Management of Chronic Cough

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1. Understand the common etiologies of chronic cough

2. Understand the Otolaryngologist's role in the treatment of chronic cough

3. Learn treatment options for sensory neuropathic chronic cough



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STATES STATES



Demographics

- Most Common Presenting Complaint for Adults seen in an ambulatory setting
- 2001-2002 US Ambulatory Care Visits

 4.3% of patients reported cough as symptom
 > 33,000,000 visits

Schappert SM, Burt CW. Vital Health Stat. 2006;13:1-66.



Neuropathogenesis

- Initiation: Reflex initiated through sensory nerve input in airway → Vagus Nerve
- 2) Processing: Complex network of central nerves
- 3) Stimulation: Efferent limb and motor output to generate cough



Phases of Cough

- Inspiratory Phase
- Compressive Phase
- Expiratory Phase

Simpson CB, Amin MR. Otolaryngology Head Neck Surg 2006;134:693-700.



Definitions

- Acute Cough: < 3 weeks
- Subacute Cough: 3-8 weeks
- Chronic Cough: > 8 weeks

Irwin RS. *NEJM* 343(23): 1715-1721,2000 Irwin RS. *Chest* 1998; 114(suppl1) :133S-181S



Common Causes of Cough

- Rhinosinusitis (Upper Airway Cough Syndrome)
- GERD
- Pulmonary
 - Asthma, Bronchiectasis
 - Non-asthmatic Eosinophilic Bronchitis



Guidelines Writing Committee. Chest 2006; 129 (Suppl. 1): 1S-292S

Pathogenic Triad of Cough



- normal CXR
- no ACE-I

Palombini BC et al. Chest 1999;116:279-84. Pratter et al. *Chest* 2006; 129 (Suppl. 1): 1S-292S.



Other Causes of Chronic Cough

- Behavioral
- Bordetella Pertussis
- Medication related
 - Angiotensin Converting Enzyme-Inhibitors
- Sensory Neuropathic
- Chronic Aspiration
- Vocal Fold Paresis



Management Strategies

- Multidisciplinary
- Multifactorial
- Ability to communicate/collaborate with Gastroenterologist, Pulmonologist
- Target underlying condition(s) of the cough rather than symptom of cough
 - Anti-tussives: Benzonatate (Tessalon),
 Dextromethorphan (Robitussin, Theraflu),
 Guaifenesin, Narcotics



Multifactorial

Percentage of Cases Presenting 1,2,3, and 4 Causative Factors



Palombini BC et al. Chest 1999;116:279-84.



Ruling Out the Obvious: ACE-Inhibitor

- 10-33% incidence for patients on ACE-I
- ACE-I related cough can present at any time after being on the ACE-I
- Discontinue ACE-I
- Symptoms should resolve or improve within 4 weeks



Addressing the major causes: Rhinosinusitis

- Upper Airway Cough Syndrome
- "Post Nasal Drip Syndrome"
- Nonspecific symptoms and signs
- Lack of objective testing
- Pathophysiology
 - mechanical stimulation of afferent limb of cough reflex in upper airway
 - Increased sensitivity of cough reflex in upper airway

Palombini BC et al. Chest 1999;116:279-84. Pratter et al. *Chest* 2006; 129 (Suppl. 1): 1S-292S.



Management of Rhinosinusitis & Allergic Rhinitis

- Avoidance
 - Allergy Testing & Desensitization
- Reduction of Inflammation/Secretions
 - Antihistamines
 - Steroid sprays
 - Anticholinergics
- Treatment of Infection
- Surgical Correction



Addressing the major causes: GERD

- Pathophysiology
 - Acid exposure in distal esophagus stimulating esophageal-tracheobronchial cough reflex (via vagus)
 - Microaspiration of esophageal contents into larynx and tracheobronchial tree

Irwin RS et al. Am Rev Respir Dis 1989;140:1295-1300. Simpson CB, Amin MR. Otolaryngol Head Neck Surg 2006;134:693-700.



Diagnosis and Treatment of GERDassociated cough

- Heartburn
- Diet related
- Throat clear, am or intermittent hoarseness, globus sensation
- Testing
 - pH Probe
 - Empiric Therapy & Low-acid Diet
 - BID PPI at 40mg for 3 months



Addressing the major causes: Pulmonary Causes

- CXR & Pulmonary Referral
- COPD, Asthma
 - Pulmonary Function Tests
- Eosinophilic Bronchitis
 - Empiric Corticosteroids
 - Sputum Test for increased Eosinophils



Pratter MR et al. Chest 2006;129:S222-31.

Other Causes: Sensory Neuropathic

- Hypersensitivity due to reduced cough threshold in response to irritative stimuli
- Sustained vagal injury
 - Postviral vagal neuropathy
- Triggers: talking, temperature change, yawning
- May accompany a motor neuropathy – Vocal Fold Paralysis/Paresis
- Does NOT wake them from sleep



Management of Sensory Neuropathic Cough

- Lower Sensory Threshold
- Can Take 3-6 weeks to have an effect
- Attempt taper after 3-6 months



Sensory Neuropathic Cough: Dosing

- Increase dosing until symptoms improve/resolve or side effects become overwhelming
- Gabapentin (Lee & Woo Ann ORL 2005;114:253-7)
 Start 100 tid increase to 300 tid
- Amitriptyline (Bastian RW OHNS 2006;135:17-21)
 - Start 10mg qhs increase to 60-100 mg qhs
- Pregabalin (Halum SL Laryngoscope 2009;119:1844-7)

Tramadol Dosing

- 25-50mg bid-tid prn
- Mechanism of Action: Central Nervous System Opiate activation and serotonin/norepinephrine inhibition
- Dion et al.
 - Prospective Trial of 16 patients
 - Cough Severity Index (23 \rightarrow 14, p=0.003)
 - Leicester Cough Questionnaire (74 → 103, p=0.005)
 - Follow-up: 15 1029 days

Dion GR, Teng SE, Achlatis E, Fang Y, Amin MR. Treatment of Neurogenic Cough with Tramadol: A Pilot Study. Otolaryngol Head Neck Surg. 2017;157:77-9.



Side Effects

- Reversible with discontinuation of medication
- Amitriptyline: somnolence, weight gain, postural hypotension, dry mouth, arrhythmias
- Gabapentin: somnolence, dizziness, rash, weakness, nausea, tremor, nightmares, blurred vision, leukopenia
- Pregabalin: somnolence, difficulty thinking clearly, dry mouth, peripheral edema, weight gain, dizziness
- Tramadol: somnolence, serotonin syndrome, dependence





Morrison M et al. J Voice 1999;13:447-55.



64F with Dyspneic Episodes x 15 years

- Difficulty on Inspiration
- Denies Dysphonia & Cough
- Began when exposed to pollution in China
- Triggers: Perfume, Cleaning Products, Fragrant Flowers, Hot Air, Exertion
- Diagnosed with Asthma: Significant Response to Bronchodilators



Laryngoscopy before and after exposure to Purell









TW 57F w/cough x 2 years



Behavioral Therapy to Address Cough

Behavioral Therapy

- Speech Language Pathologist
- Utilization of voice therapy techniques
- Identification of Triggers (Cough Diary)
- Alternative Compensatory Techniques
 - Drink Water
 - Slow breathing against pursed lips

Other Causes: Bordetella Pertussis

- Severe coughing fits that develop after URI
- Early stage Culture or PCR
- Late stage Serum IgG
 40% (19/48) tested positive
- Treatment: Anti-tussives, supportive
- Cough will gradually improve over months

Bock JM et al. Otolaryngol Head Neck Surg 2012;146:63-7.

Other Causes: Vocal Fold Paresis 44F w/ 2 year history of chronic cough

- PFTs: No Asthma
- Codeine helps
- No GERD/LPR symptoms
- 6 week trial of Amitriptyline unsuccessful
- Decreased vocal projection which does not bother her

Sulica L. Injection Augmentation for Chronic Cough. Fall Voice Conference, Oct 18, 2013, Atlanta,

44F c Chronic Cough: Stroboscopy

Injection Laryngoplasty

JL: 55M w/Cough x 10 years

- Occur 20 times per day, last seconds
- At times can be severe → Passing out
 Fell and struck his head 2 days ago
 Lost consciousness while driving once
- Alleviating Factors: Tussionex
- Aggravating: Cigarette Smoke, Perfumes, Gas, Lysol

JL: Past Medical History

- PMHx:
 - Asthma
 - GERD
- PSHx:
 - C5-7 cervical fusion
 - Nissen Fundoplication

- OSA
- Hypertension
- UPPP
- Septoplasty

JL Laryngoscopy

Treatment Plan

2-Month Follow up

- 3 visits with local SLP No effect
- Titrated Gabapentin to 300mg tid and became irritable so stopped
- Unable to tolerate Injection Laryngoplasty

Treatment Plan – Next Steps?

- Behavioral Therapy at Voice Center
- Pregabalin 150 mg PO bid

53F w/Cough x 3 years

- Productive
- Associated Symptoms: Facial Pressure, Headache, Rhinorrhea, PND
- Cough unresponsive to Antibiotics (but sinus symptoms resolved)
- Unresponsive to Tessalon, Allegra Singuilair
- On PPI, EGD showed improvement in Gastritis

53F w/Cough x 3 years

- Only regimen that resolved her cough is combination of
- Oral Steroids
- Budesonide Sinonasal Rinse bid

53F w/Cough Stroboscopy

53F w/Cough Maxillofacial CT

Treatment Plan?

Conclusions

- Otolaryngologist sees a small fraction of chronic cough
- Multi-Factorial
- Thorough Algorithm
 - Address Common Causes First
 - Develop Relationships with Pulmonary/GI
 - Address Sensory Neuropathic Cough
 - Consider Behavioral Therapy

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