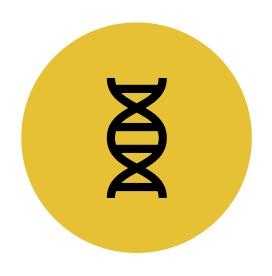


Mast Cell & Allergy Support



Researched Nutritionals® 800 755 3402 www.ResearchedNutritionals.com

Histamine: Primary Mediator of Allergic Response



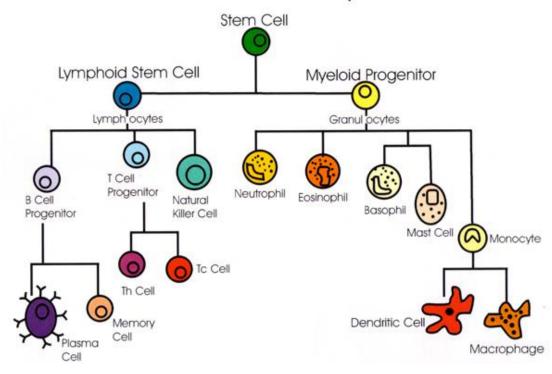


IMMUNE MESSENGER MOLECULE RELEASED FROM MAST CELLS

NEUROTRANSMITTER

Immune Cells

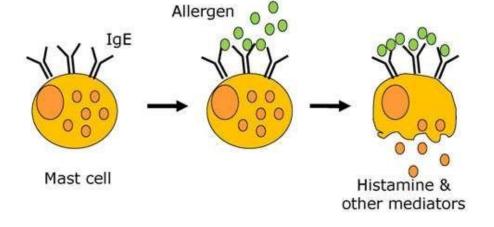
Cells of the Immune System



^{*}These statements have not been evaluated by the Food and Drug Administration. These products are not intended to diagnose, treat, cure, or prevent any disease.

Histamine Release from Mast Cells

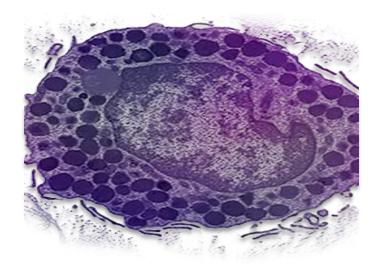
- Histamine Causes:
 - Blood vessel dilation
 - Bronchoconstriction
 - Increased heartbeat
 - Release of adrenaline
 - Increased capillary permeability
 - Increased gastric acid secretion
 - Pruritis



HistaQuel® Comprehensive Allergy Support

Two-Fold Mechanism of Action

- 1. Reduce Histamine release from Mast cells*
- 2. Block Histamine from binding to receptors via competitive inhibition*



HistaQuel®

Supplement Facts Serving Size: 2 Capsules Servings Per Container: 60 Amount Per Serving %Daily Value** Mast Cell Secure™ 375 mg † Quercefit™, Perilla Frutescens (seed) Extract, Luteolin, Fisetin HistaCheck™ 200 mg † Black Cumin (seed) Extract, Stinging Nettle (leaf) Extract ** Percent Daily Values are based on a 2,000 calorie diet. † Daily Value not established.

Other Ingredients: Organic rice hulls, hypromellose (vegetable capsule), I-leucine.

Manufactured without milk, eggs, fish, crustacean shellfish, tree nuts, peanuts, wheat, soy and gluten. Produced in a cGMP facility that may process other ingredients containing these allergens.

Suggested use: Take two capsules twice per day with or without meals.



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HistaQuel® Ingredient Blends

Ingredient Blend	Constituents	Mechanisms of Action*
Mast Cell Secure™ (Flavonoids)	 Fisetin Luteolin Perilla Frutescens QuercefitTM (Up to 20-fold more bioavailable of regular Quercetin) 	 Targeted flavonoid blend supports a decrease in mast cell release of histamine Moderates interactions between mast cells and activated T cells. Decreases Th2 release of allergy promoting cytokines IL-4 and IL-5
HistaCheck™	 Stinging Nettle leaf Black Cumin Seed Extract 	 Decreases ability of histamine to bind to its receptors by competitive inhibition Supports management of a healthy immune response to allergens

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HistaQuel[®] Mast Cell Secure™ Blend Ingredient Research

Ingredient research indicates...

Flavonoids (general)

- Inhibit IgE mediated histamine release¹
- Decrease production of proinflammatory cytokines¹
- Down-regulate mast cell activation¹

Fisetin

- Significantly inhibited infiltration of inflammatory cells including eosinophils, mast cells, CD4(+) T cells and CD8(+) T cells²
- Total serum immunoglobulin E (IgE) levels were markedly reduced by Fisetin²
- Reduced the production of interferon-gamma and interleukin-4 by activated CD4(+) T cells in a dose-dependent manner²
- Limits interactions between mast cells and activated T cells³

Quercefit™

Up to 20-fold more bioavailable of regular Quercetin⁴

HistaQuel[®] HistaCheck™ Blend Ingredient Research

Ingredient research indicates...

Perilla Frutescens

- Suppresses IgE-mediated type I hypersensitivity reactions⁷
- Significantly inhibits IgE-mediated histamine release⁷

Nettle Extract

- Shows in vitro inhibition of several key inflammatory events that cause the symptoms of seasonal allergies⁵
- Antagonist against the Histamine-1 (H(1)) receptor⁵
- Inhibition of mast cell tryptase⁵
- Inhibits prostaglandin formation⁵

Black Cumin Seed Extract

- Gastroprotective: inhibit acid secretion and block histamine receptor⁶
- Anti-Inflammatory effects in asthma⁶
- Inhibits mast cell histamine release and decreases mast cell numbers⁶

HistaQuel®

- Flavonoid blend to support the body's ability to regulate healthy histamine levels*
- Herbs to supports healthy mast cell function*
- GMO-Free
- Vegetarian



References

- 1. Flavonoids inhibit histamine release and expression of proinflammatory cytokines in mast cells. Park HH et al. <u>Arch Pharm Res.</u> 2008 Oct;31(10):1303-11.
- 2. Immunosuppressive effects of Fisetin against dinitrofluorobenzene-induced atopic dermatitis-like symptoms in NC/Nga mice. <u>Kim GD</u> . <u>Food Chem Toxi Toxicol</u>. 2014 Apr;66:341-9.
- 3. The hydroxyflavone, Fisetin, suppresses mast cell activation induced by interaction with activated T cell membranes. Nagai K. et al. *British Journal of Pharmacology*. 2009.
- 4. Improved oral absorption of quercetin from Quercetin Phytosome®, a new delivery system based on food grade lecithin. Riva A. et al. *Int. Jr. of Drug Metabolism & Pharmacokinetics*. 2018.
- 5. Nettle extract (Urtica dioica) affects key receptors and enzymes associated with allergic rhinitis. Roschek B. et al. *Phytother. Research*. 2009 Jul;23(7):920-6.
- 6. Gastroprotective Effect of an Aqueous Suspension of Black Cumin *Nigella Sativa* on Necrotizing Agents-Induced Gastric Injury in Experimental Animals *Saudi J Gastroenterol 2008 Jul;14(3):128-34. doi: 10.4103/1319-3767.41731*
- 7. A flavanone derivative from the Asian medicinal herb (Perilla frutescens) potently suppresses IgE-mediated immediate hypersensitivity reactions. Kamei R. et al. *Biochem Biophys Res. Commun*. 2017 Jan 29;483(1):674-679.



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