

BMJ 2014;349:g5703 doi: 10.1136/bmj.g5703 (Published 17 October 2014)



# PRACTICE

# CHANGE PAGE

# Avoid prescribing antibiotics in acute rhinosinusitis

Change Page aims to alert clinicians to the immediate need for a change in practice to make it consistent with current evidence. The series advisers are Sera Tort, clinical editor, and David Tovey, editor in chief, the Cochrane Library. We welcome any suggestions for future articles (changepage@bmj.com).

Wytske J Fokkens professor of otorhinolaryngology, clinical epidemiologist<sup>1</sup>, Ruth Hoffmans otorhinolaryngologist in training<sup>1</sup>, Mike Thomas professor of primary care research<sup>2</sup>

<sup>1</sup>Department of Otorhinolaryngology, Academic Medical Centre, Amsterdam, Netherlands; <sup>2</sup>Department of Primary Care Research, University of Southampton, UK

Acute rhinosinusitis (ARS) is an acute inflammatory condition of the nose and sinuses that is characterised by sudden nasal blockage, discharge, facial pain, or pressure and reduction in smell in adults or cough in children.<sup>1</sup> It is common, having a global prevalence of 6-15%,12 and it is usually managed in primary care. Despite consistent evidence of spontaneous resolution and recommendations to restrict antibiotics to severe illness, more than 80% of people with ARS receive antibiotics in Europe and North America.<sup>2-5</sup> Prescription rates might be lower (30%) in Asia, although over the counter availability of antibiotics in some settings makes accurate figures difficult to ascertain.<sup>2</sup> High prescribing results in pressure for antibiotic resistance and in adverse events. However, the primary cause of ARS is postviral inflammation. Fewer than 2% of patients have the more severe presentation of "bacterial ARS," a clinical rather than microbiological diagnosis characterised by discoloured discharge, severe local unilateral pain, fever (>38°C), raised levels of inflammatory markers (erythrocyte sedimentation rate and C reactive protein) and/or "double sickening" (deterioration after an initial milder illness).<sup>1</sup> The gold standard diagnostic test of true bacterial ARS is a positive culture from an invasive sinus puncture or meatal swab (ideally endoscopically guided); radiological opacification of the sinuses has less diagnostic value.<sup>1</sup> Neither bacteriology nor radiology is recommended in making the clinical diagnosis of bacterial ARS or guiding management. Individual symptoms are poorly predictive, but there is limited evidence to suggest that combinations of clinical factors (while not diagnostic of bacterial infection) may alert clinicians to patients with more severe and prolonged illness-for example, lasting beyond 10 days or worsening after 5-7 days-who should be monitored and considered for more intensive treatment, including antibiotics.<sup>6</sup>7(8)

The diagnostic criteria for bacterial ARS are similar between guidelines, derived from expert consensus and observational data, and the box outlines two examples.<sup>1 8</sup> These centre on the severity, character, and duration of symptoms as cited above. About a third of those with a clinical diagnosis of ARS will have bacteria identified on endoscopic sampling,<sup>9</sup> and most of these people will recover fully without antibiotic treatment. No controlled trials have shown that even bacterial ARS requires an antibiotic, although placebo controlled studies might be deemed unethical in those with a more severe illness. All current guidelines state that the combination of at least three of the severe symptoms and signs listed in the box should make the clinician at least consider antibiotic treatment.<sup>10 8</sup> We propose avoiding prescribing antibiotics in ARS unless several of the features given in the box are present.

# The evidence for change

Systematic reviews show that uncomplicated ARS resolves without antibiotic treatment.<sup>11 12</sup> A Cochrane review of antibiotics against placebo in adults with ARS found 10 trials (eight from primary care) with a low risk of bias, involving 2450 participants. Antibiotics provided no meaningful benefits; they can marginally shorten the time to cure (by less than half a day), but only five more participants per 100 will be cured by 7-14 days, and 18 participants (95% confidence interval, 10 to 115) will need to be treated for one patient to be cured more quickly. This needs to be weighed against adverse effects of antibiotics-the number needed to treat to harm was only 8  $(95\% \text{ confidence interval 6 to } 13)^{11}$  with the most common adverse events being gastrointestinal disturbances (nausea, vomiting, diarrhoea) and rash. Serious adverse events were uncommon in both arms. Given the lack of clear benefits and the pressing global problem of antibiotic resistance,13 the authors

Correspondence to: W J Fokkens w.j.fokkens@amc.nl

#### Key points

Only consider prescribing antibiotics in patients with symptoms of acute rhinosinusitis (ARS), for instance with at least three of the following more severe symptoms: purulent secretion, high fever, severe (unilateral) facial pain, prolonged illness (7 days or more), and/or "double sickening."

The prescription of antibiotics does not prevent serious complications in ARS

Combinations of clinical factors that may indicate more severe disease and consideration of antibiotic therapy

Bacterial ARS is characterised by the presence of at least three of the following symptoms in European guidelines: • Discoloured discharge (with unilateral predominance) and purulent secretion in the nasal cavity

Diagnosis of bacterial ARS requires the presence of at least two of the following symptoms, which must include item 2 or 3, and symptoms persisting beyond 10 days or worsening after 5-7 days in Canadian guidelines.<sup>3</sup>

Facial pain, pressure or fullness,

Nasal obstruction,

Nasal purulence or discolored postnasal discharge,

Hyposmia or anosmia

state: "there is no place for antibiotics for the patient with clinically diagnosed, uncomplicated acute rhinosinusitis."

A separate Cochrane review of antibiotics versus placebo for acute maxillary sinusitis (a common subgroup of ARS) with symptoms lasting at least seven days<sup>14</sup> found six controlled trials. There was a modest symptom resolution benefit with antibiotics, but improvement was high in both the placebo (80%) and the antibiotic treated groups (90%). There was also only a marginal difference in "total cure" rates between groups, with antibiotics resulting in a small reduction in relative risk of ongoing symptoms at 7 to 15 days (0.73, 0.63 to 0.85). The authors conclude that the modest benefits must be weighed against the potential for adverse effects at both individual and population levels.

The evidence shows that in a primary care setting, antibiotics have little if any role in ARS in adults and only a small treatment effect in patients with severe symptoms that persist beyond a week.<sup>14</sup>

Non-antibiotic treatment options include information on disease course, reassurance, and symptomatic treatment. Although widely used, there is no convincing evidence of clinically relevant benefits from antihistamines, steam inhalation, decongestants, or saline irrigation.<sup>1 15</sup> Topical nasal steroids have been shown to have a modest effect on symptoms and speed of recovery.<sup>16</sup> A Cochrane review found that symptoms of participants receiving this treatment were more likely to resolve at two weeks compared with those receiving placebo (73%  $\nu$  66.4%; risk ratio 1.11, 1.04 to 1.18). This modest benefit is similar to that observed for antibiotics. Although this review reported no significant adverse events, possible adverse effects can include nasal irritation and epistaxis.<sup>17 18</sup> Current topical nasal steroid preparations are not licensed for this indication.

## Barriers to change

Doctors want to prevent serious complications of ARS, such as orbital or intracranial abscess, which represent medical emergencies requiring prompt recognition and treatment. Clinical case series from specialist units treating these complications, however, suggest that they occur rarely and early in the course of the disease, and that the prevalence and the outcome are not influenced by early antibiotics in primary care.<sup>19 20</sup>

Research evaluating drivers for overprescribing antibiotics in respiratory tract infections suggest uncertainty in diagnosis and management, perceptions of patient expectation and potential conflict with patients,<sup>21</sup> availability of antibiotics over the counter,<sup>22</sup> and unawareness of local resistance problems<sup>23</sup> are important. Professional education and communication training, with or without additional near patient C reactive protein testing, can substantially reduce antibiotic use in respiratory infections.<sup>24</sup>

## How should we change our practice?

Antibiotic treatment should not be used in adults with uncomplicated ARS, and we propose that it should only considered for the small minority with features such as high fever, severe (unilateral) facial pain, purulent rhinorrhoea and "double sickening."<sup>1</sup>

Non-antibiotic treatment strategies centred on symptom control and the provision of information on the inflammatory but non-bacterial self limiting nature of the disease, and the lack of benefit and potential harm of antibiotics, should be usual first line management, for both individual and population health considerations.

Contributors: All authors agreed on the final version of this paper. WF: conception and design, drafting the article and final approval of the version to be published. Responsible for the overall content as guarantor. MT and RH: acquisition, analysis, and interpretation of the literature, revision of the manuscript and final approval of the version to be published.

Competing interests: We have read and understood BMJ policy on declaration of interests and declare the following interests: WF contributed to a study on local corticosteroids in acute rhinosinusitis for GlaxoSmithKline. The department of otorhinolaryngology received grants for studies of chronic rhinosinusitis from GlaxoSmithKline and Bioinspire and for studies on allergic rhinitis from Allergopharma and Medapharma. The department also received a contribution for travel and an honorarium for WF to produce a webcast on rhinosinusitis for general practitioners and speak on allergic rhinitis in Japan, which included assessment of effects of nasal corticosteroids in allergic rhinitis. RH has no conflict of interest. MT received an honorarium from GlaxoSmithKline to speak at the symposium sponsored by GlaxoSmithKline at the European Academy of Allergy and Clinical Immunology on quality of life in asthma and allergic rhinitis, which included an assessment of the effects of nasal steroids on quality of life.

Provenance and peer review: Not commissioned; externally peer reviewed.

 Fokkens WJ, Lund VJ, Mullol J, Bachert C, Alobid I, Baroody F, et al. European position paper on rhinosinusitis and nasal polyps. *Rhinology* 2012:3:1-298.

- 2 Wang DY, Wardani RS, Singh K, Thanaviratananich S, Vicente G, Xu G, et al. A survey on the management of acute rhinosinusitis among Asian physicians. *Rhinology* 2011;49:264-71.
- 3 Smith SS, Evans CT, Tan BK, Chandra RK, Smith SB, Kern RC. National burden of antibiotic use for adult rhinosinusitis. J Alleray Clin Immunol 2013:132:1230-2
- antibiotic use for adult rhinosinusitis. J Allergy Clin Immunol 2013;132:1230-2.
  Jorgensen LC, Friis Christensen S, Cordoba Currea G, Llor C, Bjerrum L. Antibiotic prescribing in patients with acute rhinosinusitis is not in agreement with European recommendations. Scand J Prim Health Care 2013;31:101-5.
  Hoffmans R, Schermer T, van Weel C, Fokkens W. Management of rhinosinusitis in Dutch
- 5 Hoffmans R, Schermer T, van Weel C, Fokkens W. Management of rhinosinusitis in Dutch general practice. *Prim Care Respir J* 2011;20:64-70.
- 6 Hansen JG, Schmidt H, Rosborg J, Lund E. Predicting acute maxillary sinusitis in a general practice population. BMJ 1995;311:233-6.
- 7 Williams JW, Jr., Simel DL, Roberts L, Samsa GP. Clinical evaluation for sinusitis. Making the diagnosis by history and physical examination. Ann Intern Med 1992;117:705-10.
- 8 Desrosiers M, Evans GA, Keith PK, Wright ED, Kaplan A, Bouchard J, et al. Canadian clinical practice guidelines for acute and chronic rhinosinusitis. J Otolaryngol Head Neck Surg 2011;40:S99-193.
- 9 Shintani Smith S, Ference EH, Evans CT, Tan BK, Kern RC, Chandra RK. The prevalence of bacterial infection in acute rhinosinusitis: a systematic review and meta-analysis. *Laryngoscope* 2014; published online 9 Apr.
- 10 Chow AW, Benninger MS, Brook I, Brozek JL, Goldstein EJ, Hicks LA, et al. IDSA clinical practice guideline for acute bacterial rhinosinusitis in children and adults. *Clin Infect Dis* 2012;54:e72-112.
- 11 Lemiengre MB, van Driel ML, Merenstein D, Young J, De Sutter AI. Antibiotics for clinically diagnosed acute rhinosinusitis in adults. *Cochrane Database Syst Rev* 2012;10:CD006089.
- 12 Young J, De Sutter A, Merenstein D, van Essen GA, Kaiser L, Varonen H, et al. Antibiotics for adults with clinically diagnosed acute rhinosinusitis: a meta-analysis of individual patient data. *Lancet* 2008;371:908-14.
- 13 Goossens H, Ferech M, Vander Stichele R, Elseviers M. Outpatient antibiotic use in Europe and association with resistance: a cross-national database study. *Lancet* 2005;365:579-87.
- 14 Ahovuo-Saloranta A, Borisenko OV, Kovanen N, Varonen H, Rautakorpi UM, Williams JW Jr, et al. Antibiotics for acute maxillary sinusitis. *Cochrane Database Syst Rev* 2008;16:CD000243.

- 15 Shaikh N, Wald ER, Pi M. Decongestants, antihistamines and nasal irrigation for acute sinusitis in children. *Cochrane Database Syst Rev* 2010;12:CD007909.
- 16 Zalmanovici A, Yaphe J. Intranasal steroids for acute sinusitis. Cochrane Database Syst Rev 2009;4:CD005149.
- 17 Sastre J, Mosges R. Local and systemic safety of intranasal corticosteroids. J Investig Allergol Clin Immunol 2012;22:1-12.
- Schafer T, Schnoor M, Wagenmann M, Klimek L, Bachert C. Therapeutic Index (TIX) for intranasal corticosteroids in the treatment of allergic rhinitis. *Rhinology* 2011;49:272-80.
   Babar-Craig H, Gupta Y, Lund VJ. British Rhinological Society audit of the role of antibiotics
- in complications of acute rhinosinusitis: a national prospective audit. Rhinology 2010;48:344-7.
- 20 Hansen FS, Hoffmans R, Georgalas C, Fokkens WJ. Complications of acute rhinosinusitis in the Netherlands. *Fam Pract* 2012;29:147-53.
- 21 Tonkin-Crine S, Yardley L, Little P. Antibiotic prescribing for acute respiratory tract infections in primary care: a systematic review and meta-ethnography. J Antimicrobial Chemotherapy 2011;66:2215-23.
- 22 Brookes-Howell L, Hood K, Cooper L, Little P, Verheij T, Coenen S, et al. Understanding variation in primary medical care: a nine-country qualitative study of clinicians' accounts of the non-clinical factors that shape antibiotic prescribing decisions for lower respiratory tract infection. *BMJ Open* 2012;2:4.
- 23 Wood F, Phillips C, Brookes-Howell L, Hood K, Verheij T, Coenen S, et al. Primary care clinicians' perceptions of antibiotic resistance: a multi-country qualitative interview study. *J Antimicrobial Chemotherapy* 2013;68:237-43.
- 24 Little P, Stuart B, Francis N, Douglas E, Tonkin-Crine S, Anthierens S, et al. Effects of internet-based training on antibiotic prescribing rates for acute respiratory-tract infections: a multinational, cluster, randomised, factorial, controlled trial. *Lancet* 2013;382:1175-82.

#### Accepted: 13 August 2014

#### Cite this as: *BMJ* 2014;349:g5703

© BMJ Publishing Group Ltd 2014