TOOLS FOR PRACTICE #329 | December 12, 2022



Coughing up the evidence: Bronchodilators or inhaled steroids for post-infectious cough

CLINICAL QUESTION

Do bronchodilators or inhaled corticosteroids (ICS) improve post-infectious cough in adults without asthma?

BOTTOM LINE

Data is very limited with only two ICS randomized controlled trials (RCTs) (163 patients) and one bronchodilator RCT (92 patients). For adults, post-infectious cough scores may improve ~50% on placebo and ~5-10% more with ICS over 2 weeks. Ipratropium/salbutamol may resolve cough in more patients than placebo at day 10 (69% versus 37%) but most patients (>80%, regardless of treatment) will have cough resolution by day 20.

EVIDENCE

- Differences statistically significant unless noted.
- ICS versus placebo:
 - Four systematic reviews of treatments for persistent cough (4-9 RCTs, 335-750 patients):
 Interpretation limited by inclusion of RCTs with acute (<3 weeks)^{1,2} and chronic (>8 weeks)^{2,3} cough, and multiple drug classes.⁴

- Most useful systematic review² (2 RCTs, 163 patients), mostly subacute cough (3-8 weeks):
 - ICS 0.42 (standard mean difference) better than placebo. In clinical terms, placebo improved cough scores at ~2 weeks by ~50-56% and ICS improves ~2-13% more.
 - Largest RCT⁵ (133 patients) also reported:
 - Additional outcomes (days off work, nocturnal awakenings, adverse effects): No difference.
 - Proportion of non-smokers with >50% cough improvement: 81% versus 54% (placebo), number needed to treat (NNT)=4.
 - No improvement among smokers.
 - Limitations: Not all patients had post-infectious cough; study industry funded.
- Bronchodilators versus placebo:
 - One RCT⁶ of 92 patients (cough duration 3-4 weeks): Combination of nebulized salbutamol/ipratropium versus placebo.
 - Proportion with ongoing cough at day 10: 37% versus 69% placebo, NNT=3.
 - No difference at day 20 (both >80% resolved).
 - Limitations: Small studies; non-validated cough scores; multiple outcomes.

CONTEXT

- Post-infectious cough is a cough persisting 3-8 weeks after an acute respiratory illness.
- One RCT⁸ compared beclomethasone and placebo in 72 patients with prolonged acute cough (10 days-3 weeks), so likely not true post-infectious (subacute) cough.
 - o ICS improved 3 of 6 outcomes over placebo on device-measured cough, but no difference in patient-reported symptoms.
- Guidelines suggest considering a trial of inhaled ipratropium or, if refractory, inhaled corticosteroids.⁹
- RCTs above did not include patients with COVID-19.

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