Medicines to Avoid Before Skin Allergy Testing



he American Academy of Otolaryngic Allergy (AAOA) has developed this position statement to assist allergy providers in determining which medicines patients should avoid prior to skin testing. These medicines are known to decrease or eliminate skin reactivity causing a negative histamine control. Providers should have a thorough understanding of the classes of medicines that could interfere with allergy testing. With proper patient counseling, the goal is to yield interpretable skin results without unnecessary medicine discontinuation.

Antihistamines suppress the histamine response for a variable period of time. In general, first-generation antihistamines can be stopped for 72 hours, however, several types including Cyproheptadine (Periactin) can have active histamine suppression for up to 11 days. Second-generation antihistamines also suppress testing for a variable length of time, up to 7 days. Astelin (Azelastine) nasal spray has been shown to suppress testing for up to 48 hours. 1, 2, 3, 4, 5, 6, 7

Short-term oral corticosteroids (30 mg daily for a week) do not suppress skin testing.⁸ There is a difference of opinion about the effects of long-term or relatively high-dose steroids, i.e. greater than 20 mg of prednisone per day, on the suppression of immediate skin tests.^{9, 10}

Topical glucocorticosteroids can block the histamine response.^{11, 12, 13} Application of potent topical steroids have been shown to stop the histamine response for up to three weeks.¹⁴

Tricyclic antidepressants can suppress the antihistamine response from 7 to 14 days depending upon the type. 15, 16

Benzodiazepines should be discontinued for 7 days before the testing and include clonazepam, diazepam, lorazepam, and midazolam.¹⁵ Alprazolam has also been shown to inhibit skin testing.¹⁷

H2 blockers have the potential to suppress histamine skin reactions for up to two days and include cimetidine, ranitidine, and famotidine.^{18, 19}

- Bernstein, L. et al Allergy Diagnostic Testing: an updated practice parameter. Annals of Allergy and Asthma Immunology 2008; 100:S18.
- 2 Long, WF., Taylor, RJ., Wagner, CJ., Leavengood, DC., Nelson, HS. Skin test suppression by antihistamines and the development of subsensitivity. *Journal of Allergy and Clinical Immunology* 1985; 76:113-7 (111).
- 3 Cook, TJ., MacQueen, DM., Wittig, HJ., Thornby, JI., Lantos, RL, Virtue, CM. Degree and duration of skin test suppression and side effects with antihistamines: a double blind controlled study with five antihistamines. *Journal of Allergy and Clinical Immunology* 1973; 51:7107. (111).
- 4 Phillips, MJ., Meyrick Thomas, RH., Moodley, I., Davies, RJ. A comparison of the in vivo effects of ketotifen, clemastine, chlorpheniramine and sodium cromoglycate on histamine and allergen induced weals in human skin. Br J Clin Pharmacol. 1983: 15:277-86. (11a).
- 5 Almind, M., Dirksen, A., Nielsen, NH., Svendsen, UG. Duration of the inhibitory activity on histamine-induced skin weals of sedative and non-sedative antihistamines. *Allergy*. 1988; 43:593-6 (111).
- 6 Simons, FE., Simons, KJ., Clinical pharmacology of new histamine H1 receptor antagonists. Clin Pharmacokinet. 1999; 15:277-86. (11a).
- 7 Pearlman, DS., Grossman, J., Meltzer, EO. Histamine skin test reactivity following single and multiple doses of azelastine nasal spray in patients with seasonal allergic rhinitis. *Ann Allergy Asthma Immunol.* 2003; 91:258-62. (1b).
- 8 Slottri, Zweiman B. A controlled study of the effects of corticosteroids on immediate skin test reactivity. *Journal of Allergy and Clinical Immunology* 1974; 54:292-225.
- 9 Des Roches, A., Paradis, L., Bougeard, Y.H. et al. Long-term oral corticosteroid therapy does not alter the results of immediate skin allergy prick tests. *Journal of Allergy and Clinical Immunology* 1996; 98(3):522-7.

- 10 Olson, R. et al. Skin reactivity to codeine and histamine during prolonged corticosteroid therapy. *Journal of Allergy and Clinical Immunology* 1990; 86:153-159.
- 11 Andersson M., Pipkorn U. Inhibition of the dermal immediate allergic reaction through prolonged treatment with topical glucocorticosteroids. J Allergy Clin Immunol, 1987:79:345
- 12 Gradman J., Wolthers OD. Suppressive effects of topical mometasone furoate and tacrolimus on skin prick testing in children. Pediatr Dermatol 2008; 25:26
- 13 Pipkorn U. Hammarlund A., Enerback L. Prolonged treatment with topical glucocorticoids results in an inhibition of the allergen-induced wheal-and-flare response and a reduction in skin mast cell numbers and histamine content. Clin Esp Allergy 1989; 19:19.
- 14 Narasimha, S.K., Effective topical corticosteroid application frequency and histamine induced wheals. *International Journal of Dermatology* 2005; 44(5):425-427.
- 15 Shah, K.M. et al. Predicting which medicine classes interfere with allergy skin testing. Allergy and Asthma Proceedings 2010; 31:477-482.
- 16 Bousquet, J. et al. Practical guide to skin prick testing in allergy to aeroallergens. Allergy 2012; 67:18-24.
- 17 Duenas-Laita, A. et al. Successful treatment of chronic drug-resistant urticaria without alprazolam. *Journal of Allergy and Clinical Immunology* 2009; 123:504-505.
- 18 Kupczyk M., Kuprys I., Bochenska-Marciniak M., et al. Ranitidine (150 mg daily) inhibits wheal, flare, and itching reactions in skin-prick tests. Allergy Asthma Proc 2007; 28:711.
- 19 Miller, J. et al. Suppression of immediate skin tests by ranitidine. *Journal of Allergy and Clinical Immunology* 1989; 84:895-899.

Note: American Academy of Otolaryngic Allergy's (AAOA) Clinical Care Statements attempt to assist otolaryngic allergists by sharing summaries of recommended therapies and practices from current medical literature. They do not attempt to define a quality of care for legal malpractice proceedings. They should not be taken as recommending for or against a particular company's products. The Statements are not meant for patients to use in treating themselves or making decisions about their care. Advances constantly occur in medicine, and some advances will doubtless occur faster than these Statements can be updated. Otolaryngic allergists will want to keep abreast of the most recent medical literature in deciding the best course for treating their patients.



Medicines to Avoid Before Skin Allergy Testing (continued)

Beta blockers are a risk factor for more serious and treatment resistant anaphylaxis, making the use of beta blockers a relative contraindication to inhalant skin testing.

Treatment with omalizumab (anti-IgE antibody) can suppress skin reactivity for up to six months.^{20, 21} No data exists for other biologic agents.

Topical calcineurin inhibitors have a variable affect. Pimecrolimus²² did not affect histamine testing but tacrolimus¹² did.

Herbal products have the potential to affect skin prick testing. In the most comprehensive study, ²³ using a single dose crossover study, it was felt that common herbal products did not significantly affect the histamine skin response. However, complementary and other alternative medicines do sometimes have a significant histamine response ²⁴ and included butterbur, stinging nettle, citrus unshiu powder, lycopus lucidus, Spirulina, cellulose powder, traditional Chinese medicine, Indian ayurvedic medicine.

Leukotriene receptor antagonist did not affect skin testing. ^{25, 26, 27}

Selective serotonin reuptake inhibitors (SSRIs) do not affect skin testing. 15, 28

Selective norepinephrine reuptake inhibitors (SNRIs) and protein pump inhibitors (PPIs) are felt not to need to be discontinued.¹⁵

Cyclosporin did not affect skin histamine response.²⁹

ACE inhibitors did not affect skin histamine response.30

Healthcare providers should take into consideration that many of these studies are done when the patient is taking one pharmaceutical agent for a short time. It is unclear, if a patient is taking multiple pharmaceutical/herbal agents that alone have a minor effect, whether the combination of these drugs could suppress the histamine response. Therefore, it is imperative that the provider have a positive skin histamine response before proceeding with diagnostic skin testing.

This is not a comprehensive list of medications that might affect skin testing. Physicians are expected to use their clinical judgment for other medications.

- 20 Noga O., Hanf G., Kunkel G. Immunological and clinical changes in allergic asthmatics following treatment with omalizumab. *Int Arch Allergy Immunol* 2003; 131:46
- 21 Corren J., Shapiro G., Reimann J., et al. Allergen skin tests and free IgE levels during reduction and cessation of omalizumab therapy. J Allergy Clin Immunol 2008; 121:506.
- 22 Spergel JM, Nurse N., Taylor P., PameixSpake A. Effect of topical pimecrolimus on epicutaneous skin testing. J Allergy Clin Immunol 2004; 114:695.
- 23 More, D.R., et al. Herbal supplements and skin testing. Allergy 2003; 58:492-494.
- 24 Mainardi, T. et al. *Journal of Allergy and Clinical Immunology* February 2009; 123(2).
- 25 Hill, S.L., Krouse, J.H. The effects of montelukast on intradermal wheal and flare. Otolaryngology Head and Neck Surgery 2003; 129(3):199-203.

- 26 Simons FE, Johnston L., Gu X., Simons KJ. Suppression of the early and late cutaneous allergic responses using fexofenadine and montelukast. *Ann Allergy Asthma Immunol* 2001: 86:44
- 27 Cudhadaroglu C., Erelel M., Kiyan E., et al. Role of Zafirlukast on skin prick test. *Allergol Immunopathol (Madr)* 2001; 29:66.
- 28 Isik SR, Celikel S., Karakaya G., et al. The effects of antidepressants on the results of skin prick tests used in the diagnosis of allergic diseases. Int Arch Allergy Immunol 2011; 154:63.
- 29 Munro CS, Higgins EM, Marks JM, et al. Cyclosporin A in atopic dermatitis: therapeutic response is dissociated from effects on allergic reactions. Br J Dermatol 1991: 124:43.
- 30 Joint Task Force on Practice Parameters, American Academy of Allergy, Asthma and Immunology, American College of Allergy, Asthma and Immunology, Joint Council of Allergy, Asthma and Immunology. The diagnosis and management of anaphylaxis: an updated practice parameter. J Allergy Clin Immunol 2015; 115:341

Note: American Academy of Otolaryngic Allergy's (AAOA) Clinical Care Statements attempt to assist otolaryngic allergists by sharing summaries of recommended therapies and practices from current medical literature. They do not attempt to define a quality of care for legal malpractice proceedings. They should not be taken as recommending for or against a particular company's products. The Statements are not meant for patients to use in treating themselves or making decisions about their care. Advances constantly occur in medicine, and some advances will doubtless occur faster than these Statements can be updated. Otolaryngic allergists will want to keep abreast of the most recent medical literature in deciding the best course for treating their patients.

Medicines to Avoid Before Skin Allergy Testing (continued)



Suppressant Effects of Drugs on Immediate Skin Tests*		
Medications	Mean Days Suppressed	Max Days Suppressed
First Generation Antihistamines	2	5
Second Generation Antihistamines	2	7
Antihistamine Nasal Sprays	0	1
Antihistamine Eye Drops	0	1
Tricyclic Antidepressants and Tranquilizers		14
Histamine2 Antihistamines (H2 Blocker)	0	2
Topical Corticosteroids		Up to 21

Medications that DO NOT Need to be Stopped Prior to Allergy Skin Prick Testing* Angiotensin-Converting Enzyme (ACE) Inhibitors Benazepril Captopril Enalapril Lisinopril Perindopril Quinapril Ramipril Immunosuppressant Cyclosporin **Nasal Steroid Sprays Beclomethasone Dipropionate Nasal Budesonide Nasal** Ciclesonide Nasal Fluticasone Propionate Fluticasone Furoate Nasal **Mometasone Furoate Nasal** Oxymetazoline **Triamcinolone Acetonide** Norepinephrine Reuptake Inhibitors (SNRIs) Duloxetine Venlafaxine **Protein Pump Inhibitors (PPIs)** Esomeprazole Lansoprazole Omeprazole **Pantoprazole** Rabeprazole Serotonin Reuptake Inhibitors (SNRIs) Citalopram Escitalopram Fluoxetine **Paroxetine** Sertraline

Note: American Academy of Otolaryngic Allergy's (AAOA) Clinical Care Statements attempt to assist otolaryngic allergists by sharing summaries of recommended therapies and practices from current medical literature. They do not attempt to define a quality of care for legal malpractice proceedings. They should not be taken as recommending for or against a particular company's products. The Statements are not meant for patients to use in treating themselves or making decisions about their care. Advances constantly occur in medicine, and some advances will doubtless occur faster than these Statements can be updated. Otolaryngic allergists will want to keep abreast of the most recent medical literature in deciding the best course for treating their patients.

^{1 *}Some exceptions—see prior references