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PRACTICE



UNCERTAINTIES

What treatments are effective for common cold in adults and children?

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What you need to know

- Quality evidence to say whether over-the-counter treatments work for nasal symptoms of the common cold is limited.
- For adults, consider a trial of decongestants alone, or with antihistamines or analgesics to alleviate bothersome nasal symptoms.
- Do not prescribe decongestants to children under 12, as evidence of their effectiveness is limited and associated risks may exist.

The common cold is usually caused by viruses and is mostly self limiting,¹ but it can have a substantial impact on work, school,² use of health services, and money spent on medications. Children have around 6-8 colds per year and adults have 2-4.³⁴

Many over-the-counter (OTC) treatments for the common cold claim to alleviate nasal symptoms, such as congestion, rhinorrhoea (runny nose), and sneezing. Table 1 lists commonly used drugs. Evidence for the effectiveness of these treatments is limited and of low quality, and clear guidance is lacking.⁵ Long term use of nasal decongestants is known to lead to chronic nasal congestion.⁶

What is the evidence of uncertainty?

Search strategy and study selection

We searched the Cochrane Library for systematic reviews that investigate the effectiveness of treatments for the common cold. If only a protocol or no Cochrane review was available, we searched PubMed for other systematic reviews on the topic. If no systematic reviews were found, we searched for individual randomised controlled trials of commonly used treatments (fig 1, table 2). We extracted data on the subjective severity and duration of nasal symptoms (nasal congestion, rhinorrhoea, and sneezing) and adverse events. We extracted the number of studies and participants, and where available, used pooled results. If pooled results were not available, we assessed whether the findings were in favour of the active treatment.

Adults

We found Cochrane reviews on treatments such as decongestants, antihistamines, analgesics, intranasal corticosteroids, herbal remedies, and vitamins and minerals (zinc) in adults with common cold. Commonly reported primary outcomes in the included studies are nasal resistance measures or outcomes such as clinical cure or composite symptom scores. Only a few studies included in these reviews report on bothersome nasal symptoms, such as congestion, rhinorrhoea, and sneezing. In summary, low quality evidence suggests that decongestants (either in monotherapy or in combination with antihistamines and/or analgesics) have a small effect on nasal symptoms (fig 1). Harms include an increased risk of insomnia, drowsiness, headache, or gastrointestinal upset (fig 1, table 2).⁷¹¹ Long term use can lead to chronic nasal congestion. However, the recommended safe treatment duration for decongestants varies and seems to be based on expert opinion.

A Cochrane review⁸ (four randomised controlled trials, 1466 participants) shows that sedating antihistamines are associated with relief of rhinorrhoea and sneezing compared with placebo, but not nasal congestion (two randomised controlled trials, 375 participants). Sedation was commonly reported, but there were no differences between groups (6 randomised controlled trials, 2265 participants). Studies with non-sedating antihistamines show an unclear effect on congestion (one randomised controlled trial, 53 participants), and no effect on rhinorrhoea (three randomised controlled trials, 838 participants), or sneezing (four randomised controlled trials, 456 participants) and no increased risk of adverse events compared with placebo.⁸

A Cochrane meta-analysis showed no effect of antibiotics on nasal symptoms, but the risk of adverse events was increased.¹⁵ Evidence does not exist for the effectiveness of antivirals, and intranasal corticosteroids for nasal symptoms in the common

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cold, and their use is not recommended.¹²³² Acetaminophen/paracetamol and NSAIDs are sometimes prescribed for pain relief in common cold, but they do not appear to improve nasal congestion or rhinorrhoea.^{9 10} Low quality

evidence suggests intranasal ipratropium bromide reduces rhinorrhoea compared with placebo, but not nasal congestion. Nosebleeds, nasal dryness, and dry mouth are side effects.¹³

Nasal symptoms are not reported in trials investigating the effect of echinacea,¹⁷ vitamin C,¹⁶ zinc lozenges,²⁴⁻²⁶ and heated humidified air or steam.²¹ Echinacea does not seem to improve overall symptoms.¹⁷ Zinc lozenges have been shown to reduce the duration but not severity of cold symptoms,²⁴⁻²⁶ but the optimal composition and dosage of lozenges has not been established. No evidence exists for the use of heated humidified air or steam in the common cold.²¹ A Cochrane review concludes that saline irrigations are not likely to be effective in adults.²⁰ We did not find trials studying the effect on common cold symptoms for the following treatments: probiotics,²⁷⁻²⁹ garlic,¹⁸ Chinese medicinal herbs,³⁰ vapour rub,²² eucalyptus oil, honey,³¹ ginseng,²³ and increased fluid intake.³³

Children

Trials are lacking for children under 12, who carry the highest burden of common colds. A Cochrane review found low quality evidence that saline irrigations or drops may be effective and safe in young children.²⁰ A small number of trials report contradictory results for decongestants and antihistamines on nasal symptoms and safety in children.⁷⁸¹¹ Some products that contain decongestant may improve nasal symptoms in children, but their safety, especially in young children, is unclear. We did not find evidence to support the use of other common treatments and home remedies in children (such as heated humidified air or steam, analgesics, echinacea, probiotics, herbs, or vitamins).

Decongestants, antihistamines, and analgesics in monotherapy

A Cochrane review⁷ (2 randomised controlled trials, 94 participants) comparing oral or intranasal decongestants with placebo found that 3-4 doses per day (over 5 days and up to 10 days) was associated with reduced severity of nasal congestion. Short term adverse events were no different between decongestants and placebo (7 randomised controlled trials, 1195 participants). No trials compared oral with intranasal routes.

A Cochrane review⁸ (4 randomised controlled trials, 1466 participants) shows that sedating antihistamines are associated with relief of rhinorrhoea and sneezing compared with placebo, but not nasal congestion (2 randomised controlled trials, 375 participants). Sedation was commonly reported, but there were no differences between groups (6 randomised controlled trials, 2265 participants). Studies with non-sedating antihistamines show an unclear effect on congestion (1 randomised controlled trial, 53 participants), and no effect on rhinorrhoea (3 randomised controlled trials, 838 participants), or sneezing (4 randomised controlled trials, 456 participants) and no increased risk of adverse events compared with placebo.⁸

In a Cochrane review⁹ (4 randomised controlled trials, 758 participants), investigating the effect of acetaminophen/paracetamol compared with placebo on pain and common cold symptoms, only one trial (n=60) reports specific nasal symptoms, noting an unclear effect on severity of symptoms and possible increase of nasal congestion in the acetaminophen group. Adverse events, such as sweating and

analysis of 3 trials (n=199) showed no effect of NSAIDs on nasal congestion or rhinorrhoea compared with placebo, although sneezing was reduced (2 randomised controlled trials, n=159).¹⁶ Adverse events, such as rash, oedema and gastro-intestinal complaints, were not different between groups (2 randomised controlled trials, n=220).

Combinations of decongestants, antihistamines, and analgesics

A Cochrane review¹¹ (27 randomised controlled trials, 5117 participants) evaluated the effect of different combinations of decongestants, antihistamines, and analgesics in the common cold.

Oral antihistamine decongestant combinations¹¹ and analgesic decongestant combinations may improve congestion and sneezing, but data could not be pooled because of heterogeneity. Patients taking combinations reported more adverse effects such as sedation, insomnia, and headache.¹¹

Of three trials studying oral antihistamine-analgesic combinations, two (341 participants) showed no improvement of nasal congestion compared with placebo or acetaminophen. In one trial (150 participants) the combination was associated with less sneezing. Adverse events (nasal irritation, dry mouth, gastrointestinal upset) occurred in both groups (3 randomised controlled trials, 1508 participants).¹¹

Oral antihistamine analgesic decongestant combinations¹¹ were consistently associated with reduced nasal congestion and rhinorrhoea compared with placebo (3 randomised controlled trials, 595 participants). It is unclear if adverse events were different between groups.

Ipratropium bromide

Low quality evidence finds that intranasal ipratropium bromide reduces rhinorrhoea compared with placebo, but not nasal congestion, however there is an increased risk of nosebleeds, nasal dryness, and dry mouth.¹³ A trial with 786 participants reported that decongestant ipratropium bromide combination improved both nasal congestion and rhinorrhoea compared with placebo, with similar adverse events.¹⁴

Antibiotics

Antibiotics are not indicated for viral infections such as the common cold. A Cochrane meta-analysis¹⁵ (6 randomised controlled trials, 1047 participants) showed that antibiotics did not reduce duration of purulent rhinitis (4 randomised controlled trials, 723 participants) or clear rhinitis (2 randomised controlled trials, 227 participants), but the risk of adverse events was increased (4 randomised controlled trials, 1267 participants). Effect on congestion was not reported and there was an unclear risk of bias overall.

Antivirals

A Cochrane review³² concludes that none of the licensed antivirals were effective in reducing symptoms, and adverse events make them unacceptable for use in the common cold. This review was withdrawn in 2004 as unpublished data from the original review were not accessible.

Outcomes in children

Few trials investigate the effect of common cold treatments in children, showing only small effects (fig 1, table 2). In young children (1.5-60 months) sedating antihistamines were associated with shorter duration of rhinorrhoea,⁸ and non-sedating

gastrointestinal upset, were more common with high dose

paracetamol (1000 mg) in another trial (n=392). A pooled

antihistamines with shorter duration of overall symptoms, but nasal symptoms were not reported.⁸ Adverse events were either not reported (non-sedating) or not different (sedating).⁸ The Cochrane review on combination treatments for common cold reported that a combination of acetaminophen decongestant antihistamine in children improved nasal congestion on day 5 (although not on day 3) compared with acetaminophen alone.¹¹ An NSAID decongestant combination reduced the duration of nasal congestion compared with pseudoephedrine or placebo.¹¹ Antihistamine-decongestant combinations did not show consistent effects on nasal symptoms.¹¹ Saline nasal irrigation may improve nasal congestion in older children and possibly reduce rhinorrhoea severity.²⁰ Vapour rub may improve nasal congestion (not rhinorrhoea), but at an increased risk of adverse events.²²

A trial with echinacea does not report nasal symptoms, but shows it increases the risk of a rash.¹⁷ The trial of ginseng did not report nasal symptoms,²³ nor did studies with honey.⁵ Furthermore, we did not find any trials studying the effect of the following treatments in children with common cold: decongestants in monotherapy,⁷ NSAIDs¹⁰ or paracetamol⁹ in monotherapy, intranasal corticosteroids,¹² intranasal ipratropium bromide,¹³ antivirals,³² eucalyptus oil,²² fluid intake,³³ garlic,¹⁸ heated humidified air,²¹ Chinese medicinal herbs,³⁰ Pelargonium sidoides,¹⁹ probiotics,²⁷⁻²⁹ vitamin C,¹⁶ and zinc.^{25 26}

Is ongoing research likely to provide relevant evidence?

A search of International Clinical Trials Registry Platform using the terms "common cold" or "respirat*" yielded 17 references to ongoing trials. These trials use

analgesic-decongestant-antihistamine combinations (n=3), an intranasal decongestant (n=1), Chinese (n=3) or other herbs (n=4), herbal steam inhalation (n=1), lactic acid bacteria (n=1), pelargonium (n=1), guaifenesin (n=1), and antivirals (n=2). Twelve of these trials include adults (and older children), four include only children, and one includes participants of all ages.

Most of these studies have reasonable sample sizes but few report on nasal symptoms. Five trials explicitly mention they will report on nasal symptoms, and only one of these includes children. Several traditional Chinese, Thai, and Indian herbal treatments are also studied, but none of these trials will provide information about the effect on nasal symptoms. It is unlikely that these will address the uncertainty. No evidence yet exists on the effect of guaifenesin, an expectorant used to treat cough, on nasal symptoms. This study may add to the evidence base.

What should we do in light of the uncertainty?

The common cold is self limiting and symptoms usually clear within 7 to 10 days.³⁴ Explain to patients that there are no "magic bullets" to relieve their symptoms and that very few OTC treatments are supported by evidence.

For adults with bothersome nasal symptoms, decongestants and antihistamines in monotherapy or in combination products are the best choice. However, the effect is small and although the adverse events are usually mild, some—such as sedation—can be disturbing. No evidence suggests that a tablet taken orally or a nasal spray is the more effective. Advise patients to use nasal decongestants for a maximum of 3 to 7 days.³⁵⁻³⁸ Patients often take OTC decongestants before they consult the GP and commonly for more than just a few days.⁶ They may not be aware that prolonged use can lead to chronic nasal congestion (rhinitis medicamentosa). None of the other commonly used OTC treatments have been shown to relieve nasal symptoms and many have not been studied at all. Based on the currently available evidence, reassurance that symptoms are self limiting is the best you can offer patients.

The evidence for common cold treatments in children is more limited. We do not recommend decongestant or formulations containing antihistamine in children under 6 and advise caution between 6 and 12 years.³⁵⁻³⁸ There is no evidence that these treatments alleviate nasal symptoms and they can cause adverse effects such as drowsiness or gastrointestinal upset. Serious harm, such as convulsions, rapid heart rate and death have been linked to decongestant use in very young children. None of the other commonly used OTC and home treatments, such as heated humidified air, eucalyptus oil, or echinacea are supported by adequate evidence.

Explain that a cold is distressing but should pass in 7-10 days. If parents are concerned about their child's comfort, saline nasal irrigations can be given to alleviate nasal symptoms.

Recommendations for future research

- Large, well conducted randomised controlled trials should include
 Population: children, especially young children as they carry the highest burden of common colds
 - Intervention: commonly used treatments such as nasal irrigations, steam inhalations or vaporizers with humidified air, eucalyptus or other aromatic oils, or vapour rub
 - Comparator: other commonly used treatments or head-to-head comparisons of active products (such as oral or intranasal decongestants)
 - Outcome: outcomes relevant to patients, eg, subjective nasal congestion rather than nasal patency, impact on daily life, short- and long term safety

Education into practice

- How do you discuss treatments for nasal symptoms of the common cold? With an adult? With the parent of a child?
- How would you explore duration of use for decongestant, and how would you address this issue?

What patients need to know

Common cold is usually self limiting—symptoms clear in 7 to 10 days. Your doctor may offer you medications to relieve headache, pain, or nasal congestion if these are bothersome.

Adults

o If a blocked or runny nose, or sneezing related to a cold is bothering you, you can try using nasal decongestants for up to 3 to 7 days

o Beware of unintended effects such as drowsiness, insomnia, or headache

o Do not take decongestants longer than advised as long term use may lead to chronic nasal congestion, which is difficult to treat

 $\ensuremath{\mathsf{o}}$ Other treatments have either not been effective in clinical trials or have not been studied at all

In children under 12

o Saline nasal irrigations or drops can be used safely, but this may not give the desired relief

 $\ensuremath{\mathsf{o}}$ Consult a doctor if symptoms are bothersome. Do not give children decongestants

o Vapour rub may relieve congestion but can cause skin rashes

o Other treatments, such as steam, humidified air, echinacea, or probiotics, are either not effective or have not been studied in children.

Information resources for patients

NHS Choiceshttp://www.nhs.uk/conditions/Cold-common/Pages/ Introduction.aspx

Definition of common cold, symptoms, treatment, complications, children. Free of charge. No registration needed

Mayo Clinichttp://www.mayoclinic.org/diseases-conditions/common-cold/ home/ovc-20199807#

Overview, symptoms and causes, diagnosis and treatment, self-management. Free of charge. No registration needed

How patients were involved in the creation of this article

We asked 10 customers seeking OTC treatments for the common cold in a community pharmacy in Belgium what concerned them most when they had a cold. This revealed a strong focus on managing nasal symptoms. Based on this, we decided to focus on the effect of commonly used treatments on subjective nasal symptoms in common cold. A patient reviewer acknowledged that while there is no clear cut way to resolve symptoms of nasal congestion, appropriate treatment options can be discussed for adults and for children. We have now presented the evidence for common treatments for adults and children separately and also clarified these in the section on 'what patients need to know'.

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- Andrewes CH. The natural history of the common cold. Lancet 1949;1:71-5. 10.1016/S0140-6736(49)90398-0 18236517
- 2 Hellgren J, Cervin A, Nordling S, Bergman A, Cardell LO. Allergic rhinitis and the common cold--high cost to society. *Allergy* 2010;65:776-83. . 10.1111/j.1398-9995.2009.02269.x 19958315
- 3 Gwaltney JM. Clinical significance and pathogenesis of viral respiratory infections. Am J Med 2002;112(Suppl 6A):13S-8S. 10.1016/S0002-9343(01)01059-2 11955455
- 4 Witek TJ, Ramsey DL, Carr AN, Riker DK. The natural history of community-acquired common colds symptoms assessed over 4-years. *Rhinology* 2015;53:81-8. . 10.4193/Rhin14.149 25756083
- 5 Allan GM, Arroll B. Prevention and treatment of the common cold: making sense of the evidence. CMAJ 2014;186:190-9. 10.1503/cmaj.121442 24468694
- 6 Mehuys E, Gevaert P, Brusselle G, etal . Self-medication in persistent rhinitis: overuse of decongestants in half of the patients. J Allergy Clin Immunol Pract 2014;2:313-9. 10.1016/j.jaip.2014.01.009. 10.1016/j.jaip.2014.01.009 24811023
- Deckx L, De Sutter AI, Guo L, Mir NA, van Driel ML. Nasal decongestants in monotherapy for the common cold. *Cochrane Database Syst Rev* 2016;10:CD009612.
 10.1002/14651858.CD09612.pub2.https://www.ncbi.nlm.nih.gov/entrez/query.fcgi? cmd-Batriave&th-PubMed&list_uide=272489558/dot-Abstract 27248955
- cmd=Retrieve&db=PubMed&list_uids=27748955&dopt=Abstract 27748955
 8 De Sutter AI, Saraswat A, van Driel ML. Antihistamines for the common cold. *Cochrane Database Syst Rev* 2015;11:CD009345. 10.1002/14651858.CD009345.pub2. 26615034
- 9 Li S, Yue J, Dong BR, Yang M, Lin X, Wu T. Acetaminophen (paracetamol) for the common cold in adults. *Cochrane Database Syst Rev* 2013;7:CD008800. 10.1002/14651858.CD008800.pub2.https://www.ncbi.nlm.nih.gov/entrez/query.fcgi? cmd=Retrieve&db=PubMed&list_uids=23818046&dopt=Abstract 23818046
- 10 Kim SY, Chang YJ, Cho HM, Hwang YW, Moon YS. Non-steroidal anti-inflammatory drugs for the common cold. *Cochrane Database Syst Rev* 2015;9:CD006362. 10.1002/14651858.CD006362.pub4.https://www.ncbi.nlm.nih.gov/entrez/query.fcgi? cmd=Retrieve&db=PubMed&list_uids=26387658&dopt=Abstract 26387658
- De Sutter AI, van Driel ML, Kumar AA, Lesslar O, Skrt A. Oral antihistamine-decongestant-analgesic combinations for the common cold. *Cochrane Database Syst Rev* 2012;15:CD004976.10.1002/14651858.CD004976.pub3.https://www.

ncbi.nlm.nih.gov/entrez/query.fcgi?cmd=Retrieve&db=PubMed&list_uids=22336807& dopt=Abstract 22336807

- 12 Hayward G, Thompson MJ, Perera R, Del Mar CB, Glasziou PP, Heneghan CJ. Corticosteroids for the common cold. *Cochrane Database Syst Rev* 2015;10:CD008116. 10.1002/14651858.CD008116.pub3.https://www.ncbi.nlm.nih.gov/entrez/query.fcgi? cmd=Retrieve&db=PubMed&list_uids=26461493&dopt=Abstract 26461493
- AlBalawi ZH, Othman SS, Alfaleh K. Intranasal ipratropium bromide for the common cold. Cochrane Database Syst Rev 2013;6:CD008231.
 10.1002/14651858.CD008231.pub3.https://www.ncbi.nlm.nih.gov/entrez/query.fcgi?
- cmd=Retrieve&db=PubMed&list_uids=23784858&dopt=Abstract 23784858
 Eccles R, Martensson K, Chen SC. Effects of intranasal xylometazoline, alone or in combination with ipratropium, in patients with common cold. *Curr Med Res Opin* 2010;26:889-99. 10.1185/03007991003648015 20151787
- 15 Kenealy T, Arroll B. Antibiotics for the common cold and acute purulent rhinitis. Cochrane Database Syst Rev 2013;6:CD000247. 10.1002/14651858.CD000247.pub3.https://www. ncbi.nlm.nih.gov/entrez/query.fcgi?cmd=Retrieve&db=PubMed&list_uids=23733381& dopt=Abstract 23733381
- 16 Hemilä H, Chalker E. Vitamin C for preventing and treating the common cold. Cochrane Database Syst Rev2013;(1):CD000980.10.1002/14651858.CD00980.pub4.https://www ncbi.nlm.nih.gov/entrez/query.fcgi?cmd=Retrieve&db=PubMed&list_uids=23440782& dopt=Abstract 23440782
- 17 Karsch-Völk M, Barrett B, Kiefer D, Bauer R, Ardjomand-Woelkart K, Linde K. Echinacea for preventing and treating the common cold. *Cochrane Database Syst Rev* 2014;2:CD000530. 10.1002/14651858.CD000530.pub3. 24554461
- 18 Lissiman E, Bhasale AL, Cohen M. Garlic for the common cold. Cochrane Database Syst Rev 2014;11:CD006206. 10.1002/14651858.CD006206.pub4
- 19 Timmer A, Günther J, Motschall E, Rücker G, Antes G, Kern WV. Pelargonium sidoides extract for treating acute respiratory tract infections. *Cochrane Database Syst Rev* 2013;10:CD006323. 10.1002/14651858.CD006323.pub3. 24146345
- 20 King D, Mitchell B, Williams CP, Spurling GK. Saline nasal irrigation for acute upper respiratory tract infections. *Cochrane Database Syst Rev* 2015;20:CD006821. 10.1002/14651858.CD006821.pub3. 25892369
- 21 Singh M, Singh M, Jaiswal N, Chauhan A. Heated, humidified air for the common cold. Cochrane Database Syst Rev 2017;8:CD001728. 10.1002/14651858.CD001728.pub6. 28849871
- Paul IM, Beiler JS, King TS, Clapp ER, Vallati J, Berlin CMJr. Vapor rub, petrolatum, and no treatment for children with nocturnal cough and cold symptoms. *Pediatrics* 2010;126:1092-9, 10.1542/peds.2010-1601 21059712
- 23 Vohra S, Johnston BC, Laycock KL, etal. Safety and tolerability of North American ginseng extract in the treatment of pediatric upper respiratory tract infection: a phase II randomized, controlled trial of 2 dosing schedules. *Pediatrics* 2008;122:e402-10. 10.1542/peds.2007-2186 18676527
- 24 Singh M, Das RR. WITHDRAWN: Zinc for the common cold. Cochrane Database Syst Rev 2015;4:CD001364. 10.1002/14651858.CD001364.pub5. 25924708
- 25 Hemilä H. Zinc lozenges and the common cold: a meta-analysis comparing zinc acetate and zinc gluconate, and the role of zinc dosage. JRSM Open 2017;8:2054270417694291. 10.1177/2054270417694291 28515951
- 26 Hemilä H, Petrus EJ, Fitzgerald JT, Prasad A. Zinc acetate lozenges for treating the common cold: an individual patient data meta-analysis. Br J Clin Pharmacol 2016;82:1393-8. 10.1111/bcp.13057 27378206
- 27 Araujo GV, Oliveira Junior MH, Peixoto DM, Sarinho ES. Probiotics for the treatment of upper and lower respiratory-tract infections in children: systematic review based on randomized clinical trials. J Pediatr (Rio J) 2015;91:413-27. . 10.1016/j.jped.2015.03.002 26054771
- 28 Wang Y, Li X, Ge T, etal . Probiotics for prevention and treatment of respiratory tract infections in children: A systematic review and meta-analysis of randomized controlled trials. *Medicine (Baltimore)* 2016;95:e4509. . 10.1097/MD.00000000004509 27495104
- 29 Robinson JL. Probiotics for modification of the incidence or severity of respiratory tract infections. *Pediatr Infect Dis J* 2017;36:1093-5. . 10.1097/INF.000000000001714 28767618
- 30 Wu T, Zhang J, Qiu Y, Xie L, Liu GJ. Chinese medicinal herbs for the common cold. Cochrane Database Syst Rev 2007;1:CD004782.
- 10.1002/14651858.CD004782.pub2. 17253524
 Smith SM, Schroeder K, Fahey T. Over-the-counter (OTC) medications for acute cough in children and adults in community settings. *Cochrane Database Syst Rev*
- 2014;11:CD001831. 10.1002/14651858.CD001831.pub5. 25420096
 Jefferson TO, Tyrrell D. WITHDRAWN: Antivirals for the common cold. *Cochrane Database Syst Rev* 2007;3:CD002743. 17636705
- Guppy MP, Mickan SM, Del Mar CB, Thorning S, Rack A. Advising patients to increase fluid intake for treating acute respiratory infections. *Cochrane Database Syst Rev* 2011;2:CD004419.21328268
- 34 Lorber B. The common cold. J Gen Intern Med 1996;11:229-36. 8744881
- 35 NHS Choices. Decongestants 2016. https://www.nhs.uk/conditions/decongestants/#howto-use-decongestants
- 36 Therapeutic Guidelines Ltd. Ear, nose and throat infections Melbourne, Australia 2014 https://tgldcdp.tg.org.au/viewTopic?topicfile=ear-nose-throat-infections& guidelineName=Antibiotic#toc_d1e888.
- 37 Government of Canada. Guidance document—Nonprescription topical nasal decongestants labelling standard 2014. https://www.canada.ca/en/health-canada/services/drugs-healthproducts/drug-products/applications-submissions/guidance-documents/nonprescriptiondrugs-labelling-standards/nonprescription-topical-nasal-decongestants-labelling-standard. html#s5.1
- 38 Therapeutic Goods Administration. OTC medicine monograph: Topical nasal decongestants: Australian Government Department of Health. 2014. https://www.tga.gov. au/otc-medicine-monograph-topical-nasal-decongestants.

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Tables

Table 1 Drugs used to treat symptoms in c	common cold
Decongestant	
Sympathomimetic: oral	Ephedrine hydrochloride, pseudoephedrine hydrochloride, phenylephrine hydrochloride
Sympathomimetic: intranasal	Naphazoline nitrate, oxymetazoline hydrochloride, tramazoline hydrochloride, xylometazoline hydrochloride
Antihistamine	
Older, "first generation," sedating	Alimemazine tartrate, chlorphenamine maleate, clemastine, cyproheptadine hydrochloride, dimethindene maleate, hydroxyzine hydrochloride, ketotifen, promethazine hydrochloride
Newer, "second generation," non-sedating	Acrivastine, azelastine hydrochloride, bitati ne, cetirizine hydrochloride, desloratadine, fexofenadine hydrochloride, levocabasti be by drochloride, levocetirizine hydrochloride, loratadine, mizolastine, olopatadine hydrochloride
Analgesic	Acetaminophen/paracetamol
	Non-steroidal anti-inflammatory drugs (NSAIDs):
	acetylsalicylic acid (aspirin), ibuprofen, naproxen
Nasal corticosteroids	Beclomethasone diproprionate, budesonide, ciclesonide, fluticasone furoate/propionate, flunisolide, mometasone furoate, triamcinolone acetonide
Antimuscarinic	Ipratropium bromide
Saline nasal irrigation	Sodium chloride 0.9% (saline)

Availability and OTC status of the products vary by country

* Both oral and intranasal

** Intranasal

Table 2| Efficacy (subjective nasal symptoms) and harm of common cold treatments in adults and children: overview of the available evidence from Cochrane reviews and clinical trials

			Adults	
Treatment option	Outcome	Included	Adults Effect aaaaaaaaaaaaaaaaaaaaaaaaaaaaaaaaaaaa	
.	o	studies	aaaaaaaaaaaaaaaaabb	
Decongestants Cochrane review ⁷	Congestion	n=94; 2 RCTs	Small effect on severity : Pooled effect after 3 hours: SMD 0.49 (95% CI 0.07 to 0.92)	
	Dhimaushaaa		Fooled effect after 3 hours. SMD 0.43 (93% CI 0.07 to 0.92)	
	Rhinorrhoea	Not reported		
	Sneezing	Not		
	Ū	reported		
	Adverse	n=1195;	No increased risk:	
	events	7 RCTs	Pooled risk: OR 0.98 (95% CI 0.68 to 1.40)	hł
Sedating antihistamines	Congestion		No effect on severity	
Cochrane review ⁸		RCTs	Pooled effect after day 1: MD -0.07 (95% CI -0.29 to 0.15); scale 0-4	
		n=27; 1 RCT	aaaaaaaaaaaaaaaaaaaaaaaaaaaaaaaaaaaa	
	Dhim a mula a			ak
	Rhinorrhoea	n=1466; 4 RCTs	No pooling: 1 RCT shows no effect Gaaaaaaaaaaaaaaaaaaaaaaaaaaaaaaaaaaa	
		n=1465;	Pooled effect after day 2: MD =0.18 (95% CI =0.13 to 0.00); scale 0-4	
		4 RCTs		
		n=27; 1	Inclear effect on duration	
		RCT	No pooling: 1 RCT no effect	
				- 1-
	Sneezing	n=1466;	Addadaaaaaaaaaaaaaaaaaaaaaaaaaaaaaaaaa	ac
		4 RCTs	Pooled effect after day 1: MD -0.07 (95% CI -0.15 to 0.00); scale 0-4	
		n=1465;	Pooled effect after day 2: MD -0.29 (95% Cl -0.38 to -0.21) scale 0-4)	
		4 RCTs	Unders affect on describer	
		n=27; 1 RCT	Unclear effect on duration	
	Advaraa			
	Adverse events	n=2265; 6 RCTs	Pooled OB 1 13 (95% CI 0 80 to 1 59)	
Non-sedating antihistamines	Congestion	n=53; 1	Unclear effect on severity	-
Cochrane review ⁸	0	RCT	No pooling: 1 RCT no effect	
	Rhinorrhoea	n=383; 3	No effect on severity	Ð
		RCTs	Pooled effect after day 4: MD -0.08 (95% CI -0.26 to 0.09); scale 0-4	
	Sneezing	n=456; 4	Possibly no effect on severity	
		RCTs	No pooling: 4 RCTs no effect	
	Adverse	n=215; 3	No increased risk	
	events	RCTs	Pooled OR 1.21 (95% CI 0.52 to 2.81)	
Paracetamol/	Congestion	n=60; 1	Unclear effect on severity	
Acetaminophen	-	RCT	No pooling: increased nasal congestion	
Cochrane review9				_
	Rhinorrhoea	Not	Fooled OK 1.13 (95% Cl 0.60 to 1.39) Unclear effect on severity No pooling: 1 RCT no effect aaaaaaaaaaaaaaaaaaaaaaaaaaaaaaaaaaa	
		reported		

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			Adults	c
Treatment option	Outcome	Included studies	Effect	
	Sneezing	Not reported		
	Adverse events	1 RCT (n=392)	Unclear risk	
ISAIDs	Congestion	· ,	No pooling: more minor AE, not clear if differences between groups aaaaaaaaaaaaaaaaaaaaaaaaaaaaaaaaaaaa	ahh
cochrane review10	0	RCTs	Pooled SMD –0.15 (95% Cl –0.43 to 0.13)	abc
	Rhinorrhoea	n=199; 3 RCTs	aaaaaaaaaaaaaaaaaaaaaaaaaaaaaaaaaaaaaa	aaa
	Sneezing	n=159; 2	Pooled SMD 0.03 (95% CI = 0,25 to 0.30) 	aab
	Sheezing	RCTs	Pooled SMD –0.44 (95% Cl –0.75 to –0.12)	
	Adverse	n=220; 2	No increased risk	
ntihistomino docon sostant	events	RCTs	Pooled RR 2.94 (95% CI 0.51 to 17.03)	
ntihistamine-decongestant ombination cochrane review ¹¹	Congestion	n=678; 5 RCTs	Unclear effect on severity No pooling: 3 RCTs (n=568) significant effect, 2 RCTs (n=110) no effect	
	Rhinorrhoea	n=660; 4 RCTs	Unclear effect on severity No pooling: 2 RCTs (n=369) significant effect, 2 RCT (n=291) no effect	
	Sneezing	n=574; 3 RCTs	Possible effect on severity No pooling: 3 RCTs significant effect	
			No pooling. 5 hors significant enect	
	Adverse events	n=842; 7 RCTs	aaaaaaaaaaaaaaaaaaabbaaaaaa baaaaaabaaaaaa	aa
	erenta		Pooled OR 1.58 (95% CI 0.78 to 3.21)	
Antihistaming.opplaceic.combinetion	Concession	n_1500.	Lineloor offeet on constitu	
Antihistamine-analgesic combination	Congestion	n=1508; 3 RCTs	aaaaaaaaaaaaaaaaaaaaaaaaaaaaaaaaaaaaaa	

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Table 2 (continued)

			Adults
Treatment option	Outcome	Included studies	Effect
	Sneezing	n=150; 1 RCT	Unclear effect on severity No pooling: significant effect on day 5
	Adverse events	n=1508; 3 RCTs	Possibly no increased risk No pooling: 3 RCTs no difference in AE
Analgesic-decongestant combination Cochrane review ¹¹	Congestion	n=1627; 5 RCTs	Possible effect on severity No pooling: 4 RCTs (n=1436) significant effect; 1 RCT (n=191) no effect

	Rhinorrhoea	n=679; 3 RCTs	Possibly no effect on severity No pooling: 3 RCTs no effect
	Sneezing	n=621; 2 RCTs	Possibly no effect on severity No pooling: 2 RCTs no effect
	Adverse events	n=1440; 5 RCTs	Increased risk Pooled OR 1.71 (95% CI 1.23 to 2.37; NNH 14)
Antihistamine-analgesic-decongestant combination Cochrane review ¹¹	Congestion	n=595; 3 RCTs	Possible effect on severity No pooling: 3 RCTs significant effect

Rhinorrhoea n=595; 3 RCTs Possible effect on severity No pooling: 3 RCTs significant effect

Sneezing	n=70; 1	Unclear effect on severity
	RCT	No pooling: no effect
Adverse	n=595; 3	Unclear risk
events	RCTs	No pooling: unclear if differences between groups

Congestion			
	reported		
Rhinorrhoea	Not		
	reported		
Sneezing	Not		
	reported		
Adverse	n=200; 1	Unclear risk	
events	RCTs	No pooling: no differences	
Congestion	n=1081;	Possibly no effect on severity	
	4 RCTs	No pooling: 4 RCTs no significant effect	
Rhinorrhoea	n=1959;	Possible effect on severity	
	4 RCTs	No pooling: 4 RCTs significant effect	
Sneezing	Not		
	reported		
Adverse	No RCTs	Increased risk	
events	available	Epistaxis: OR 3.21 (95% CI 1.68 to 6.13)	
		Nasal dryness: OR 2.55 (1.50 to 4.33)	
		Dry mouth: OR 3.59 (1.38 to 9.38)	
		Other AE: not significant	
Congestion	n=786; 1	Unclear effect on severity	
	RCT	No pooling: significant effect after 24 hrs	c
	Sneezing Adverse events Congestion Rhinorrhoea Sneezing Adverse events	reported Rhinorrhoea Not reported Sneezing Not reported Adverse n=200; 1 RCTs Congestion n=1081; 4 RCTs Rhinorrhoea n=1959; 4 RCTs Sneezing Not reported Adverse Not available Congestion n=786; 1	reported Rhinorrhoea Not reported Sneezing Not reported Adverse n=200; 1 Adverse n=200; 1 Vot No pooling: no differences Congestion n=1081; 4 RCTs No pooling: 4 RCTs no significant effect Rhinorrhoea n=1959; 4 RCTs No pooling: 4 RCTs no significant effect Sneezing Not reported Adverse No RCTs available Increased risk Epistaxis: OR 3.21 (95% CI 1.68 to 6.13) Nasal dryness: OR 2.55 (1.50 to 4.33) Dry mouth: OR 3.59 (1.38 to 9.38) Other AE: not significant Congestion n=786; 1 Unclear effect on severity

Table 2 (continued)

			A studies	
			Adults	<u> </u>
Treatment option	Outcome	Included studies	Effect	irst published
Systematic review14	Rhinorrhoea	n=786; 1	Unclear effect on severity	i le
		RCT	No pooling: significant effect after 24 hrs	0 20 20
	Sneezing	Not reported		8 IO. I
	Adverse	n=786; 1	Unclear risk	<u>.</u>
	events	RCT	Unclear risk No pooling: significantly higher incidence of blood-tinged mucus, epistaxis, nasal passage irritation, a with ipratropium bromide No effect on duration/persistence of purulent rhinitis	and nag
Antibiotics	Congestion	Not reported		.K370
Cochrane review ¹⁵	Rhinorrhoea	n=723; 4	No effect on duration/persistence of purulent rhinitis	0
		RCTs	Pooled RR 0.73 (95% CI 0.47 to 1.13)	OT
		n=227; 2	No effect on duration/persistence of clear rhinitis	0
		RCTs	Pooled RR 0.58 (95% Cl 0.23 to 1.48)	Cc
	Sneezing	No RCTs available		
	Adverse	n=1267;	Increased risk	2018.
	events	4 RCTs	Pooled RR 2.62 (95% Cl 1.32 to 5.18)	0. F

Vitamin C	Congestion,	Not	
Cochrane review ¹⁶	rhinorrhoea	reported	
	and		
	sneezing		aaaaaaaaaaaaaaaaaaaabbb
	Adverse	n=4556;	No increased risk
	events	7 RCTs	No pooling: no difference, nature of AE not reported
Echinacea	Congestion,	No RCTs	
Cochrane review ¹⁷	rhinorrhoea	available	
	and		
	sneezing		
	Adverse	n=1108;	Possibly no increased risk
	events	7 RCTs	No pooling: 1 RCT increased risk, 6 RCTs no difference, AE not reported

rlic	Congestion, No	ot		
chrane review ¹⁸	rhinorrhoea repor	ted		
	and sneezing			
	Adverse n=14	,	Unclear risk	
	events RC	T	No pooling: Unclear if different	
argonium sidoides extract	Congestion n=10	3; 1	Unclear effect on severity	
chrane review ¹⁹	RC	T	No pooling: Significant effect by day 5	
	Rhinorrhoea No	ot		
	repor	ted		
	Sneezing No	ot		
	repor	ted		
	Adverse n=10	3; 1	Unclear risk	
	events RC	т	No pooling: No difference in AE	•
	Adverse n=10	3; 1		

PRACTICE

			Adults
reatment option	Outcome	Included studies	Effect
Saline nasal irrigation Sochrane review ²⁰	Congestion	n=205; 2 RCTs	Possibly no effect on severity No pooling: 2 RTCs no effect on day 3
	Rhinorrhoea	No RCTs available	
	Sneezing	No RCTs available	
	Adverse events	n=143; 1 RCT	Unclear risk No pooling: no difference in AE
leated humidified air Cochrane review ²¹	Congestion, rhinorrhoea, and sneezing		
	Adverse events	n=203; 3 RCTs	Unclear risk No pooling: minor AE, unclear if different
/apour rub RCT ²²	Congestion	No RCTs available	
	Rhinorrhoea	No RCTs available	
	Sneezing	No RCTs available No RCTs	
	events	available	
	Congestion,	No RCTs available	

Table 2 (continued)

			Adults	irst
Treatment option	Outcome	Included studies	Effect	publis
	Adverse events	No RCTs available		hed as
				10
Antivirals ^{v 20} ,	Congestion,	No RCTs		1
zinc, ²⁴ - ²⁶ probiotics, ²⁷ - ²⁹ Chinese medicinal	rhinorrhoea,	available		36/bmj
herbs, ³⁰ honey, ³¹ eucalyptus oil, ²² fluid	sneezing			br
intake ^{vi 36}	and adverse			<u>, </u>
	events			N

"No effect" indicates that data were pooled and the overall effect estimate was not statistically significant. A "possible effect" is based on a qualitative appreciation of the effects reported size was based on what the authors reported and on the Cochrane Handbook (eg, a standardised mean difference of 0.2 to 0.49 represents a small, 0.5 to 0.79 a moderate, and 90.8 results were consistent, we concluded there was a possible effect or possibly no effect. Quality of evidence was based on the GRADE assessment reported in the review (indicated with review we assigned a GRADE assessment (see supplementary table); NSAIDs: non-steroidal anti-inflammatory drugs; GI: gastrointestinal; AE: adverse events; CI: confidence in loval; risk.

Positive scores represent treatment benefit;

This meta-analysis included one study that might have included children (n=60), although the age of the participants was not clear

It was not clear from the review which studies recorded adverse events, therefore we were not able to differentiate between adults and children

The review on antivirals has been withdrawn, no new updated Cochrane review has been published

Whe review on fluid intake did not identify any relevant trials

Figure

ADULTS	EFFICACY			HARM	
	Congestion	Rhinorrhoea	Sneezing	Risk of harm	Reported adverse events
Analgesic-decongestant combination	**	*	*		Sedation, insomnia, nervousness, palpitations, light headednes Gl symptoms, dizziness, dry mouth, headache, fever
Antihistamine-analgesic-decongestant combination	**	**			Sedation, insomnia, dizziness, palpitations, headache and GI symptoms
Antihistamine-decongestant combination			**		Sedation, dry mouth, insomnia, GI symptoms, dizziness, palpitations, nervousness and headache
Decongestants					Insomnia and headache
Intranasal ipratropium bromide	*	**			Nosebleeds, dryness of nose/eyes/mouth, nasal irritation, headache, tachycardia
NSAIDs					Drowsiness, GI symptoms, rash, oedema, hyperactivity, flushed face, insomnia, light-headedness, dry mouth
Sedating antihistamines					Sedation, GI symptoms, insomnia, dry nose, headache, dizziness and dry mouth
Non-sedating antihistamines			*		Sedation, GI symptoms, insomnia, dry nose, headache, dizziness and dry mouth
Antihistamine-analgesic combination				***	Nasal dryness/irritation, drowsiness, dizziness dry mouth, Gl symptoms, appetite loss, headache, depression, epistaxis
Decongestant - ipratropium bromide combination					Blood-tinged mucus, epistaxis, nasal dryness/irritation, headache, dry throat/mouth, nausea, throat irritation
Paracetamol/Acetaminophen					Sweating and GI symptoms
Pelargonium sidoides extract					GI symptoms and allergic skin reactions
Saline nasal irrigation	*				Dry nose, nasal pain/irritation
Antibiotics					GI symptoms
Vitamin C					Unclear which adverse events occurred
Echinacea				***	Unclear which adverse events occurred
Garlic; Heated humidified air; intranasal corticosteroids					Garlic: rash and odour; Heated air: nasal/lip irritation, light-headedness, congestion, discomfort of mask

Vapour rub, Zinc

s,	No effect /*possibly no effect
	Small effect/ **possible small e
	Unclear

Not reported

lo increased risk / **possibly no increased risk

CHILDREN EFFICACY HARM Congestion Rhinorrhoea Sneezing Risk of harm Reported adverse events ** Difficulty tolerating delivery of nasal drops, nosebleeds Saline nasal irrigation Analgesic-decongestant combination Antihistamine-analgesic-decongestant combination No adverse events occurred Sedating antihistamines Sedation Increased sleepiness Antihistamine-decongestant combination Non-sedating antihistamines No adverse events occurred Antibiotics GI symptoms Skin rash/redness, burning of skin/nose/eyes, hyperactivity, Vapour rub sleepiness, headache Echinacea - Ginseng Echinacea: skin rash: ginseng: no AE occurred Honey No trials available for: Antivirals¹, Chinese medicinal herbs, Decongestants in monotherapy, Decongestant-ipratropium Small effect/ **possible small effect No increased risk /***possibly no increased risk Unclear Unclear bromide combination, Ipratropium bromide, Eucalyptus oil, Not reported

Fluid intake², Garlic, Heated humidified air, intranasal corticosteroids, NSAIDs, Paracetamol, Pelargonium

sidoides extract, Probiotics, Vitamin C, Zinc

Fig. 1 Benefit and harm of common cold treatments in adults and children: a summary based on analysis of the evidence from Cochrane reviews and clinical trial. "No effect" indicates that data were pooled and the overall effect estimate was not statistically significant. A "possible" effect is based on a qualitative appreciation of the effects reported in individual trials that could not be pooled. Interpretation of the size of the effect was based on what the authors reported and on the Cochrane Handbook (eg, a standard mean difference of 0.2 to 0.49 represents a small, 0.5 to 0.79 a moderate, and ≥0.8 a large clinical effect). If no pooling was available but results were consistent, we concluded there

[📒] Unclear

was a possible effect or possibly no effect. * The Cochrane review on antivirals has been withdrawn, no new updated Cochrane review has been published. [†] The Cochrane review on fluid intake did not identify any relevant trials