She indicated that her symptoms worsened, especially after administering bilastine and montelukast. Her physical examination revealed widespread urticarial lesions across her body, but her systemic examination was otherwise normal.

Laboratory tests, which encompassed complete blood count, renal and liver function tests, blood glucose and serum electrolytes, urinalysis, complement component 4 (C4), total immunoglobulin E (IgE), and serum triptase levels, were within standard parameters. Rheumatologic markers such as cytoplasmic anti-neutrophil cytoplasmic antibodies (c-ANCA), perinuclear anti-neutrophil cytoplasmic antibodies (p-ANCA), anti-double-stranded DNA (anti-dsDNA), and antinuclear antibodies (ANA), along with hepatitis serology, were negative. No pathogens were identified in the urine culture or the parasitic stool examination.

To manage the patient's urticaria symptoms, oral methylprednisolone 40 mg/day was initiated. Due to

the patient's drug history that involved simultaneous use of bilastine and montelukast, LTRA and antihistamine were halted pending further assessments. The patient's symptoms receded within five days following systemic steroid administration, after which the treatment was discontinued. Drug provocation tests were subsequently planned. In the interim, when the patient sought treatment at the dermatology clinic, ebastine was prescribed following a biopsy of the lesions, which were found to be consistent with urticaria. On her follow-up visit to our clinic, she reported taking ebastine 20 mg daily for three months without any issues. Urticarial plaques did not reappear during the 2-month follow-up after discontinuing ebastine. Both her asthma and rhinitis were managed effectively with inhaler and intranasal treatments. Subsequent tests included a prick-to-prick skin test with montelukast 10 mg tablets, which yielded a negative result. An oral provocation test with montelukast began with an initial dose of 0.833 mg, followed by doses of 1.66 mg, 2.5 mg, and 5 mg at one-hour intervals. Itching, redness, and swelling manifested around the oral area and on both gluteal regions seven hours after the final dose [Fig. 1]. These reactions subsided naturally within 5-6 hours without any residual marks. Given the urticarial response seven hours after the montelukast oral provocation test, the patient was diagnosed with a montelukast allergy.

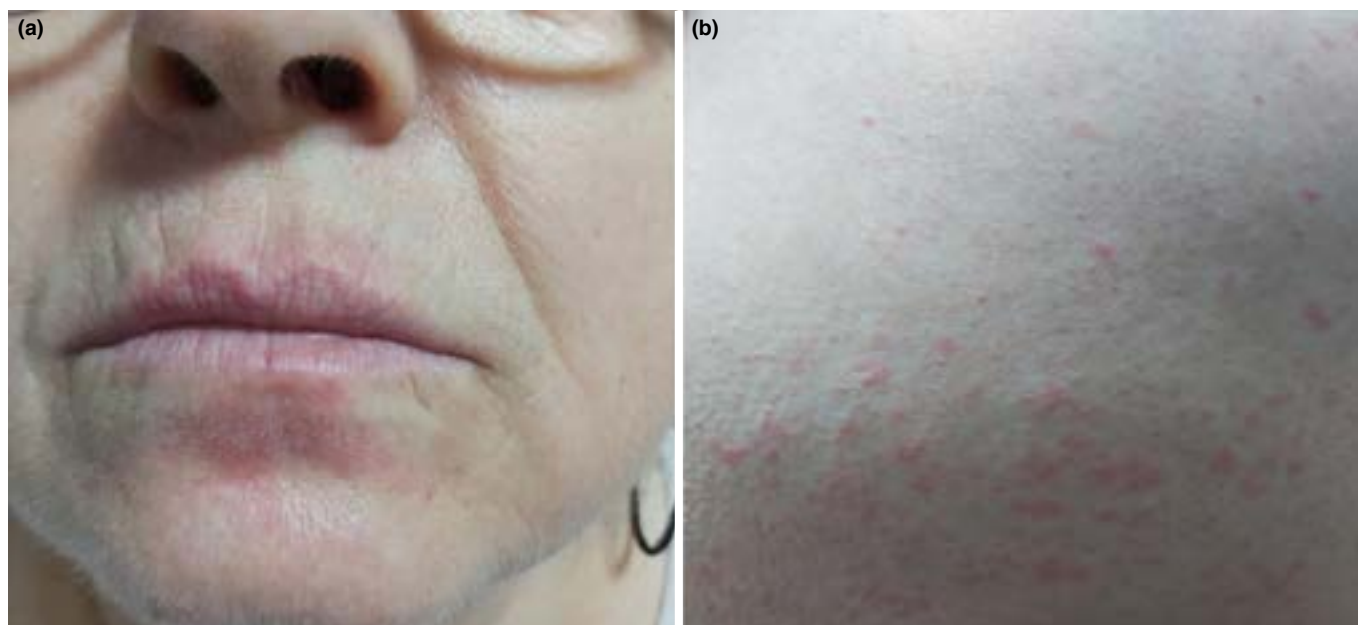


Figure 1: Redness in the peroral area (a) and the gluteal region (b)

## Discussion

For asthma management, it is advised to commence treatment with inhaled corticosteroids (ICS) as an initial step and to integrate combination therapies as the treatment intensifies. Present Global Initiative for Asthma (GINA) guidelines advocate LTRAs as an alternative treatment option beginning from the second step of asthma management.<sup>[1]</sup> Montelukast, utilized for treating asthma, allergic rhinitis, and urticaria, is generally well-received by patients. Nonetheless, although rare, side effects such as anaphylaxis, angioedema, muscle weakness, elevated transaminases, eosinophilia, sleep disturbances, depression, and agitation can occur.<sup>[6]</sup> Eosinophilic granulomatous polyangiitis, a seldom-seen vasculitis associated with asthma, has had cases linked to montelukast use, even though its exact pathogenesis remains elusive.<sup>[7,8]</sup> There have been reports in literature of dermatitis and urticaria arising post-montelukast consumption. Often, these instances feature concurrent use of other medications alongside montelukast.<sup>[9, 10]</sup>

In the case presented, we report urticaria development following montelukast use due to Samter's syndrome. The patient's allergy to montelukast was confirmed through an oral drug provocation test. In previously reported montelukast allergy cases, unlike this one, diagnoses were often based on patient histories, many of which involved the concurrent use of various drugs.

While montelukast allergies are extremely rare, clinicians, especially those in pulmonary diseases and allergy clinics where this drug is frequently prescribed, should be aware of the potential for rare hypersensitivity reactions to montelukast.

### Informed Consent

Written informed consent was obtained from the patient for the publication of the case report and the accompanying images.

### Conflicts of interest

There are no conflicts of interest.

### Financial support and sponsorship

Nil.

## Peer-review

Externally peer-reviewed.

## Authorship Contributions

Concept – F.D.Ç., H.Ç.T., M.Y., Ö.A., O.T., K.A.; Design – F.D.Ç., H.Ç.T., M.Y., Ö.A., O.T., K.A.; Supervision – F.D.Ç., H.Ç.T., M.Y., Ö.A., O.T., K.A.; Funding – F.D.Ç., H.Ç.T., M.Y., Ö.A., O.T., K.A.; Materials – F.D.Ç., H.Ç.T., M.Y., Ö.A., O.T., K.A.; Data collection &/or processing – F.D.Ç., H.Ç.T., M.Y., Ö.A., O.T., K.A.; Analysis and/or interpretation – F.D.Ç., H.Ç.T., M.Y., Ö.A., O.T., K.A.; Literature search – F.D.Ç., H.Ç.T., M.Y., Ö.A., O.T., K.A.; Writing – F.D.Ç., H.Ç.T., M.Y., Ö.A., O.T., K.A.; Critical review – F.D.Ç., H.Ç.T., M.Y., Ö.A., O.T., K.A.

## References

1. Global Initiative for Asthma. Global Strategy for Asthma Management and Prevention, 2023. Updated May 2023. Available at: www.ginasthma.org. Accessed May 29, 2023.
2. Bousquet J, Schünemann HJ, Togias A, Bachert C, Erhola M, Hellings PW, et al.; Allergic Rhinitis and Its Impact on Asthma Working Group. Next-generation Allergic Rhinitis and Its Impact on Asthma (ARIA) guidelines for allergic rhinitis based on Grading of Recommendations Assessment, Development and Evaluation (GRADE) and real-world evidence. *J Allergy Clin Immunol* 2020;145(1):70–80.e3. Erratum in: *J Allergy Clin Immunol* 2022;149(6):2180.
3. Wan KS. Efficacy of leukotriene receptor antagonist with an anti-H1 receptor antagonist for treatment of chronic idiopathic urticaria. *J Dermatolog Treat* 2009;20(4):194–7. [[CrossRef](#)]
4. Breslow RG, Caiado J, Castells MC. Acetylsalicylic acid and montelukast block mast cell mediator-related symptoms during rapid desensitization. *Ann Allergy Asthma Immunol* 2009;102(2):155–60.
5. Lee RU, Stevenson DD. Aspirin-exacerbated respiratory disease: evaluation and management. *Allergy Asthma Immunol Res* 2011;3(1):3–10. [[CrossRef](#)]
6. Callero-Viera A, Infante S, Fuentes-Aparicio V, Zapatero L, Alonso-Lebrero E. Neuropsychiatric reactions to montelukast. *J Investig Allergol Clin Immunol* 2012;22(6):452–3.
7. Cuchacovich R, Justiniano M, Espinoza LR. Churg-Strauss syndrome associated with leukotriene receptor antagonists (LTRA). *Clin Rheumatol* 2007;26(10):1769–71. [[CrossRef](#)]
8. Currie GP, McKinlay L, Kerr KM. Histological appearances of putative montelukast related Churg-Strauss syndrome. *Thorax* 2008;63(12):1120. [[CrossRef](#)]
9. Herzinger T, Ludolph-Hauser D, Przybilla B. Urticaria triggered by antiallergy treatment. *Clin Exp Dermatol* 2008;33(4):519–20. [[CrossRef](#)]
10. Tedeschi A. Paradoxical exacerbation of chronic urticaria by H1-antihistamines and montelukast. *Eur Ann Allergy Clin Immunol* 2009;41(6):187–9.