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A COMPREHENSIVE REVIEW OF ALLERGIC DISEASES: PATHOPHYSIOLOGY, DIAGNOSIS, AND TREATMENT OPTIONS

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ABSTRACT

Allergic reactions are a complex and frequently misunderstood aspect of the immune system's response to external stimuli. These abstract aims to succinctly describe allergic reactions, focusing on their underlying mechanisms, common triggers, and management strategies. The primary components in allergic responses include immunoglobulin E (IgE) antibodies, mast cells, and basophils. When an allergic individual encounters an allergen, these antibodies prompt the release of inflammatory mediators, such as histamine, resulting in symptoms like itching, swelling, hives, and, in severe cases, anaphylaxis. Allergic reactions are intricate immune responses that can present in various forms and intensities

Keywords: Allergen, Anaphylaxis, Food Allergy, Atopic Dermatitis.

INTRODUCTION

An allergic reaction occurs when the immune system responds to a typically harmless substance as if it were a threat. This substance, called an allergen, can provoke the release of various chemicals in the body, causing a range of symptoms. Normally, the immune system defends against harmful invaders like bacteria and viruses. However, in allergies, the immune system mistakenly identifies certain substances as harmful and initiates a defense against them.

Immunoglobulin E (IgE) antibodies, produced by the immune system, are crucial in allergic responses. When an allergic individual encounters an allergen, their immune system produces IgE antibodies specific to that allergen. These antibodies then attach to specialized cells called mast cells and basophils, found in many tissues throughout the body, especially in areas prone to exposure like the skin, lungs, and digestive tract [1].

Upon subsequent exposure to the same allergen, it binds to IgE antibodies on the surfaces of mast cells and basophils. This interaction prompts these cells to release potent compounds such as histamine. Histamine and other mediators lead to the characteristic symptoms of an allergic reaction. These symptoms can affect various body systems, and the severity of the reaction can differ significantly between individuals.



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Signs and Symptoms:

Allergens are peptide molecules found in various substances and can affect multiple organ systems, including the circulatory, cardiac, gastrointestinal, and respiratory systems. The severity and speed of sensitization to allergens can lead to a range of symptoms such as swelling (edema), skin reactions (cutaneous responses), low blood pressure (hypotension), bronchoconstriction, fainting, and even coma [2].

Anaphylaxis is a rapid, life-threatening hypersensitive reaction that can be fatal if not treated immediately. Allergens like latex can cause skin rashes and irritation, leading to conditions such as angioedema, contact dermatitis, and other severe cutaneous and systemic effects. These allergens can be inhaled, ingested, or come into contact with the skin.

Many allergens, such as dust particles, are microscopic and airborne, making them easily inhaled and capable of causing symptoms in exposed organs like the nasal passages, lungs, and eyes. Common symptoms include mucosal irritation, a runny nose, and sneezing, often associated with allergic rhinitis (hay fever) [3].



Figure No 1: Allergy Reaction Symptoms

Common allergic reactions include:

- 1. Respiratory Symptoms:
- Sneezing
- Runny or stuffy nose
- Itchy or watery eyes

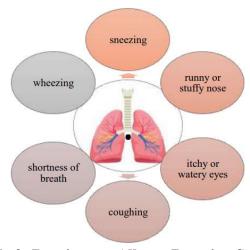


Figure No 2: Respiratory Allergy Reaction Symptoms



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Skin Symptoms:

- 1. Itching
- 2. Hives (raised, red, itchy welts on the skin)
- 3. Eczema or dermatitis

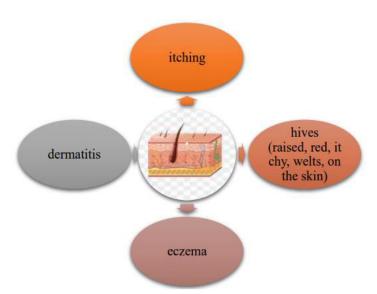


Figure No 3: Skin Allergy Reaction Symptoms

Bacterial Infections in Allergy

The relationship between bacterial infections and allergies is intricate, with various factors influencing their interaction. Both bacterial infections and allergic reactions involve the immune system, and their interplay can affect the development and severity of allergic conditions. Here's an in-depth look at this connection [4]:

Immune System Modulation:

a. Toll-Like Receptors (TLRs):

Bacterial infections trigger the immune system through pattern recognition receptors, such as Toll-like receptors (TLRs). When TLRs are activated, they initiate an inflammatory response that can affect the immune pathways involved in allergic reactions.

Bacterial Infections in Allergy (Continued)

b. Th1/Th2 Balance:

Bacterial infections typically stimulate a Th1 response, associated with cell-mediated immunity. This Th1 response may counterbalance the Th2 response involved in allergic reactions, potentially influencing the development and severity of allergic conditions [5].

Early-Life Exposure:

a. Hygiene Hypothesis:

The hygiene hypothesis suggests that reduced exposure to infections, including bacterial infections, during early life may increase the risk of developing allergies. Lack of exposure to a variety of microbes, including bacteria, can affect the proper development of the immune system.



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Bacterial Infections in Allergy (Continued)

Role of Microbiota:

a. Gut Microbiota:

The gut microbiota plays a crucial role in modulating the immune system. Imbalances in the gut microbiota, often influenced by factors like antibiotic use, may increase the risk of allergic conditions.

b. Microbiota and Immune Tolerance:

A healthy and diverse microbiota is associated with immune tolerance, helping to prevent inappropriate allergic responses. Disruptions in the microbiota composition can lead to a breakdown in immune tolerance and an increased susceptibility to allergies.

Atopic Diseases:

a. Eczema (Atopic Dermatitis):

Bacterial skin infections can exacerbate symptoms of atopic dermatitis, a common allergic skin condition. Itchiness, redness, and inflammation may worsen during bacterial infections.

b. Respiratory Infections and Asthma:

Severe bacterial respiratory infections, particularly in early childhood, can influence the development of asthma. Recurrent or severe infections may contribute to airway inflammation and hyperreactivity.

Protective Effects:

a. Microbial Exposure and Allergy Protection:

Research indicates that early exposure to microorganisms, such as bacteria, may reduce the risk of developing allergies. This exposure is believed to help train the immune system to handle a wide range of environmental stressors effectively.

Bacterial Infections in Allergy (Continued)

Management Strategies:

a. Antibiotic Use and Microbiota Disruption:

Frequent or prolonged use of antibiotics can disrupt the microbial balance. This disruption may affect immune regulation and increase the risk of allergic conditions.

b. Probiotics and Prebiotics:

Modulating the gut microbiota through probiotics (beneficial bacteria) and prebiotics (substances that promote the growth of beneficial bacteria) is a promising area of research for potential strategies to prevent or manage allergies.

Vaccination:

a. Preventive Measures:

Vaccination against certain bacterial infections can help prevent these infections and may indirectly influence the immune response, potentially impacting the development of allergic conditions.



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Bacterial Infections in Allergy (Continued)

Environmental Exposures:

a. Endotoxin Exposure:

Exposure to bacterial endotoxins, components of bacterial cell walls, has been linked to a reduced risk of allergies. This exposure is more common in environments with higher microbial diversity, such as farms.

In conclusion, the interaction between bacterial infections and allergies is complex and influenced by factors like the type of bacteria, individual susceptibility, and the timing of exposures. While bacterial infections can affect immune responses and potentially modulate allergic conditions, further research is necessary to fully understand this relationship and to develop targeted preventive and therapeutic strategies [6].

Treatments

Advances in allergy research have significantly impacted the treatment of mild to severe allergic disorders. Various medications are available for specific symptoms of allergic conditions, including pharmaceuticals that effectively manage and treat atopic symptoms. Epinephrine injections for anaphylaxis can be carried by individuals, while anti-allergic and antihistamine drugs are commonly used to alleviate adverse effects.

The treatment of allergic diseases in children follows a similar approach to that in adults. Treatment options include allergen avoidance through environmental control, pharmacotherapy, and immunotherapy. The primary goal of treatment is to control symptoms without affecting the child's daily functioning. An equally important goal is to prevent the progression of allergic diseases [7]. For children at high risk of developing allergies, early implementation of dietary and environmental control measures can reduce sensitization, and timely recognition and treatment of allergic symptoms are crucial.

Various allergen-based diagnostic and treatment strategies have been developed to address allergic reactions. The following drugs are commonly used to treat allergic conditions:

- Epinephrine injections for anaphylaxis.
- Anti-allergic drugs to manage symptoms.
- Antihistamines to reduce adverse effects.

In conclusion, the ongoing development of diagnostic and therapeutic strategies continues to improve the management of allergic disorders, offering better outcomes for both children and adults.

1. Allergen Avoidance

The cornerstone of allergy treatment always revolves around meticulously avoiding specific allergens that trigger allergic reactions. The most crucial and effective strategy to mitigate allergic responses in sensitive individuals is strict allergen avoidance. For food allergies and certain insect sting allergies, avoidance remains the primary treatment approach, which can be highly effective with proactive measures and preparedness[8].

However, airborne allergens that are difficult to control or detect pose challenges to avoidance strategies. In such cases where avoidance is impractical, alternative strategies are necessary to navigate these obstacles effectively.



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2. Pharmacotherapy

Pharmacotherapy becomes essential when allergen avoidance is not feasible, and the development of sensitivity is unavoidable. Numerous medications have been developed to antagonize and mitigate the actions of allergic mediators. Antileukotrienes and antihistamines are common examples that prevent the onset of allergic symptoms and suppress inflammatory mediators. FDA-approved medications include adrenaline (epinephrine), antihistamines, glucocorticoids, and theophylline, all of which serve as anti-inflammatory agents[9].

Additionally, decongestants, mast cell stabilizers, and eosinophil chemotactic agents such as zafirlukast (Accolate) or montelukast (Singulair) are frequently prescribed medications to diagnose and prevent chronic and severe allergic conditions.

3. Immune Cell Therapy

Allergen-specific immunotherapy involves gradually increasing allergen doses to achieve immunological and clinical tolerance. Immunotherapy induces T-cell tolerance through various mechanisms, including modulation of cytokine levels, suppression of allergen-specific T-cell proliferation, promotion of apoptosis, and generation of regulatory T cells. These processes reduce inflammatory mediators and cells within affected tissues, promote the production of blocking antibodies, and induce IgE desensitization [10].

Studies have demonstrated the effectiveness of this type of therapy, showing long-term benefits in preventing the progression of atopy. A recent advancement in immunotherapy includes intravenous administration of monoclonal anti-IgE antibodies, which bind to both B-cell-associated and free IgE, thereby neutralizing and eliminating them. Sublingual immunotherapy offers an oral treatment approach that relies on inducing local immune tolerance to non-pathogenic substances like environmental microbes and food allergens.

Allergy shot therapy, or allergen immunotherapy, may emerge as a prominent treatment method in the future. It requires rigorous monitoring and long-term commitment to achieve optimal individualized therapy outcomes.

CONCULISION

Allergies: A Global Concern; Allergies pose a significant challenge affecting millions worldwide. Identifying and avoiding rare or elusive allergens can be daunting, making symptom management crucial for allergic patients. Current diagnostic methods and treatment strategies aim to alleviate symptoms, yet long-term relief remains elusive. Researchers are exploring novel theories and testing methodologies to advance allergy treatments. Progress in biochemical, proteomic, and genomic techniques has provided new insights into allergies and sensitivities. A major challenge in allergy bioinformatics is the analysis and integration of vast data sets. Developing bioinformatics tools for B-cell and T-cell epitope prediction is essential for understanding allergic responses comprehensively. Resources and databases that consolidate information from diverse sources are critical for advancing allergenicity prediction and understanding allergen cross-reactivity. Recent advancements focus on antibody-specific epitope prediction algorithms, crucial for improving diagnostic and



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therapeutic methods for allergic diseases. The application of IgE-binding epitope prediction algorithms is pivotal for developing more effective allergy immunotherapy (AIT) strategies, recognized as precise or personalized medicine. Bioinformatics-driven breakthroughs in AIT approaches and allergen bioinformatics promise to enhance our understanding of allergic disorders and inspire further research in the field.

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