Aspirin Desensitization for Aspirin-Exacerbated Respiratory Disease (AERD)

Nkiruka Erekosima, MD MPH



AERD - Description

- Also known as Samter's Triad
- Refers to the combination of:
 - Asthma
 - Chronic rhinosinusitis (CRS) with nasal polyposis (NP)
 - Acute reactions to aspirin and other COX-1 inhibiting NSAIDs
 - Causing upper and lower respiratory symptoms
- Prevalence:
 - 7% among patients with asthma
 - 15% among patients with severe asthma
 - 9-10% among patients with NP or CRS
 - Up to 30% of asthmatics with NP

- Pathophysiology: Not fully understood but involves dysregulation in arachidonic acid metabolism
 - Overproduction of Leukotrienes



Source: Trevor AJ, Katzung BG, Masters SB: Pharmacology Examination & Board Review, 9th Edition: www.accesspharmacy.com Copyright © The McGraw-Hill Companies, Inc. All rights reserved.

AERD - Clinical Features

- <u>Chronic upper and lower respiratory disease (CRS w/ NP, Asthma)</u> and <u>reaction to NSAIDs is an exacerbating factor</u>.
- Worsening of asthma and chronic sinusitis occurs even with NSAID avoidance
- Usually diagnosed in adulthood
- The 3 components tend to develop serially over years
- Acute exacerbation after ingestion of NSAID typically begins 30 minutes to 3 hours after ingestion
 - Dose-related: small doses produce milder symptoms; large doses produce more severe symptoms
 - Urticaria and/or angioedema occur in 15% of AERD patients.
 - (Similar symptoms may occur with high doses of acetaminophen)

AERD - Diagnosis & Management

- **Diagnosis** made clinically; may require a diagnostic aspirin challenge
- Management: Medications, Surgery (polypectomy), Aspirin Desensitization
 - Leukotriene modifying agents (montelukast, zafirlukast, zileuton)
 - Protect patients from severe exacerbations due to accidental NSAID exposure
 - NSAID avoidance (unless patient has been desensitized to aspirin)
- Aspirin Desensitization: Induces one's ability to temporarily tolerate aspirin. After successful desensitization, the patient must continue aspirin/NSAID daily in order to maintain the desensitized state.
 - Indications:
 - Nasal polyposis that worsen or recur after surgery despite optimal medical management
 - Other conditions requiring daily use of NSAIDs (e.g. CAD, arthritis)
 - Reduces the expression of leukotrienes
 - Successful in majority of AERD patients

Aspirin Desensitization

• If sinus surgery/polypectomy is required, schedule desensitization 2-4 weeks after surgery.

Within 1-2 weeks prior to desensitization:

- FEV1 > 70% predicted (optimally)
- Start or continue leukotriene modifier and other asthma medications
- Systemic steroid burst if necessary

Day of Procedure:

- Obtain informed consent, baseline vital signs and spirometry
- Vital Signs, FEV1 (or peak flow) and clinical assessment prior to each dose and with symptoms
- Treatment of reactions (ocular, nasal, laryngeal, bronchial, cutaneous): antihistamines, albuterol, epinephrine
- Lower resp. tract reaction: 15% decrease in the FEV1 from baseline FEV1
- After successful desensitization, start daily aspirin 650 mg twice daily
 - Taper over 6 months to 325 mg twice daily
- Subsequent desensitization is performed if aspirin is missed for more than 48 hours

Protocol

Time	Aspirin dose
0	20.25 mg
1.5 hours	40.5 mg
3.5 hours	81 mg
5.5 hours	162 mg
7.5 hours	325 mg

- Most take 2 days. May adjust doses and extend dosing interval based on patient history and assessment.

- If reaction occurs, observe patient for at least 3 hours.

- When stabilized, repeat provoking dose; escalate if tolerated. (Provoking dose is usually 81 mg.)

AERD - Summary

- AERD is a distinct and important subtype of asthma and polypoid sinus disease.
- Early identification is important given the increased morbidity and costs associated with asthma exacerbations and recurrent sinus surgeries.
- Benefits of Aspirin Desensitization followed by daily aspirin in AERD:
 - Decrease in nasal polyp formation, nasal congestion, sinus infections, sinus surgery, and need for systemic corticosteroids
 - Increase in asthma control and sense of smell
 - Effects on CRS symptoms are more dramatic than effects on asthma

References

Solensky, et al. Drug Allergy: an updated practice parameter. Ann Allergy Asthma Immunol 2010;105:259-273.

Macy, et al. Aspirin challenge and desensitization for aspirin-exacerbated respiratory disease: a practice paper. *Ann Allergy Asthma Immunol* 2007;98:172–174. White AA, Stevenson DD. Aspirin desensitization in aspirin-exacerbated respiratory disease. *Immunol Allergy Clin North Am* 2013;33(2):211-22.

Contact information:

T: 301-879-7700 | www.AlphaAllergy.com

Email: Info@AlphaAllergy.com

