



## BILASTINE USED IN TREATMENT OF ALLERGIC RHINITIS AND URTICARIA

### Dermatology

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### ABSTRACT

Bilastin is a new approved second generation oral H1 Antihistaminic drug which is generally used in several countries for symptomatic treatment of Allergic Rhinitis (AR), Chronic Urticaria (CU) and hives. Oral bilastine is a non-sedating H1 antihistaminic antagonist. H1-receptor played most vital role in control allergic inflammation by directly interfering with histamine H1 receptor. Allergic rhinitis is not a serious disease but it is globally health problem. Bilastine does not cross blood brain barrier (BBB). Bilastine has not cytochrome p450 activity. Absolute bioavailability of Bilastine 60 – 61 %. Bilastine bioavailability much more in fasting as compare to with meal. With meal decrease the absorption time of bilastine and time increase to reach maximum concentration (Cmax). Bilastine have a great affinity to protein binding 84-94%. Bilastine approximately 95 % excreted out, faeces or stool (67%), urine (33%).

### KEYWORDS

Urticaria, Bilastine, blood brain barrier, faeces.

### INTRODUCTION

Bilastine in urticaria management: Bilastin is a new approved second generation oral H1 Antihistaminic drug which is generally use in several countries for symptomatic treatment of Allergic Rhinitis (AR), Chronic Urticaria (CU) and hives.<sup>1,2</sup> Oral bilastine is a non-sedating H1 antihistaminic antagonist. H1-receptor played the most vital role in control allergic inflammation by directly interfering with histamine H1 receptor.<sup>3,4</sup>

Urticaria named is derived from nettle plant (Latin, urtica), which is now known to contain histamine. Histamine has played a mediator role control in urticaria. Histamine has been recognized for many years to symptomatic treatment of urticaria.<sup>9</sup> urticaria is a common disease generally affecting approximately 15-20% population at list once during in life time. Even it may happen in any age, it is most common in age group 20 to 40 years.<sup>10</sup> It mostly found in developing (Asian) country. Allergenic symptoms of acute urticaria can be identified up to 60 to 80% while this figure is lower as compare to chronic urticaria.<sup>10</sup> Urticaria patient develop typically wheals (hives), angioedema, inflammation and flare.<sup>8</sup> Wheals are small lived elevated erythematous lesions which range from a few millimeters to centimeters in diameter and can confluent.<sup>10,11</sup> Erythematous lesion burning and unbearable and worse in the evening and night time. At least half of urticaria patient reported sleep disturbance.<sup>11</sup>

Bilastin H1-antihistamines is a nonsedating agent it is newly approved the symptomatic treatment of acute and chronic urticaria, and current guideline for the treatment of acute and chronic urticaria.<sup>6</sup> Main goal of urticaria management provide symptomatic treatment, relief to patient by antagonizing the effect (etching, hives, inflammation) of histamine from H1 receptor.<sup>5</sup> Bilastine have a high affinity and selectivity for histamine H1 receptor, and great influence anti allergic activity.<sup>7</sup>

Allergic rhinitis: The rate of allergy is increasing in middle and low class country especially in Asian country.<sup>14</sup> Allergic rhinitis is not a serious disease but it is globally health problem.<sup>13</sup> Allergic rhinitis is a worldwide common disorder which mostly effect 10-40% population. AR show the negative effect in patient, and it can be affect quality of life of patient and decrease night sleep.<sup>12</sup> AR patient suffer nasal congestion, etching, rhinorrhea and sneezing result of IgE mediated inflammation. And those who has suffer AR show the symptoms in eyes for example redness, teary, and etching in eyes. allergic rhinitis patients suffer some nonnasal symptoms thirst, headache, cough, wheezing, sore throat etc.<sup>12,13,14</sup> Allergy is a common and often recognized cause of respiratory distress, and discomfort among in primary care patient.<sup>14</sup>

Bilastine show great efficacy in allergic rhinitis and urticaria. Bilastine 20 mg oral once a day is more effective in allergic rhinitis and other allergic disease.<sup>15</sup> Bilastine is as effective fexofenadine 120 mg, cetirizine 10 mg it show great onset of action, after take a dose after one hour it should reduce allergic symptoms. Single dose of bilastine is effective upto 26 hour after intake.<sup>15,16,23,25</sup> Bilastine take a single dose daily orally and no show sedation and cardiac toxicity. Bilastine has a good binding property to H1 receptor.<sup>16</sup>

Brief pharmacokinetic of Bilastine: Bilastine is not structurally derived from any antihistamine it belongs to piperidine derivative activity.<sup>1</sup> Bilastine is new discovered second generation H1 antihistamine.<sup>15</sup> Bilastine has great affinity to H1 receptor, and H1 inverse agonist. His affinity is 3-6 times more than as compare to cetirizine and fexofenadine.<sup>1</sup> Absorption of Bilastine fully dose dependent, taken in fasting stage fast absorption. Bilastine not taken with juice, food and meal delay absorption. Bilastine is recommended to be taken at least 1 hr before meal and 2 hr after meal. Empty stomach increased absorption.<sup>5</sup> Bilastine rapidly absorbed after oral administration. Bilastine show great affinity and antihistamine activity in time interval 30 minutes to 8:17 hrs.<sup>5</sup> Bilastine does not interact with benzodiazepenes.<sup>15,16</sup>

Bilastine has taken maximum time (Tmax to reach Cmax 1hour 13 minute. Absolute bioavailability of Bilastine 60 – 61 %. Bilastine protein binding value was 84 to 94%.<sup>4,19</sup> No accumulation in repeated dose.<sup>5,19</sup>

Bilastine has no interact on cytochrome p450 system and does not undergo significant metabolism in human system. Bilastine does not cross blood brain barrier (BBB).<sup>19</sup> It did not show the sedation neurotoxic activity.<sup>5</sup> Bilastin 20 mg orally is safe in hepatotoxic patient.<sup>4,18,19</sup>

Bilastine do not show hepatic metabolism and it excreted out by renal excretion. Bilastine approximately 95 % excreted out, faeces or stool (67%), urine (33%).<sup>4</sup> It is excreted out in unchanged form.<sup>5</sup>

### Few steps to therapeutic use of bilastine in allergic rhinitis and urticaria.

1	Bilastine is second generation H1 antihistaminic. <sup>5</sup>
2	Bilastine 20 mg dose once a day daily dose for symptomatic treatment of urticaria and AR. <sup>2</sup>
3	Bilastine has been no sedative and cardio toxic effect. <sup>2</sup>
4	Bilastine is well tolerated in seasonal allergic rhinitis and urticaria. <sup>19</sup>

5	Adverse event of bilastine 0.5% and not any death and serious event recorded.19
6	According to international guideline second generation H1 antihistamine first line treatment in elderly urticaria and allergic rhinitis patient.21
7	Dose adjustment not required in hepatic and renal patient.21
8	Bilastine 95% excreted out faeces and urine.5
9	Bilastine approved in Europe in treatment of urticaria and allergic rhinitis in a children >6 year old patient.22
10	Nonsedating antihistamine has been low lipid solubility.20
11	Bilastine show effectiveness in reducing the symptoms of urticaria and allergic rhinitis.23,24

## DISCUSSION:

The efficacy and safety is much required for patient quality life. Bilastine show great bioavailability in oral dose. Bilastine is safe in hepatic and renal failure patient but give dose once a day some study is required in this field. Bilastine is much safer as compare to other antihistaminic drug. Bilastine is safe or not in pregnant women, more study is required in this topic.

## CONCLUSION:

In this review article important resources have been focused on reducing the incidence of the allergic response by new anti-allergic drugs as bilastine. Bilastine epitomizes the evolution of research on antihistamines concerning both efficacy and safety in humans. Bilastine 20 mg once a day is safe in hepatic and renal patient. Bousquet et al reviewed the available literature and found in his study bilastine 20 mg OD in a day improved nasal airway symptoms in allergic rhinitis. Bilastine is low potential for central nervous system. Bilastine has no drug-drug interaction. No drug adjustment need in urticaria and allergic rhinitis patient.26, 27 Bilastine has a fast onset of action, bilastine reach Cmax 30 minutes to 1 hour and his 95 % protein binding ability.22,25,26 Non sedating antihistamine agent has been low lipid solubility.20

## REFERENCES:

1. Jeet Gandhi, Kiran Godse, Gauri Godse Consultant Dermatologist, Sakhiya Skin Clinic, Surat, Gujarat, Professor of Dermatology, D Y Patil Hospital, Shree Skin Centre, Navi Mumbai, Maharashtra, India: Bilastine, A Novel Antihistamine. Indian Journal of Drugs in Dermatology. January-June 2018; 4(1).
2. Zoltan Novak, Anahi Yanez, Ildiko Kiss, Piotr Kuna, Miguel Tortajada-Girbes & Roman Valiente 6: Safety and tolerability of bilastine 10 mg administered for 12 weeks in children with allergic diseases & the "Bilastine Paediatric Safety Study Group. Accepted for publication 20 February 2016 DOI:10.1111/pai.12555. *Pediatr Allergy Immunol* 2016; 27: 493–498.
3. Bachert P, Kuna P & Zuberbier T: Bilastine in allergic rhinoconjunctivitis and urticaria. *C. Accepted for publication 13 April 2010 DOI:10.1111/j.1398-9995.2010.02404.x. Allergy* 2010; 65 (Suppl. 93): 1–13.
4. Erminia Ridolo, Marcello Montagni, Laura Bonzano, Cristoforo Incorvaia and Giorgio Walter Canonica & Ridolo et al: Bilastine new insight into antihistamine treatment *Clinical and Molecular Allergy*. (2015); 13:1 DOI 10.1186/s12948-015-0008-x.
5. Kiran Godse, Abhishek De, Vijay Zawar, Bela Shah, Mukesh Girdhar, DS Krupa Shankar, Nidhi Sharma, Murlidhar Rajagopalan, & Anant Patil: Review Article Bilastine for the treatment of chronic spontaneous urticarial. Consensus statement for Indian patients *IP Indian Journal of Clinical and Experimental Dermatology* 5 (2019); 180–185.
6. Amalia Leceta, Ander Sologuren, Roman Valiente, Cristina Campo, Luis Labeaga, Leceta A, Sologuren A, Valiente R, Campo C & Labeaga L: Bilastine in allergic rhinoconjunctivitis and urticarial. a practical approach to treatment decisions based on queries received by the medical information department Bilastine. *Drugs in Context* 2017; 6: 212500. DOI: 10.7573/dic.212500 I of 12 ISSN: 1740-4398.
7. Michinori Togawa, Hidetoshi Yamaya, Monica Rodriguez & Hirotaka Nagashima: Pharmacokinetics, Pharmacodynamics and Population Pharmacokinetic/Pharmacodynamic Modelling of Bilastine, a Second-Generation Antihistamine, in Healthy Japanese Subjects. *Clin Drug Investig* (2016); 36:1011–1021 DOI 10.1007/s40261-016-0447-2.
8. Yadav narayanan, pongsakorn tantilipikorn et al, Treatment of allergic rhinitis and urticaria: A review of the newest antihistamine drug bilastine. *Therapeutics and Clinical Risk Management* April 2016 DOI: 10.2147/TCRM.S105189.
9. Sudha Yadav, Amitabh Upadhyay & AK Bajaj: Chronic urticarial. An overview. *Year* 2006; 51 (3):171-177.
10. Ashimav Deb Sharma & Shimav Deb Sharma: Use of patch testing for identifying allergen causing chronic urticarial. July 11, 202; IP: 103.104.54.58.
11. Bettina Wedi, Ulrike Raap, Dorothea Wiecek & Alexander Kapp: Urticaria and infections Published. 1 December 2009 *Allergy, Asthma & Clinical Immunology* 2009; 5:10 doi:10.1186/1710-1492-5-10.
12. Martin K. Church, Marysia Tiongco-Recto, Erminia Ridolo & Zoltan Novak: Bilastine a lifetime companion for the treatment of allergies. 2020; 36(3): 445–454 https://doi.org/10.1080/03007995.2019.1681134 Article ST-0354.R1/1681134.
13. I Jáuregu, J Bartra, A del Cuvillo, I Dávila, M Ferrer, J Montoro, J Mulla, J Sastre & A Valero: Bilastine and Quality of Life. *J Investig Allergol Clin Immunol* 2011; 21 (Suppl. 3): 16-23.
14. Ralph Mösges, Dennis Lip Yen Lee, Jovilia Abong, Bella Siasoco, Steven KW Chow, Jern-Lin Leong, Harvinder Singh, S Kuljit & Benjamin Campomanes: Role of bilastine in the management of allergic rhinitis and urticarial. An Asia-Pacific consensus statement, <http://dx.doi.org/10.5415/apallergy.2016.6.1.56> *Asia Pac Allergy* 2016; (6):56-66.
15. Erminia Ridolo, Marcello Montagni, Laura Bonzano, Cristoforo Incorvaia & Giorgio Walter Canonica: Bilastine, new insight into antihistamine treatment. *Clinical and Molecular Allergy* (2015); 13:1 DOI 10.1186/s12948-015-0008-x.

16. Reggie Bosmaa, Jelle van den Bora, Henry F. Vischera, Luis Labeaga, Rob Leursa: The long duration of action of the second generation antihistamine bilastine coincides with its long residence time at the histamine H1 receptor. *nal of Pharmacology*, Volume 838, 5 November 2018; Pages 107-111.
17. Xue Yan Wang, Margaret Lim-Jurado, Narayanan Prepageran, Pongsakorn Tantilipikorn & De Yun Wang: Treatment of allergic rhinitis and urticarial. A review of the newest antihistamine drug bilastine. 2016; 12: 585–597.
18. Akiko YAGAMI, Masataka FURUE, Michinori TOGAWA, Akihiro SAITO & Michihiro HIDE: One-year safety and efficacy study of bilastine treatment in Japanese patients with chronic spontaneous urticaria or pruritus associated with skin diseases. doi: 10.1111/1346-8138.13644 *Journal of Dermatology* 2017; 44: 375–385.
19. F. SCAGLIONE: Safety profile of Bilastine, 2nd generation h1-antihistamines. 2012; 16: 1999-2005.
20. A. DEMONTE, M.B. GUANTI, S. LIBERATI, A. BIFFI, F. FERNANDO, M. FAINELLO, P. PEPE: Bilastine safety in drivers who need antihistamines new evidence from high-speed simulator driving test on allergic patients. 2018; 22: 820-828.
21. Ander Sologuren, M.D., Rosa Vinas M.D., Esther Cordon M.D., Susana E. Riesgo M.D., Maria del Mar Fores M.D., Maria Rosa Senan M.D., Sonia Fernandez Pharm.D., Luis Labeaga, B.Sc., and Manuel Ruiz-Mijang M.D: Open-label safety assessment of bilastine in elderly patients with allergic rhinoconjunctivitis and/or urticaria. DOI: 10.2500/aap.2018; 39:4136.
22. Atsushi Fukunaga, Yoshiko Oda, Ken Washio, Takashi Omori, Yasumasa Kakei, Michihiro Hide, Ch150kako Nishigori & Fukunaga et al: Efficacy of switching to bilastine, a histamine H1 receptor antagonist, in patients with chronic spontaneous urticaria (H1-SWITCH). Study protocol for a randomized controlled trial *Trials*. (2020); 21:23 <https://doi.org/10.1186/s13063-019-3878-2>.
23. K. Krause, A. Spohr, T. Zuberbier, M.K. Church & M. Maurer: Up-dosing with bilastine results in improved effectiveness in cold contact urticaria. 2013; 68: 921–928. Accepted for publication 13 March 2013 DOI:10.1111/all.12171.
24. Kazuhiro Hashiguchi a, Ken-ichiro Wakabayashi b, Michinori Togawa c, Akihiro Saito c, Kimihiro Okubo d & K. Hashiguchi et al: Therapeutic effect of bilastine in Japanese cedar pollinosis using an artificial exposure chamber (OHIO Chamber). / *Allergology International* (2017); 66: 123e131.
25. Michihiro Hide a, Akiko Yagami b, Michinori Togawa c, Akihiro Saito c, Masataka Furue d. M. Hide et al: Efficacy and safety of bilastine in Japanese patients with chronic spontaneous urticarial. A multicenter, randomized, double-blind, placebo-controlled, parallel-group phase II/III study *lergology International* (2017); 66: 317e325.
26. Nikolaos G. Papadopoulos and Torsten Zuberbier: The safety and tolerability profile of bilastine for chronic urticaria in children. (2019); 9:55 <https://doi.org/10.1186/s13601-019-0294-3>.
27. Xue Yan Wang, Margaret Lim-Jurado, Narayanan Prepageran, Pongsakorn Tantilipikorn & De Yun Wang: Treatment of allergic rhinitis and urticarial, a review of the newest antihistamine drug bilastine. 2016; 12: 585–597.
28. Yagyesh Kapoora & Kapil Kumara: Structural and clinical impact of anti-allergy agents. An overview *Bioorganic Chemistry* (2020); 94: 103351.