

# Acute Otitis Media & Other Respiratory Tract Infections

Summer / Fall 2024

# Did you know?

- Firstline is a free, localized, pointof-care antimicrobial stewardship app / website with guidance for pediatrics (IPCH) or adults & penicillin allergy de-labeling (SHA).<sup>1,2</sup> See back page.
- Analgesics are sometimes underdosed for pain/fever in respiratory tract infections. Expert Advise caregivers to use weight-based, scheduled dosing for the first 48 hours while awake and then PRN.
- Infection-related complications are rare even without an antibiotic:

3 Thompson'09, 4 Peterson'07

- mastoiditis: <3.8 events per 10,000 **acute otitis media** cases.
- peritonsillar abscess (quinsy), 0-15 years: <6 events, ≥16 years: <21 events per 10,000 **pharyngitis** cases.
- pneumonia, <65 years: <126 events per 10.000 **bronchitis** cases.
- Reserve amox/clav CLAVULIN, g for AOM or sinusitis treatment failure.

  5 CPS 16,6 CSO-HNS 11 If prescribing, strive for clavulanate ≤ 10 mg/kg/day in pediatrics & 125 mg/dose in adults to decrease adverse events e.g. diarrhea. 7-9
- Macrolide *S. pneumoniae* resistance is increasing ~21% <sup>10 Saskatoon 23</sup> ~38% <sup>11</sup> Regina <sup>23</sup>. Reserve use for AOM or sinusitis in those with a <u>severe</u> penicillin allergy. <sup>5,6</sup> >80% of people outgrow their penicillin allergy after 10 years, even if severe. <sup>12 Blumenthal 19</sup>
- Group A Streptococcus (GAS), the main **pharyngitis** pathogen of concern, is **100%** susceptible to penicillin in SK.<sup>10,11,13</sup> CPS<sup>21</sup> Prescribe **penicillin V PEN-VK**, **g in those able to swallow tablets** (scored) as narrower spectrum than amoxicillin.

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www.RxFiles.ca

In Canada, SK is the jurisdiction with the third highest antimicrobial use per capita. 14 PHAC'21

- ~90% of all antibiotic prescriptions are dispensed in the community sector (Canada). 14 PHAC'21
- ~30% of all outpatient antibiotic prescriptions are not required (USA). 15 Fleming-Dutra'16

  Overuse leads to harms e.g. by 2050, the UN projects there will be as many deaths due to antimicrobial resistance as there are currently deaths due to cancer (> 10 million/year). 16-19

Patient education can provide reassurance on the self-limiting nature & usual course of infections to support watchful waiting / "wait & see" strategies. 20,21 This approach found: 22 Cochrane'23, 23 Mas-Dalmau'21

- · similar duration or severity of symptoms as with antibiotic use
- · maintained parental satisfaction (>90%)
- · less antibiotic use vs immediate antibiotic prescription (\$\sigma^65-85\%)
- fewer adverse events e.g. diarrhea NNH=17 / 30 days



# Optimize Acute Otitis Media (AOM) Management

~2 out of 3 cases resolve without antibiotics at 72 hrs. 24 Marchetti 105 If needed, amoxicillin AMOXIL, g remains 1st line & is preferred over amox/clav per local SK antibiogram. 10,11

# COMMON QUESTIONS RELATED TO AMOXICILLIN DOSING IN ACUTE OTITIS MEDIA:

When to use "high-" vs "standard-dose" amoxicillin?

Penicillin resistant *S. pneumonaie* (PRSP) rates are currently low, SK ~0% to <1.3%.<sup>10,11</sup> Recommend high-dose in select patients with PRSP risk factors: received an antibiotic in the previous 3 mos, or attending daycare, or <2 years, or unimmunized/underimmunized.<sup>5 CPS'16, Bugs & Drugs'23, JPCH, Expert</sup>

What is the maximum dose for amoxicillin in pediatrics?

The usual dose is 4 g/day.<sup>29-31</sup> For adolescents ≤15 years, may use up to 4 grams per day (pediatric max dosing) due to increased renal elimination of amoxicillin.<sup>Expert</sup>

How frequently should amoxicillin be dosed throughout a 24 hour period?<sup>32, Expert</sup>

To maintain adequate middle ear drug levels, divide **TID** if prescribing **standard-dose** amoxicillin. If prescribing **high-dose** amoxicillin, may divide **BID** as higher drug levels are achieved.

When should I prescribe amoxicillin 5 days vs 10 days?

Children ≥2 years: 5 days adequate and fewer AEs NNH=29 vs 10 days (even if risk factors for PRSP). 33 Cochrane 10 Children <2 years: 10 days better than 5 days. 34 Hoberman 16

# Reassess Penicillin/Amoxicillin Allergy

≥90% of individuals reporting a *penicillin allergy* turn out to actually be beta-lactam tolerant.<sup>35</sup> AAAAI'¹0 False allergy labels lead to ↑risk of poor outcomes.<sup>36-39</sup> Risk-stratify & de-label when possible!<sup>40,41</sup>

# History of:

- a) predictable adverse effects e.g. GI upset, headache, or
- b) family member with allergy, orc) same drug taken again without reaction.

NO true allergy risk.<sup>44</sup> Direct de-labeling suitable. The low hanging fruit! {Avoid false labeling!} If <u>Firstline</u><sup>42</sup> app, or <u>PEN-FAST</u><sup>43</sup> suggest very-low, or low-risk. e.g. history of <u>non-severe delayed</u> reaction (rash with no systemic or blistering/mucous membrane symptoms).

blistering/mucous membrane symptoms),

>5-10 years since reaction
(childhood rash)

Low-risk & may be suitable for direct de-labeling or Direct Oral Challenge\*

(e.g. adults: amoxicillin 500mg x 1 dose; monitor x 1 hour) Immediate (vs delayed)
Severe (vs non-severe)
(e.g. anaphylaxis, angioedema; systemic reaction, blistering of skin/mucous membranes; treatment required)

Recent (vs >5-10 years)

Moderate to high-risk Identify those with history of true IgE allergy or severe reaction. Avoid de-labeling; (may direct challenge, or refer to allergist).

\*Systematic review (N=13, n=1202) of direct oral challenge: ~96% of low-risk individuals were successfully de-labeled; ~4% had a mild immediate or delayed reaction; no reports of serious adverse events. 45 Cooper 21

- Treat the ear pain. Calculate weight-based analgesia & schedule for the first 48 hours.
- Watchful waiting for up to 48 hours is recommended for most children ≥6 months old.
- If antibiotics are required, amoxicillin is the empiric drug of choice in most cases.
- In uncomplicated AOM, children ≥2 years can usually be treated with 5 days of antibiotic therapy, however, children <2 years of age should be treated for 10 days.
- Effusion does not mean tx failure. It is sterile fluid & can take up to 3mos to resolve.
- Patient Tools:<sup>7-9</sup> Mayo Clinic interactive tool, CDC handout, Choosing Wisely handout

# Overview

- No gold standard for diagnosis & it is especially challenging in young children, leading to overdiagnosis/overtreatment. Often post viral URTI.
- Diagnosis: 5 CPS'16 Acute onset (<48hours) of symptoms e.g. otalgia or suspected otalgia (sx may be non-specific in non-verbal children) AND</li> 2 Middle ear effusion AND 3 Inflammation: bulging tympanic membrane OR perforated tympanic membrane with discharge (i.e. otorrhea)
- Etiology: viral or bacterial; difficult to distinguish between & both are often present at the same time (up to ~65% of pts). S. pneumoniae (more virulent, ~20% spontaneously resolve), H. influenzae non-typeable or M. catarrhalis (less virulent, ~50-75% spontaneously resolve), GAS (uncommon, linked perforation). 20-24
- Some evidence suggests routine childhood immunizations (e.g. pneumococcal conjugate PREVNAR-13, H. influenza e type b, influenza vaccine) may ↓ episodes of AOM & alter pathogen prevalence e.g. ↓ S. pneumoniae pneumococcal vaccine serotypes & ↑ H. influenzae (non-typeable). <sup>25-28</sup>

# An Approach to Treatment

Treat the ear pain (whether or not an antibiotic is prescribed, schedule analgesia for the first 48 hours) 29 Cochrane (23 (4 RCTs, 411 children; low quality evidence)

see RxFiles Pediatric Pain Chart page 139, caregiver dosing infographic

- Acetaminophen TYLENOL, g 10-15mg/kg q4-6h PRN (max=75 mg/kg/d term neonate)
- Either acetaminophen or NSAID provided pain relief at 48hr ± antibiotic (NNT=6-7/48hr, analgesic ~90% vs placebo ~75%). 29 Cochrane 23 (4 RCTs, 411 children: low quality) NSAIDs e.g. ibuprofen ADVIL, g 5-10 mg/kg q6-8h PRN (max=40mg/kg/d limited data <amous / limited evidence of additional benefit when routinely combining or alternating. May lead to confusion/dosing errors; may suggest select patients. Experi
- Possible benefit: ?topical anaesthetics POLYSPORIN Plus Pain Relief ear drops, orc >6/yrs \$20/15ml; lidocaine, polymyxin (≥3yr: ↓pain for 10-30min; At not well studied; Ci perforation). 30 Cochrane '06, NICE (consider during watchful waiting) If used, give in addition to po analgesics Anecdotal benefit (only if no eardrum perforation): hot/cold compress; oil instillation e.g. mineral, olive. Some recommend against. If used, give in addition to oral analgesics.

Not recommended: antihistamines (no benefit, ?^AE, ?^effusion duration), decongestants (no benefit, ^AE), nasal or systemic corticosteroids (largely unstudied, low quality evidence). 31,32 Cochrane '08 8'18'

- Determine if watchful waiting / "wait & see" is appropriate (the usual course of AOM is ~3 days, but can last up to 1 week<sup>3 NICE</sup> and most do not require immediate antibiotic see Table 1 below). 33 Choosing Wisely Primary Care, Emerg Medicine • Watchful waiting: antibiotics are withheld & treatment is limited to analgesia & caregiver observation for up to
  - 48 hours 5 CPS'16 ). 3 NICE'228'18 / 4 MUMS'19 / 6 AAP'13 up to 72hour This should be considered in healthy children ≥6 months. - Caregiver may be instructed to return for reassessment or given an antibiotic e.g. "wait & see" / delayed
  - prescription if the child does not improve or worsens at any time during the watchful waiting period.34 Cochrane'23
- In eligible children, watchful waiting may \u2211 antibiotic use (n=1277, ~66% recovered without an antibiotic @72hrs & 30 days). 35 Marchetti'05
- Whether children receive antibiotic or placebo, ~60% had ear pain resolution at 24hrs. Most had resolution at 48hrs (NNT≈20, ~90% antibiotic vs ~85% placebo), but some are more likely to benefit from antibiotics (<2 years with bilateral AOM, otorrhea: NNT≈3-4, 70-76% vs 40-45%). ↑N/V/rash with antibiotic vs placebo (NNH≈14, 27% vs 20%). 36 Cochrane 23 (13 RCTs, 3401 children) Infection-related complications are rare (e.g. <3.8 events per 10,000 AOM cases). 37,38 ↑ resistance with ↑ antibiotic use.

# Table 1. Watchful Waiting is NOT appropriate in the Following Patients<sup>1-6</sup>

- <6 months
- Perforated tympanic membrane
- Otorrhea i.e. ear discharge (any age)
- Moderate to severe presentation
  - Severe otalgia, poor response to antipyretics OR
  - Symptoms do not improve after ≥48hrs OR
  - Temperature ≥39°C (with or without antipyretic use)
- Tympanostomy tubes, cochlear implants
- Recurrent AOM (see below)
- <2 years with bilateral AOM
- Medical co-morbidities (e.g. craniofacial abnormalities, immunodeficiency)
- Caregiver unable to support child or follow-up

#### Table 2. Empiric Drug Regimens<sup>1-6</sup> \* see Antibiotic Formulations Chart page 7 for price, etc. If watchful waiting is not appropriate (table 1) or no improvement/worsening of signs ± symptoms after >48 hours. • Amoxicillin is the drug of choice for AOM. Rates of penicillin Standard-dose (SD) Amoxicillin © resistant S. pneumoniae (PRSP) are low (SK<sup>2023</sup><0 - 1.3%).<sup>39,40</sup> 40 to 50mg/kg/day AMOXIL, g Consider high-dose option if risk of PRSP e.g. daycare. ÷ TID **Moxilen Forte** under or unimmunized, <2 years, or antibiotic within 3 months. High-dose (HD) AE: no head-to-head, appear similar; 36 Cochrane' 23 Meta-analysis: 41 diarrhea SD Max total daily dose ~9% vs HD ~14%, rash SD ~3% vs HD ~6.5% <mark>overlapping CI, 个heterogeneity</mark>.41 4g/day 80 to 90mg/kg/day • Amox resistant to beta-lactamase positive *H. influenzae* (20% SK 2023) & ©= tastes good ÷ BID (may give TIDSanfords) M. catarrhalis (most strains) but these likely to spontaneously resolve. PENICILLIN ALLERGY: NON-SEVERE (e.g. delayed >72 hours rash) see RxFiles Beta-lactam Allergy Chart page 6, SHA Firstline. 15 to 30 mg/kg/day Cefprozil • Broader spectrum vs amox (greater selective pressure on other bacteria) ÷ BID-TID CEFZIL, g © • Cefprozil suspension is ~\$5-10 less compared to cefuroxime. • Cefuroxime: increase bioavailability by dividing total daily dose 20 to 30mg/kg/day Cefuroxime TID vs BID, or giving with food.42 CEFTIN, g ☺ ÷ BID-TID (cc preferred) PENICILLIN ALLERGY: SEVERE (e.g. anaphylaxis, angioedema) see RxFiles Beta-lactam Allergy Chart page 6, SHA Firstline

#### Clarithromycin 15 mg/kg/day ÷ BID BIAXIN, g Azithromycin 10mg/kg Day 1, then ZITHROMAX, g 5mg/kg Days 2-5

2 to 5mg/kg/day po

Doxycycline

DOXYCIN, g

- Avoid use. ↑ S. pneumoniae resistance ~21%<sup>39 Saskatoon'23</sup> to ~38%<sup>40</sup> Regina'23. If used, clarithromycin preferred as potentially less resistance and major **DI** uncommon in pediatrics.<sup>43,44</sup>
- Doxy: lack tooth staining <8yrs, may use ≤21d;<sup>45 AAP'21, 46 Ravindra'23</sup> admin: no commercially available suspension, see Geri-RxFiles. capsule (may open & mix applesauce) and tablet (may crush).
- Other alternative: clindamycin 20-30mg/kg/day ÷ TID. \*potential exceptions i.e. use amoxicillin-clavulanate 1st line or 2nd/3rd generation cephalosporin e.g. cefprozil, ceftriaxone if: amoxicillin in previous 30 days, history of AOM unresponsive to amoxicillin, immunocompromised. <sup>2 Bugs & Drugs</sup> <sup>23, 5 CPS</sup> <sup>16</sup> (<sup>22), 6 AAP</sup> <sup>13</sup>
- Controversy: concurrent purulent conjunctivitis (some suggest more likely to be due to H. influenzae or M. catarrhalis [may produce beta-lactamases  $\rightarrow$  amoxicillin resistance]<sup>5 CPS</sup> others suggest still likely to be *S. pneumoniae* and amoxicillin 1st line adequate<sup>Expert</sup>).

Treatment Duration: (M: response to antibiotic usually within 2-3 days of antibiotic treatment) 6 AAP'13 Antibiotic x 5 days: uncomplicated / healthy children ≥2vrs. 1 Sanford'24, 3 NICE'228/18, 5 CPS'16 ('22), 6 AAP'13, 47 Cochrane'10 (49 RCTs, n=12,045) Antibiotic x 10 days: <2 years of age, see RxFiles Trial Summary: treatment in children (6-23 months)<sup>48 Hoberman'16</sup> or AOM with perforated tympanic membrane, treatment failure, recurrent AOM. 5 CPS'16('22)

Table 3. Managing Treatment Failure<sup>1-6</sup> (no symptomatic improvement after ~2-3 days of antibiotics) Middle ear effusion does not equal treatment failure, ~60-70% of individuals will have persistent middle ear effusion after an episode of acute otitis media and ~90% of all cases will self-resolve by 3 months. 49,50

Amoxicillin	in High-dose regimen (see left for dosing) • Use if failure of standard-dose amoxicilling			
Amoxicillin/ Clavulanate CLAVULIN, g >3mos: do not use 4:1 product †volume, AEs  Dose listed as amoxicillin component	If failure of high-dose amoxicillin: Standard-dose: Amox/clav 7:1 ratio 45 mg/kg/day ÷ TID x 10 days cc • Standard-dose amox/clav provides coverage of beta-lactamase positive H. influenza & M. catarrhalis without excessive clav (associated with ↑diarrhea).51	Reserve when able: broader spectrum & ↑AE vs amox e.g. diarrhea (RR 1.15), yeast infection (RR 1.33).51 Some suggest high-dose amox/clav for all failures as high-dose amoxicillin (e.g. 90mg/kg/day) overcomes some PRSP.¹Sanfords'24,6AAP'13 In Canada, requires 2 prescriptions to create 14:1 ratio which limits total clav dose (↑diarrhea >10mg/kg/day clavulanate).52,53 (USA: 1 prescription as 14:1 ratio commercially available) - {		
Ceftriaxone ROCEPHIN, g x ▼	50mg/kg IM/IV daily x <b>3 days</b>	Role: unable to tolerate oral, fail amoxicillin/clav.  If IM: follow local policies e.g. dilute/reconstitute with lidocaine 1% (& may buffer) to ↓pain, ≥1 injection if ↑volume. Expert  May also consider cefprozil or cefuroxime, see left.		

ENT referral: persistent tx failure ?tympanocentesis for culture of middle ear fluid, or middle ear effusion >3 months.

Managing Recurrence AOM incidence decreases with increasing age

• Definition: ≥3 AOM episodes within 6 months or ≥4 episodes in 12 months (with ≥ 1 episode in the past 6 months) <sup>2</sup> Bugs & Drugs'23, 5 CPS'16 ('22), 6 AAP'13 No indication for prophylactic antibiotic as not effective & ↑resistance. <sup>1,54,55</sup>

No antibiotic in the past 4-6 weeks	Usually amoxicillin standard or high dose x <b>10 days</b> (see left for dosing).
Antibiotic in the past 4-6 weeks	Usually amox/clav x 10 days (see managing treatment failure above).

- Consider ENT referral as tympanostomy tubes may be explored.
- Prevention strategies: handwashing, breastfeeding (exclusively for 6 months), smoke avoidance, vaccination (pneumococcal, annual influenza, H. influenzae type b), ?xylitol (gum, lozenge available OTC ~\$10-20/month).54

Note: AOM is uncommon in adults, if required, 1st line empiric: amoxicillin 500-1000mg TID x 5 days. 2 Bugs & Drugs'23, Expert

- Most cases do NOT require antibiotics as most infections are viral (~98% adult, ≥90% children) and self-limiting.
- Viral and bacterial sinusitis have similar symptoms.
  - Consider watchful waiting and advise on symptomatic treatment: analgesics, saline nasal drops/rinses, warm facial packs, decongestants, & intranasal corticosteroids.
- Patients should see a healthcare provider if symptoms worsen after initial improvement or are prolonged (take >10 days to improve) as potential bacterial etiology.
- If required, amoxicillin is the antibiotic of choice for bacterial sinusitis. Avoid macrolides if possible due to increasing resistance. See RxFiles Beta-lactam allergy Chart or SHA Firstline.

## Definitions / Overview

- Acute uncomplicated: inflammation of the paranasal sinuses (sinusitis) & nasal mucosa (rhinosinusitis) for <4 weeks usually in the setting of a preceding viral URTI.</li> - ~12% of adults/yr develop acute sinusitis<sup>15</sup> (females ↑~1.5x vs males; ↑45-64yrs ~15%)¹⁶ and ~7% of pediatrics/yr (more common ≥2yrs, ~50% concurrent AOM).¹¹⁻¹¹¹
  - Most cases are viral e.g. rhinovirus, influenza, adenovirus, RSV, parainfluenza virus; however, some bacterial (adults: 20~0.5-2% & peds: 21,22 ~5-10%) S. pneumoniae, non-typeable H. influenzae, M. catarrhalis (more common in children), other less common: S. pyogenes (GAS), S. aureus, local anaerobes (associated dental infections).
- Diagnosis is clinical and imaging is not recommended. 1-8,23 Distinguish bacterial from viral sinusitis based on symptoms & their duration. be used to diagnose bacterial sinusitis (indicative of inflammation, but not of bacteria and may be present during viral). 24 Shalikh'23, 25 The following suggests bacterial: 1-8,23
  - Presence of symptoms: purulent nasal discharge AND nasal obstruction OR facial pain-pressure-fullness that usually worsens when bending forward AND
  - ② Duration of symptoms: persistent, without improvement ≥10 days OR sudden onset of worsening sx within 10 days after initial improvement "double sickening".
  - Acute viral sinusitis symptoms tend to improve within 7-10 days. 5.7.8,26 Gwaltney 67 There is a 60% risk of bacterial sinusitis if symptoms >10 days. 27 Gwaltney 92 Note: some suggest bacterial rhinosinusitis if purulent nasal discharge & severe symptoms e.g. fever ≥39°C x 3-4 days at beginning of illness.7 IDSA
- Acute complicated: symptoms extend outside the paranasal sinuses & nasal cavity e.g. neurologic, ophthalmologic, surrounding soft tissue. 9:14,23 Symptoms for Urgent Referral to Emergency (imaging usually required): systemic toxicity; altered mental status; severe headache; swelling of the orbit or vision changes.9-14,23

# An Approach to Treatment (acute uncomplicated) 1-8

- Watchful waiting should be considered (provide education and symptomatic care, see Table 1 below):
  - Present with symptoms that have not worsened OR have had symptoms for less than 10 days, and
  - Confidence in patient's ability for follow-up (e.g. discuss antibiotic if symptoms fail to improve after 7-10d or worsen at any time).
- Patient tools: Infographic (RxFiles, Choosing Wisely), Viral Prescription Pad (adult, pediatric, delayed).
- Most acute bacterial sinusitis cases are self-limiting (low risk of complications even without an antibiotic) and improve within 2 weeks without antibiotics. Whether treated with an antibiotic or not:
  - Complications e.g. orbital, intracranial or soft tissue infections appear to be rare (<0.1%) in outpatients. 50,51
  - ~50% of adults will improve by ~7-10 days & ~70% within 2wks. 50 Cochrane' 18 (antibiotics clinical improvement: adults 8 RCTs, n=1687: NNT=20/1-2wks (60% antibiotic vs 55% placebo);<sup>50</sup> peds 4 RCTs, n=382: OR 2 (95% CI 1.16-3.47) vs placebo);<sup>51</sup>
- It is unclear whether antibiotics shorten the duration of illness (trials report no reduction up to a 2 day reduction).
- \underset AE with antibiotics: adults, NNH=9 / 1-2wks (7 RCTs: 27% vs 15% placebo); 50 peds 4 RCTs ~15% vs ~8% placebo. 51

# **Table 1. Symptom Management**

see RxFiles OTC Products Chart page 215 for more details Non-pharmacologic: Limited or lack of evidence to support use. Anecdotally, may help for some e.g. ↑mucus drainage

- Saline nasal drops/sprays/irrigation X ▼ OTC 29-34 150mL hypertonic saline nasal irrigation e.g. NETI POT daily
- Steam inhalation, humidifier<sup>28 (ensure cleaning</sup> Warm facial packs
- Saline: improve QOL/symptoms & ↓use OTC meds? Most data in recurrent or chronic sinusitis. Use sterile/bottled H<sub>2</sub>O.
- If used with intranasal corticosteroid, admin saline first. - SALINEX X ▼ \$13/125mL <2vrs: 4 drops 1-6x/d PRN, Adults: 1 spray 2-6x/d PRN

Analgesics: Limited or lack of evidence to support use. Anecdotally, may provide pain relief for some.

**Ibuprofen ADVIL**, g ▼ **OTC** (suspension X ▼) Peds: 5-10 mg/kg q6-8hr PRN (max 40mg/kg/day)

Peds: 10-15mg/kg q4-6hr PRN (max 75mg/kg/day) Adult: 1000mg q4-6hr PRN (max 4g/day) ~\$10/120tab

Acetaminophen TYLENOL, g X ▼ OTC

Intranasal Corticosteroids: Some suggest as an option e.g. mild-moderate and symptoms >7-10d.3.4,7,8,38-40 Off-label.

Fluticasone FLONASE, g X ▼ \$23/120 sprays OTC: ≥18yrs ≥4yrs: 1 spray (50mcg) in each nostril once daily Adults: 2 sprays (100mcg) in each nostril once daily Mometasone NASONEX, g \$22/140 sprays

Adults: 400mg q6-8hr PRN (max 2.4-3.2g/day) ~\$10/60tab

≥3yrs: 1 spray (50mcg) in each nostril once daily Adults: 2-4 sprays (100-200mcg) in each nostril daily to BID

See RxFiles Intranasal Corticosteroid Chart pg 40 for more options.

≥12yrs, adults: 1-3 drops/sprays per nostril TID PRN

Pseudoephedrine e.g. ELTOR X OTC \$5/12tab many @

≥12yrs, adults: 60mg po q4-6hr PRN (max 240mg/day)

6-11yrs: 30mg po q4-6hr PRN (max 120mg/day)

- Some suggest monotherapy<sup>3 MUM, 4 NICE, 8 CSO-HNS</sup> or in combo to antibiotics. 3 MUM, 4 NICE, 7 IDSA More useful if allergies. Expert
- A 2013 Cochrane review of intranasal corticosteroid vs placebo (3 RCTs, n=1792): symptom resolution/improvement 73% vs 66% NNT=15/2-3wks.  $^{35}$  May  $\downarrow$  symptoms by  $^{\sim}$  3.5 days (6 days vs 9.5 days). 36 Dolor'01 ?dose response. 37 Meltzer'05, 4 NICE
- AE: epistaxis (use contralateral technique), 41 nasal itching/irritation. - Alt: RHINOCORT AQUA \$22/120 sprays; NASACORT X ▼ OTC: ≥12yrs \$21/120 spray

Decongestants: Limited / lack of evidence to support use. 42 Cochrane 14,43 Some suggest use, 1-4,8 while others do not. 7 IDSA Xylometazoline e.g. OTRIVIN X ▼ OTC \$8/20mL - Some guidelines prefer topical over oral preparations. 1,4,44

- nasal: less systemic absorption and less adverse events.
- po: <u>Caution</u>: CVD, BPH, diabetes; <u>AE</u>: insomnia, ↑glucose ↑BP. Phenylephrine nasal/po not recommended. 45 FDA'23
- Most guidelines recommend short-term use e.g. < 3 to 5 days. Prolonged nasal use can lead to rebound congestion.
- Oral Corticosteroids: most guidelines do not address or do not recommend for acute sinusitis.
- If use, reserve for severe sinusitis in adults. Cochrane 2014 po steroid vs placebo/NSAID (5 RCTs, n=1043 acute sinusitis): symptom resolution/improvement 70% vs 54% but limitations e.g. heterogeneity. No benefit without concomitant antibiotic. 48,49 Dosing e.g. prednisone 30 to 60mg po daily x 3-7 days (then stop, no taper required).
- Not recommended: antihistamines (unless allergic component), 1-4,8 mucolytics i.e. guaifenesin. 2,4

Table 2. Empiric Drug Regimens<sup>1-8,50-62</sup> see RxFiles Oral & IV Antibiotics Char page 81 for price, etc. MILD (symptoms <10 days or no worsening in symptoms) No antibiotic Mostly viral. See Table 1 (left). Tools: RxFiles, Choosing Wisely, Viral Rx Pad (adult, pediatric, delayed). MODERATE to SEVERE (symptoms ≥10 days or symptoms worsens after 5-7 days "double sickening") Peds: Standard-dose 40-45mg/kg/day ÷ TID x 7-10 days Consider high-dose if ↑risk of Amoxicillin © penicillin resistant PRSP e.g. **High-dose** 90mg/kg/day ÷ BID x 7-10 days AMOXIL, g antibiotic within 3 months, daycare. Adults: Standard-dose 500mg TID x 5 - 7 - 10 days\* **Moxilen Forte** <2 years, under / unimmunized. High-dose 1000mg TID x 5 - 7 - 10 days\* High-dose covers most PRSP. PENICILLIN ALLERGY: NON-SEVERE (e.g. delayed >72 hours rash) see Beta-lact

PENICILLIN ALLERGY: NON-SEVERE (e.g. delayed >72 hours rash) see Beta-lactam Allergy Chart page 86, SHA Firstline.				
Cefuroxime	Peds: 20-30mg/kg/day ÷ BID-TID x 7-10 days (max 1000mg/day)	- Some no longer recommend		
CEFTIN, g ☺	Adults: 250mg to 500mg BID x 5 - 7 - 10 days* (cc preferred)	(↑resistance <sup>2 Bugs&amp;Drugs</sup> ), but		
Cefprozil	Peds: 15-30mg/kg/day ÷ BID-TID x 7-10 days (max 500mg/day)	SK, high susceptibility (~98% S.		
CEFZIL, g ☺	Adults: 250mg to 500mg BID x <b>5</b> - 7 - 10 days*	pneumoniae, ~95% H. influenzae).		

pneumoniae, ~95% H. influenzae). PENICILLIN ALLERGY: SEVERE (e.g. anaphylaxis, angioedema) see Beta-lactam Allergy Chart page 86, SHA Firstline.

Peds:\* 4mg/kg/day ÷ BID (max 200mg/day) x 7-10 days Doxycycline \*Lack of tooth staining <8yrs. Ravindra'23 May use <8yrs short duration ≤21d. AAP'21 No commercial suspension, capsules may be opened & mixed with applesauce. DOXYCIN, g Adults: 200mg day 1, then 100mg BID x 5 - 7 - 10 days\*

peds, but \resistance in SK & other. If required, dosing pg 81. Table 3. Managing Treatment Failure 1-8,50-62 no symptomatic improvement in ~3-5 days of antibiotics -If failure of standard-dose. Exp High-dose amoxicillin regimen (see above for dosing)

**Amoxicillin** -CLAVULIN: broader vs Amoxicillin / If failure of high-dose amoxicillin: amoxicillin, including beta-Standard-dose Amox/clav<sup>7:1 ratio</sup> 45mg/kg/dav ÷ TID x 7-10d Clavulanate lactamase positive H. CLAVULIN. g Adults: **Standard-dose** 500mg 4:1 ratio TID x **5** - 7 - 10 days\* **or** influenzae & M. catarrhalis. Dose listed as 875mg 7:1 ratio BID x 5 - 7 - 10 days\* -1st line empirically (highamoxicillin dose) immunocompromised. component Admin: preferred to take with food to ↑tolerability -Some suggest high-dose & ↑clavulanate absorption. 65-68 amox/clav for all failures, pg 82.

\*5 days of therapy should be sufficient in uncomplicated, otherwise healthy, adults with less risk of harm.

- Meta-analysis (12 RCTs, n=4430) found no difference in clinical success (cure or improvement of symptoms) with short courses (3 to 7 days) versus longer courses (6 to 10 days) of the same antibiotic. Sensitivity analysis (7 RCTs, n=2715) comparing 5 versus 10 days did not find a difference in clinical success either. 63 Falagas'09
- Older patients with comorbidities were excluded from the trials, and therefore we do not have evidence to support a shorter course of therapy in this population. 63 Falagas'09

## When should patients with sinusitis be referred to a specialist?9-14

- Recurrent Sinusitis: ≥4 episodes of acute bacterial sinusitis/yr. Consider assessment for allergies, immunologic deficiency, or surgery. Neither antibiotics nor intranasal steroids have shown a decrease in recurrent sinusitis.
- Chronic Sinusitis: >12wks of symptoms and evidence on physical exam (e.g. anterior rhinoscopy) or radiography. Ask about OTC topical nasal decongestants long-term use can lead to rebound congestion.
  - Consider intranasal corticosteroids ± saline irrigation (if used together, do saline irrigation first). 64 AFP'17(A)
  - Antibiotics are not recommended, but may consider if acute exacerbation. 64 AFP'17(C)
  - Consider Ears/Nose/Throat specialist referral if above measures fail after 3 months.

- Macrolides: an option e.g.

- Pharyngitis is usually self-limiting (4-5d; up to ≤10d); most cases do <u>NOT</u> require antibiotics as they are viral infections (80-90% in adults, >70% in children).
- Scoring systems e.g. modified Centor score, <u>FeverPAIN</u> can help identify low risk patients who do not require diagnostic testing or antibiotics.
- For confirmed Group A Streptococcus (GAS)
   pharyngitis, penicillin x10d (started within 9 days of
   symptom onset) is 1st line. There is no documented
   GAS resistance to penicillin. Consider stewardship
   strategies e.g. delayed antibiotic pending throat swab results.
- Advise on treatments for symptomatic relief: e.g. NSAIDs, acetaminophen, lozenges, topical anesthetics, warm liquids, saltwater gargle.
- Patients should see their prescriber if: 1 symptoms worsen, 2 symptoms take longer than 4 to 5 days to resolve, &/or 3 unilateral neck swelling, shortness of breath or severe dysphagia develops (r/o abscess).

### Overview

- Etiology: viral 80-90% of adults (>70% of children); therefore, the majority do NOT require antibiotics; minority bacterial Group A Streptococcus (GAS); rarely other bacteria (e.g. Chlamydia, F. necrophorum) or fungal.
   GAS pharyngitis most common in kids 5-11 years and in winter/spring.<sup>CPS</sup>
- Scoring systems e.g. modified Centor score (94% sensitivity; 54% specificity) can help with clinical assessment.
   Exception: modified Centor score may not accurately predict risk during epidemics or in groups at high-risk for acute rheumatic fever / complications e.g. remote Indigenous communities, history of acute rheumatic fever, valvular heart disease, immunosuppression. Use clinical judgment & consider testing (RADT/throat swab) more broadly.
- Diagnostic tests (RADT, throat swab) and/or antibiotics <u>not</u> recommended if: 

  Modified Centor score ≤1 (Table 1).

  symptoms of viral infection e.g. rhinorrhea, cough, oral ulcers, hoarseness. DSA'12 (strong, high)
- 3 <3 years, unless risk factors present e.g. sibling with GAS, outbreak. DSA'12 (strong, moderate), CPS
- asymptomatic household contacts of patient with GAS pharyngitis. IDSA'12 (strong, moderate)
- Positive RADT confirms diagnosis (high specificity, 88-99%). Throat swabs suggested in kids (e.g. 5-15 years<sup>105A</sup>) & high-risk groups with negative RADT (throat swabs have ↑sensitivity). Negative RADT acceptable in adults (low GAS incidence).
- GAS pharyngitis is often self-limiting; however, antibiotics recommended to ↓complications, including:
- Suppurative complications e.g. peritonsillar abscess (quinsy), sinusitis, otitis media, lymphadenitis, mastoiditis.
- Non-suppurative complications e.g. acute rheumatic fever, rare in Canada (0.3 cases per 100,000 children/year)

  Templeton<sup>107</sup> but higher in resource-poor settings (e.g. lower socioeconomic status, household crowding, limited access to health).

  Antibiotic ↓ acute rheumatic fever (RR ~70%, ARR 1%), but studies outdated & not reflective of current Canadian incidence. Spinks 13

Clindamycin

DALACIN C, g

Clarithromycin

Azithromycin

ZITHROMAX, g

BIAXIN, g

Table :	Table 1. Modified Centor (or McIssac) Score			
	Crite	ria	Points	
Temp >	38°C (>100.5	$^\circ$ F) oral temp	1	
Absenc	e of cough		1	
Swoller	n, tender ante	rior cervical nodes	1	
Tonsillar swelling or exudate		1		
Age 3 years to 14 years		1		
Age 15 to 44 years		0		
Age ≥45 years -1			-1	
Score Risk of GAS Suggested Manage		ement		
-1 to 0	-1 to 0 1 - 2.5% Symptomatic tx. No RADT,		RADT,	
1	E 100/	swah ar antibiatic n	aadad *	

-1 to 0	1 - 2.5%	Symptomatic tx. No RADT,
1	5 - 10%	swab, or antibiotic needed
2	11 - 17%	RADT or throat swab. If
3	28 - 35%	GAS positive
4+	51 - 53%	Using a cutoff of 3 is recommended by some to $\sqrt{false}$ positives. Expert
	•	

Macrolide considerations:

- Clarithromycin x10 days superior to

azithromycin x5 days for bacterial

eradication (NNT=9) in adults, but

equivalent for clinical cure. Kaplan'01

some uncertainty whether 3-5d of

Azithromycin: no head-to-head trials of 3d

vs 5d; both provide same total dose over

course of tx (i.e. 60mg/kg; 1.5g). There is

azithromycin is sufficient to  $\sqrt{ARF}$  risk.

See exceptions in Overview section

# An Approach to Treatment

- Most cases do <u>NOT</u> require antibiotics due to viral etiology. Advise on symptomatic management (see Table 2).
- Strategies: watchful waiting (e.g. 4-5d), delayed ABX (await throat swab results), empiric ABX (stop if throat swab negative).
- Use validated clinical decision tool (see Table 1) to determine risk of GAS infection. If ≥2, RADT or throat swab.
   Patients with a positive throat swab should receive an antibiotic (Table 3) to decrease risk of complications.
- The turn-around-time for throat swab results can take a few days. Antibiotics <u>started within 9 days of symptom</u>
   onset and given for 10 days in confirmed GAS will help prevent rheumatic fever (see <a href="Light-18">Light-18</a>). Casey '05, BMI' 19

01100	tana given for 10 days	prevent medinate level (see =).
Table	2. Symptom Management	see RxFiles OTC Products Chart page 215 for more details
ANALGESICS	NSAIDs e.g. Ibuprofen ADVIL, g ▼ OTC Peds: 5-10 mg/kg po q6-8hr PRN (susp X ▼) (max 40mg/kg/day) Adults: 400mg q6-8hr PRN (\$7/50 tabs g) (max 2.4-3.2g/day)  Acetaminophen TYLENOL, g X ▼ OTC Peds: 10-15mg/kg po q4-6hr PRN (max 75 mg/kg/day) Adults: 1g q4-6hr PRN (max 4g/d) (\$8/120 tabs g)	<ul> <li>Reduce fever.</li> <li>Ibuprofen decreased associated pain more than acetaminophen and placebo. Gehanno'03</li> <li>Alternative: Naproxen, g prescription, susp &amp; tab; on SPDP, ▼ ALEVE, g X ▼ OTC:≥12 years (\$12/100 tab g)</li> <li>-Peds, &gt;2yrs: 5-7mg/kg/dose q8-12hr (max 1g/day)</li> <li>-Adults: 220-500mg BID (max=1-1.5g/day)</li> </ul>
LOCAL	Benzocaine  CEPACOL, CHLORASEPTIC X ⊗ OTC  10mg lozenge q2hr PRN (\$6/18 lozenges)	- Alleviates throat pain if used frequently. Chrubasik' 12 - Avoid in peds: choking & methemoglobinemia risks Alternative: hard candy e.g. HALLS, honey (>1 year)
1 ∢	Phenol CHLORASEPTIC X ⊗ OTC  5 sprays q2hr PRN (\$15/177 mL)	No evidence, but anecdotally may provide relief from associated pain.
RINSES	Warm liquids e.g. tea, warm saltwater gargle (recipe: ¼ to ½ tsp salt per 240mL warm water)  Benzydamine, PHARIXIA, g ✗ ⊗  15mL gargle/rinse q1.5-3hr PRN (\$38/250mL)	Little evidence, but anecdotally provide relief from associated pain/discomfort.

Systemic corticosteroids (dexamethasone 10mg adults or 0.6mg/kg pediatrics oral x 1 dose) not recommended for symptom management; NICE'18, IDSA'12 (weak, moderate) however, opinions vary. Shared decision making may be used to consider corticosteroids in select cases. BMJ'17 (weak), AFP'23

- Oral/IM corticosteroids x1-2 doses have been shown to decrease pain vs placebo (NNT≈5) but have no effect
  on clinical course or days missed from school/work. No difference in adverse events (but poor reporting).
- <u>Caution</u>: there is concern that corticosteroids may mask possible underlying complications in children. Cochrane'20 (9 RCTs, n=1319), Chiappini'17

Management of Chronic GAS Carriage and Recurrent GAS Pharyngitis: antibiotics <u>not</u> routinely recommended for chronic GAS carriers (unlikely to transmit infection, low risk for complications). CPS For high-risk patients, eradication therapy (e.g. amoxicillin-clavulanate, clindamycin) may be considered.

Mostly viral. Antibiotics only in confirmed bacterial p	pharyngitis. Choosing Wisely See Table 2.
Peds: ≤27 kg: 300mg po BID or TID x10 days  No commercially available suspension  >27kg or Adults: 300mg TID or 600mg BID  x10 days	- 1st line due to narrow spectrum, efficacy safety & low cost No documented resistance to GAS Admin: preferred when given on an empty stomach ↑absorption.
Peds: 40-50mg/kg/day ÷ BID  x10 days (max 1g/day), or GAS pharyngitts indication 50mg/kg/day daily x10 days (max 1g/day) <sup>CPS</sup> Adults: 500mg BID x10 days	- Compared to penicillin: 1 broader spectrum ↑selective pressure, 2 as effective, 1 liquid more palatable for peds 4 if mononucleosis, may cause skin rash.
Peds: 25-50mg/kg/day ÷ BID <b>or</b> QID x10 days (max 1g/day) Adults: 250mg QID x10 days, <b>or</b> 500mg BID x10 days	- 1 <sup>st</sup> generation (cephalexin, cefadroxil) preferred over 2 <sup>nd</sup> gen (cefuroxime, cefprozil) due to narrower spectrum Alternatives: <b>Cefuroxime CEFTIN</b> , g
Peds: 30mg/kg daily or ÷ BID x10 days (max 1g/day) No commercially available suspension Adults: 500mg BID x10 days, or 1000mg daily x10 days	Peds: 20mg/kg/day cc ÷ BID x10 days (max 500mg/day) Adults: 250mg BID cc x10 days Cefprozil CEFZIL, g Peds: 15mg/kg/day ÷ BID x10 days (max 500mg/day) Adults: 250mg BID x10 days
	Peds: ≤27 kg: 300mg po BID or TID x10 days  No commercially available suspension  >27kg or Adults: 300mg TID or 600mg BID  x10 days  Peds: 40-50mg/kg/day ÷ BID  x10 days (max 1g/day), or GAS pharyngitis indication 50mg/kg/day daily x10 days (max 1g/day)CPS  Adults: 500mg BID x10 days  RGY: NON-SEVERE (e.g. delayed >72h rash) see Betillin or amoxicillin direct challenge/de-labeling and patillin direct challenge/de-labeling and patillin direct challenge/de-labeling and patillin direct challenge/de-labeling and patillin direct challenge/de-labeling and patilli

Only use in confirmed GAS & serious reaction to penicillin, due to \*\textstyre\textstyre{\textstyre{18}}\textstyre{18}\textstyre

Peds: 20-30mg/kg/day ÷ TID

Adults: 300mg TID x10 days

Adults: 250mg BID x10 days

500mg daily x3 days

x10 days (max 900mg/day)

Peds: 12mg/kg/day daily x5 days, or

Adults: 500mg Day 1, 250mg x Days 2-5, or

Peds:  $15mg/kg/day \div BID x10d (max 500mg/day)$ 

20mg/kg/day dailyx3 days (max 500mg/day)

Table 3. Empiric Drug Regimens see RxFiles Oral & IV Antibiotics Chart page 81 for available products, price, etc

# Acute Bronchitis: Management Considerations 1 CHEST '20, 2 NICE'19, 3 Bugs&Drugs'23, 4 Sanford'24, 5 MUMS'24

## Clinical Pearls

- Antibiotics are NOT recommended, as acute bronchitis is predominantly viral.
  - o Advise on therapies that will provide symptomatic relief: maintaining hydration, increase humidity; honey may help in those ≥1 year. Antitussive or expectorant may be considered ≥6-12 years (mixed data, ?↓ coughing episodes). Role of inhaled bronchodilators e.g. beta-2 agonist is mainly in those with underlying airway disease e.g. asthma. Oral or inhaled corticosteroids are not recommended unless concomitant asthma.
- Patients should see their prescriber if: 1 symptoms worsen, 2 new symptoms develop (e.g. dyspnea, fever, vomiting), 3 cough >1 month, or 4 >3 episodes per year.

# Overview<sup>1-12</sup>

- Bronchitis is a lower respiratory tract infection of the bronchi consisting of **persistent cough** ± sputum production, mild wheezing or dyspnea. Symptoms typical of a common cold (e.g. headache, nasal congestion, sore throat) may be present if preceding/coinciding upper respiratory tract infection.
- Acute: self-limiting, cough usually lasts for 1 to 3 weeks (median 18 days), 13 up to 25% will have a cough beyond 3 weeks as airway hyperactivity may last up to 6 weeks. 14,15
- Incidence: ~5% of adults per year will develop acute bronchitis, ~12% of those age <5yrs per year.</li>
- Etiology: ≥90% of cases are viral e.g. Influenza A or B, Parainfluenza, Respiratory Syncytial Virus (RSV), Adenovirus, Coronavirus; minority bacterial e.g. B. pertussis, M. pneumoniae, C. pneumoniae.
- Chronic: cough usually lasts ≥3 months (note usually current/former smokers).
- Diagnosis of acute bronchitis is clinical (history and physical exam). Differential diagnosis tips:
- Rule out **pneumonia** with chest x-ray if the following: heart rate >100bpm, respiratory rate >24-30, oral temp >38°C (mental status/behavioural changes in those who may not mount a fever e.g. older adults), or findings of local consolidation on physical exam.
- Also consider pertussis, COVID-19, influenza, or other respiratory viruses. Treat with an antimicrobial if appropriate see RxFiles Charts: Community Acquired Pneumonia page 116, Common Infections page 96, COVID-19 page 122. Influenza page 90. See Q&A (right) for bronchiolitis.
- Coloured sputum does not reliably differentiate between bacterial / viral origin. Choosing Wisely Observational study of n=241 pts with acute cough found positive likelihood ratio 1.46 (indicating minimal increase in the likelihood of bacterial infection given coloured sputum). 17 Theory: WBC release of peroxidases → coloured sputum.
- Those with complicating factors (e.g. smoking, impaired lung function COPD, asthma, chronic heart disease, immunocompromised) may require further investigations (e.g. lung function tests, chest x-ray).

# An Approach to Treatment<sup>1-12</sup>

- Reassurance & symptom management (see Table 1) are the cornerstones of care. Tools:
- RxFiles Viral Prescription: adult, pediatric
- RxFiles Clinic Bronchitis Poster "Gone Viral"
- CDC Patient Handout "Preventing & Treating Bronchitis"
- Choosing Wisely in Primary Care: "don't prescribe antibiotics for bronchitis"
- Antibiotics are not recommended for acute bronchitis (no/minimal benefit, 1-5, Choosing Wisely
- A 2017 Cochrane review (17 RCTs, n=5099) of antibiotic vs placebo found at 14 days: 18
- No difference in clinical improvement ~73% vs ~66% (RR 1.07, 95% CI 0.99 to 1.15)
- ↓mean cough duration by ~0.5 days (95% CI -0.87 to -0.04 days)
- ↑AE e.g. N/V/D, headache, rash, vaginitis ~23% vs ~19% NNH≈25 (RR 1.2, 95% CI 1.05 to 1.36)
- A prospective cohort study (n=28,779) found low risk of hospital admission/death in those ≥16 years with lower respiratory tract infection after initial assessment whether prescribed immediate antibiotic (0.9%), delayed antibiotic (0.4%), or no antibiotic (0.3%). 19 Little '17
- A retrospective cohort study (n=749,389) found the risk of developing pneumonia after acute bronchitis was low in those 0-64 years: 0.25% (antibiotic treatment) vs ~1.2% (no antibiotic). However, the risk for those ≥65yrs was higher: 1.5% (antibiotic treatment) vs 4% (no antibiotic) and these individuals may warrant closer follow up post-acute bronchitis episode. Petersen'07

#### Table 1. Symptom Management<sup>1-12</sup> Other options: throat lozenges HALLS, hot tea, etc Non-pharmacologic: Limited or lack of evidence to support use. Anecdotally, can help for some. Hydration: caution in heart failure or chronic kidney disease. Hydration<sup>20,21</sup> e.g. PRN to maintain 30-50% humidity; clean frequently \$\psi\$ bacteria/fungi Humidity A 2021 Systematic review of people with URTIs (majority peds) Honey 2.5 to 10mL po HS PRN Available in Oproducts but likely found honey superior vs usual care for cough frequency & severity (4-6 RCTs, n=~450-600) but ?clinical superiority. No subtherapeutic dose

Antitussive: Limited/mixed evidence. Some suggest may use, 3-5 others do not. 1,2 See OTC chart pg 215.

# Dextromethorphan (DM) BENYLIN DM \$11/250mL X ⊗ OTC

<1vrs: not recommended (botulism concerns)

ROBITUSSIN DM \$11/250mL; \$11/20 gel caps X 🛇 OTC

Be aware of many combo products ©

?may ↓ coughing episodes, but duration of illness limited data.<sup>25</sup> Cochrane'14?reserve HS if sleep issues. ?↑AE e.g. GI, dizziness.2 NICE CI: <6yrs, safety & efficacy concerns. 26 Health CDN (check, some only ≥12yrs).

difference vs dextromethorphan (2 RCTs, n=137).<sup>22 BMJ'21,23,24</sup>

DI: serotonergic, avoid MAOIs. Other antidepressants: monitor for sx given short DM duration RxFiles Antidepressant DIs Chart.

Expectorant: Limited/mixed evidence. Some suggest may use,<sup>2</sup> others do not.<sup>1</sup> See OTC chart page 215.

# Guaifenesin X ⊗ OTC \$6/100mL "Mucous & Phlegm" Be aware of many combo products ©

?may \$\square\$ number of coughing episodes, other studies found no benefit. 25 Cochrane'14 ?more beneficial if excess mucus. 27 Albrecht, Expert CI: <6yrs, safety & efficacy concerns. 26 Health CDN (check, some only ≥12yrs).

**Bronchodilators**: Not for routine use, 1 questionable benefit, potential AE, carbon footprint. Some suggest not offering unless underlying airway disease e.g. asthma.<sup>2 NICE</sup> Others indicate may consider e.g if long-lasting cough, especially if dyspnea or wheezing.<sup>3-5</sup> See RxFiles Asthma Chart page 206.

Salbutamol VENTOLIN, g
100mcg 2 puffs TID-QID\$17/device
Salbutamol VENTOLIN, g 100mcg 2 puffs TID-QID <sup>\$17/device</sup> Terbutaline BRICANYL <b>Ø</b>

500mcg 1 puff TID-QID\$22/device

Clinical trials dosing was scheduled, but may consider PRN

**Ipratropium ATROVENT** 20mcg 4 puffs QID\$32/device **Ipratropium + Salbutamol** COMBIVENT 20/100 mcg 1 puff QID\$43/device A 2015 Cochrane review of beta-2 agonist (most po) vs placebo: 28-30

- No benefit for cough among adults (3 RCTs) or children (2 RCTs).
- AE (1 RCT, n=73 adults): ↑ tremor ± palpitations over 7 days (49% inhaled fenoterol vs 0% placebo). 31 Melbye'91
- Subgroup with wheezing/airflow obstruction (1 RCT, n=35) adults): potential benefit;  $\downarrow$  average total symptom score. days 2-7 (54% fenoterol vs 32% placebo, NNT=5).31 Melbye'91

Limited data. URTI post-infectious cough, ?daytime & nighttime improvement (1 RCT, n=14 x 3 weeks using inhaler; 32 Holmes '92 1 RCT, n=92 x 10 days using nebules with concomitant salbutamol 33 Zanasi'14).

# Not recommended for symptom management:1

- Oral or inhaled corticosteroids are not recommended in patients with acute bronchitis without asthma.<sup>2,5</sup> Post-infectious cough: 2 RCTs, n=163 with cough 3-8 weeks, placebo improved cough scores at 2 weeks by ~50% and inhaled corticosteroids improves score by ~2-13% more.<sup>34 TFP'22</sup>
- Codeine syrup 5mg/mL \$40/500mL; 3.3mg/5mL \$10/100mL is not recommended in patients with acute bronchitis.<sup>2</sup> NICE'19 While reductions in cough have been found, there is no difference between codeine syrup (120mg/d) vs non-medicated syrup. 35 Eccles 92,36,37 Anecdotally can help by ↑drowsiness HS.
- N-acetylcysteine is not recommended in patients with acute bronchitis. 2 NICE'19 6 RCTs, n=497: found a reduction in cough at seven days but questionable clinical benefit and no benefit at 28 days compared to placebo (very low quality evidence). 38 Cochrane 2013

Encourage prevention strategies e.g. smoking cessation & avoid secondhand smoke see RXFiles Smoking Cessation page 212, avoid exposure to chemicals/dust/air pollution, ensure vaccines are up to date, see RxFiles Vaccines page 91 e.g. influenza, RSV, COVID-19, pertussis, pneumococcal, haemophilus.

**Q&A:** Is bronchiolitis the same as bronchitis? Bronchiolitis is a lower respiratory tract infection of the bronchioles (more distant, smaller airways) caused by viruses in children <2 years (and usually <1 year). Supportive care is the mainstay of therapy. Antibiotics are not recommended.<sup>39</sup> Choosing Wisely Bronchiolitis: Less is Best. 40 Lirette CFP'23, 41 Greenky CFP'22

- 1) **Demographics**: ~10% of the population will report a penicillin allergy, but <1% are truly allergic. Overreporting leads to poor patient outcomes. Thus, there is potential to reassess potential allergies in low-risk patients and de-label when appropriate.
- Avoid classifying common side effects as "allergy" to proactively prevent incorrect pen allergy labeling.
- β-lactam hypersensitivity (HSR) wanes with time: 1.5
   ~60% loss of HSR at 5 years for cephalosporins, &
   ~80% loss of HSR at 10 years for penicillin<sup>per skin tests</sup>.
- Rashes coinciding with viral infections in childhood, are often non-allergic in nature e.g. maculopapular.
- Risk Calculator/De-labeling Tools (see below) are useful for assessing a previously reported β-lactam "allergy", identifying those at low-risk based on risk stratification, & who may be suitable for de-labeling.
   a) SHA Firstline Penicillin Allergy De-labeling Tools
   b) PEN-FAST-adults<sup>9</sup> c) 1-1-1-Criterion<sup>10</sup>
- 6) Patient Tools: <u>JAMA Peds: Allergy De-labeling</u><sup>11</sup>
  SHA Poster: <u>"Do you actually have a pen allergy?"</u>
- Penicillin-Cephalosporin Cross-reactivity is now estimated at 1-2%. Agents with dis-similar side chains will be even less likely to cross-react.<sup>1,12</sup>

# Introduction:

Beta-lactams, especially penicillins, are often implicated in hypersensitivity reactions (HSR). Such allergies are over-reported, resulting in poor outcomes e.g. treatment failures, \(\triangle \) use of less effective 2<sup>nd</sup> line or broad spectrum antibiotics, \(\triangle \) adverse events (\(\circ 25\% \triangle \) in C. diff infections vs those without<sup>35</sup>), & antibiotic resistance (\(\circ 18\% \triangle VRE, \(\circ 40\% \triangle MRSA^{36}\)]. (5.7,13,14,33 In pen/amox allergy, do not order non beta-lactam without an appropriate evaluation. (\(\circ \circ \circ \))

# **Definitions, Classifications, and Additional Info:**

- Beta-lactams: antibiotics with a distinctive beta-lactam ring (i.e. penicillins, cephalosporins, carbapenems, & monobactams). Although an allergy may occur to the beta-lactam ring, most true IgE mediated allergies are likely due to the unique side chain (e.g. R1), with cross-allergy only to specific agents (See Table 1).
- IgE: immunoglobulin type E antibody. After encountering a specific allergen, IgE can trigger an immediate (<1hr) or accelerated (~<6hr) immune response.
- De-labeling: removal of an allergy label from a patient's medical records (facilitated by: reaction history +/- direct challenge +/- skin test + patient education)
- Direct De-label: If allergy assessment indicates no true allergy, and no increased risk of a severe reaction, a β-lactam may be given along with de-labeling. 15
- Direct Challenge: A drug is administered to determine tolerance (often without prior skin testing).<sup>1,7,8,26,27</sup> It is **suitable for low-risk patients** and done in a setting where patient can be observed for ~30-60 minutes in case of rare anaphylaxis (i.e. in an office/clinic with access to epinephrine). If there is no IgE-type immediate hypersensitivity reaction during observation, and no delayed reaction on follow-up (e.g. after ~5 days), the allergy label may be removed. For a **1-step challenge**, typically amoxicillin is used [500mg x 1 (adult) or 15-30mg/kg x1 (child; chewable tab 125-250mg option)]. For a cautious **2-step graded challenge**, a small dose of the potential allergen (e.g. 10-25% of a full dose)<sup>7</sup> is given, followed by the balance of, or full dose, ~20-60 min later. In the inpatient setting, an inappropriate penicillin allergy label resulting in use of vancomycin or clindamycin may result in suboptimal outcomes, e.g. clinical failure, *C. difficile infection*, acute kidney injury, etc. Often a β-lactam can be used. Take advantage of setting to direct-challenge, monitor, and de-label when possible. <sup>17,18</sup>}
- Penicillin skin-testing: pricking the skin with a minute quantity of penicillin; if a localized reaction not observed, an IgE-mediated allergy is unlikely. It is suggested primarily for patients with history of a possible IgE-mediated Type I reaction (e.g., immediate anaphylaxis, or immediate onset urticaria/hives).

Table 1: Oral beta-lactam groups with identical/? similar R1 side chains & a higher risk of cross-reaction: 1,12,23 (Immediate cross-reactivity HSR related to R1 sidechain)

Amoxicillin: cefadroxil, & cefprozil; ?cephalexin

Ampicillin: cephalexin; ?cefadroxil, ?cefprozil

See cross-reactivity resource links below.

Of note: cefazolin has a unique side chain dissimilar to all other beta-lactams; it can safely be used in patients reporting a penicillin allergy/anaphylaxis. 1,12,19

Table 2: Classification of Adverse Drug Reactions that May Be Reported as "Penicillin/Beta-lactam Allergy":

Adverse I	Orug Reactions -	NON-IMMUNE	Common Features/Presentation		Considerations for Risk Stratification, Management and/or De-labeling
Non-seve	re, non-immune	Predictable	Side effects: isolated nausea, vomiting, diarrh	ea, headache	Such effects are NOT an allergy; a β-lactam can be given along with direct de-labeling. <sup>7</sup>
Adverse I	Orug Reactions -	ug Reactions - IMMUNE-MEDIATED Common Features/Presentation (Use tools, see Firstline or PEN-FAST below and/or guideline recommendations to guide course of action)			AST below and/or guideline recommendations to guide course of action)
Type I	Immediate; Usually ≤30-60 minutes (up to 6 hrs) <sup>7</sup>	<mark>IgE</mark> -mediated <i>True Allergy</i>	Urticaria/hives, see photos angioedema, anaphylaxis, bronchospasm; assess for risk of severe anaphylaxis with re-exposure; risk wanes such that ~80% will not react after 10 years lg \$\psi\$ over time risk wanes such that ~80% will not react after 10 years lg \$\psi\$ over time risk wanes such that ~80% will not react after 10 years lg \$\psi\$ over time risk wanes such that ~80% will not react after 10 years lg \$\psi\$ over time risk wanes such that ~80% will not react after 10 years lg \$\psi\$ over time risk wanes such that ~80% will not react after 10 years lg \$\psi\$ over time risk wanes such that ~80% will not react after 10 years lg \$\psi\$ over time risk wanes such that ~80% will not react after 10 years lg \$\psi\$ over time risk wanes such that ~80% will not react after 10 years lg \$\psi\$ over time risk wanes such that ~80% will not react after 10 years lg \$\psi\$ over time risk wanes such that ~80% will not react after 10 years lg \$\psi\$ over time risk wanes such that ~80% will not react after 10 years lg \$\psi\$ over time risk wanes such that ~80% will not react after 10 years lg \$\psi\$ over time risk wanes such that ~80% will not react after 10 years lg \$\psi\$ over time risk wanes such that ~80% will not react after 10 years lg \$\psi\$ over time risk wanes such that ~80% will not react after 10 years lg \$\psi\$ over time risk wanes such that ~80% will not react after 10 years lg \$\psi\$ over time risk wanes want was also		
Type II	Delayed; usually after	IgG-mediated or other (non-IgE);	Cytotoxic: blood cell dyscrasias (e.g. hemolytic anemia, thrombocytopenia); onset hours-days	Rare! If suspicion, high risk; not for	Risk stratification guides management: <u>Firstline</u> tool/app helps guide options.  If history = high-risk/severe (severe cutaneous SCARs, or organ involvement, often requiring ER
Type III	<mark>several</mark> days - weeks	onset >7days	Fever, vasculitis/rash, arthralgia; SSLR*	rechallenge!	visit/ hospitalization, Type II & III) do <u>not</u> administer a β-lactam. See below re "SSLR*"  If history = low-risk reactions (delayed onset, <u>no</u> systemic symptoms; mild, <u>not</u> involving
Type IV	<mark>(&gt; 72hrs)</mark>	T-cell mediated, or other (non-lgE)			blistering of skin or mucous membranes), may consider a direct challenge.
Other: al	Family Hy of alle	ray Inc. association of fam	ily by with individual allormy) or h) Same antihiotic tak	en again tolerated	Not a true allergy: a B-lactam can be given along with direct de-labeling: 7 address fears

# Table 3: Tools to Assess Risk of True Penicillin (Beta-lactam) Allergy and Evaluate for Possible De-labeling:

144.6 0. 100.6 10. 100.6 11. 11. 1. 1. 1. 1. 1. 1. 1. 1. 1. 1. 1				
SHA Firstline: Penicillin Allergy De-labeling App	PEN-FAST: an Allergy Decision Rule (for adults only) 9,22,28 PALACE	<b>1-1-1 Criterion in Urticaria</b> for Risk Stratification; β-Lactam Allergy <sup>10</sup>		
App/online guideline to aid in assessing & managing patients	PEN = Penicillin allergy reported (if yes, continue)	1) Did the urticaria develop after the first dose?		
with suspected or proven beta-lactam allergy. Includes:	F = ≤ Five years since reaction or interval unknown (2 points)	2) Did the urticaria develop < 1 hour of inciting dose?		
assessment tool, risk categories, algorithms, oral challenge	A,S = Anaphylaxis, Angioedema, or Severe cutaneous reaction (2 points)	3) Did the urticaria <u>resolve within 1 day</u> of stopping?		
protocol, what to do with the po challenge result, photo gallery	T = Treatment was required for the allergy (1 point)	If three "yes" answers: patient has a 90% positive predictive		
of dermatological reactions, etc. (adult - SHA, children - BC Children's, CHEO)	Score: 0=very low risk (<1%), 1-2=low risk (5%), 3=moderate risk (20%), 4-5=high (50%)	value for probable β-lactam allergy (ie. high-risk). <sup>10</sup> (adult & child)		
*SSI P-serum sickness like reactions: are more common than the other serious reactions listed; they generally occur after 7-10 days of therapy and relate to				

How likely is a beta-lactam allergy?		
10,000 In a given group of 10,000 patients		
<1,000 (<10%)	⇒ will <b>report</b> they have a penicillin allergy,⁴	
<100 (<1%)	⇒ will <b>have</b> a true IgE-mediated penicillin allergy, <sup>6</sup>	
1-3 (<0.03%)	⇒ will have some cephalosporin cross-reactivity, <sup>20</sup>	
1 (<0.01%)	⇔ will have anaphylaxis when given penicillin. <sup>5,21</sup>	

\*SSLR=serum sickness like reactions: are more common than the other serious reactions listed; they generally, occur after 7-10 days of therapy and relate to immune complexes of IgG. Symptoms include urticarial vasculitis, and joint pain. Skin testing not helpful. Challenges & desensitization contraindicated!!!

Cephalosporin allergy: similar to penicillins; less studied. Cross-reactivity Resources: 1) Bugs & Drugs 23 2) Firstline 3, 3) Zagursky et al. 22, 4) UNMC, 5) CSACI 2020 1

- Cross-reactivity data: Cephalosporin and penicillin cross-reactivity rate was thought to be up to 10%; however, the true rate is likely 1-2%. 37 In general, a cephalosporin can be given if history of non-severe (non-anaphylactic) pen-allergy. If non-anaphylactic ceph-allergy, consider direct challenge with cephalosporin with dissimilar side chain. Evidence suggests that carbapenems have 16 cross-reactivity with penicillins, are appropriate in penallergies any time a cephalosporin could be prescribed. [Cefazolin IV: unique side-chain; no cross-allergy; see above. Aztreonam IV: see online extras.]

Skin tests in SK are available via referral (but currently 2+ year wait, if routine 2024). Option for questionable severe IgE allergy history.

In patients with a history of low-risk reactions (delayed onset, not accompanied by any systemic symptoms, and not involving blistering of skin or mucous membranes), one may consider treatment with a beta lactam using a test dose direct challenge) approach.

Abbreviations: AAAAI/ACAAI=American Academy of Allergy, Asthma and Immunology; American College of Allergy, Asthma and Immunology DRESS=drug reaction with eosinophilia and systemic symptoms HSR=hypersensitivity reaction Ig=immunoglobulin

SCARs=severe cutaneous adverse reaction SHA=Saskatchewan Health Authority SJS=Stevens-Johnson syndrome SSLR=serum sickness like reactions TEN=toxic epidermal necrolysis VRE.

		osing recommendation also available via S	<u>SK Antimicrobial Stewardship</u> & Firstline (app or <u>JPCF</u>	
Generic/TRADE	Formulations	Comments	<b>Dosing</b> suspension \$: assumes 10kg & 20kg child	\$/ <mark>5day</mark>
	o penicillin binding proteins on bacterial cell wa			1
Penicillin V	no commercially available	-Drug of choice for <b>GAS pharyngitis</b> (if antibiotic required*). 1 CPS'21 \$\square\$ selective	≤27kg: 300mg po BID ac <u>or</u> 300 mg po TID ac >27kg: 300mg po TID ac or 600mg po <b>BID</b> ac	\$13-15 <sup>tab</sup> \$15-16
Potassium 者	suspension (~\$80 if compounded) tab: 300mg <sup>c</sup> (480,000 unit)	pressure on other bacteria vs amox.	*for GAS pharyngitis, dose for <b>10 days</b> to prevent	\$15-10
PEN-VK, g	{tab. 300 mg (460,000 unit) {tab scored & may be crushed}	-Preferred, 1hr before food ↑ absorption. <sup>2</sup>	rheumatic fever (\$16-21).1 CPS'21	
Amoxicillin PL	suspension: 25mg/mL, 50mg/mL & 🙂	-Drug of choice for <b>AOM</b> or <b>acute</b>	Peds standard-dose:	
AMOXIL, g	(strawberry, banana, sugar-free, berry)	sinusitis (if antibiotic required).3 CPS'16, 4	40 to 50mg/kg/day po ÷ <b>TID</b> AOM, sinusitis	\$14-17
Moxilen Forte	chew tablet: 125mg, 250mg (cherry)	-Narrower spectrum (less selective	40 to 50mg/kg/day po ÷ <b>BID</b> -TID <sup>GAS</sup> pharyngitis 50mg/kg/day po once daily <sup>GAS</sup> pharyngitis, 1 CPS'21	\$14-17
amoxicillin = amox	capsule: 250mg, 500mg	pressure on other bacteria) & fewer AE e.g. diarrhea, yeast infection than	Peds high-dose: select AOM patients (see left)	\$14-17
		amoxicillin/clavulanate. 5 Savage'23	80 to 90mg/kg/day ÷ <b>BID</b> -TID (max 4g/day)	\$17-22
	Why standard BID dosing for Group A Streptococcus (GAS) pharyngitis?	- <u>Standard dose</u> : for most individuals.	Adult (≥16 years <sup>expert</sup> ) <u>standard-dose</u> :	
	GAS is an easier bacteria to kill than	- <u>High dose</u> : for those with risk factors  S. pneumoniae resistance i.e. daycare,	500mg po TID AOM, sinusitis	\$13
	S. pneumoniae, which is the main	<2 years, under / unimmunized, or	500mg po BID GAS pharyngitis	\$12
	AOM/sinusitis pathogen so BID is adequate.	antibiotics within previous 3 months.	Adult <sup>≥16yrs</sup> <u>high-dose</u> : 1000mg TID <sup>AOM, sinusitis</sup>	\$15
Amoxicillin 🚹	susp: 25mg / 6.25mg/mL (4:1) *	-Reserve for treatment failure to 1 <sup>st</sup>	Peds standard-dose:	4
/ Clavulanate	50mg / 12.5mg/mL (4:1) * 40mg / 5.3mg/mL (7:1) *	line agent in <b>AOM</b> or <b>acute sinusitis</b> (if antibiotic required). 3 CPS'16, 4 CSO-HNS'11	Amox/clav 7:1 ratio 45mg/kg/day ÷ TID cc	\$25-40
CLAVULIN, g 者	80mg / 11.4mg/mL (7:1) *	-Broader spectrum & ↑AE e.g.	Peds <u>high-dose</u> : requires 2 prescriptions 14:1 ratio {	\$39-57
amoxicillin = amox	(raspberry-orange)	diarrhea (RR 1.15), yeast infection	2 Amoxicillin 45mg/kg/day} ÷ BID-TID cc	
clav = clavulanate	tab: 250mg / 125mg (2:1)	(RR 1.33) compared to amox. 5 Savage'23		
	500mg / 125mg (4:1)	-≤3 months of age: use 4:1 ratio (limited data with 7:1 ratio). 6, Expert	Adult (≥16 years <sup>expert</sup> )	\$17
	875mg / 125mg (7:1)	-Preferred max clav ≤10mg/kg/day <sup>peds</sup>	standard-dose: 500/125m <sup>4:1 ratio</sup> g po TID cc <u>or</u> 875/125mg po BID cc	\$17
	Canada: <b>no</b> commercially available 14:1	& 125mg/dose <sup>adults</sup> to $\downarrow$ <b>AE</b> diarrhea. <sup>6-9</sup>	high-dose: { 1 Amox/clav 875mg 7:1 ratio BID cc	\$30
	USA :: commercially available 14:1AUGMENTIN	-Preferred to take with food (cc)	PLUS 2 Amoxicillin 1000mg BID} sinusitis	
	Sample Prescription <sup>6</sup> (recurrent AOM, 22 mont	↑tolerability & ↑clav absorption. 6-9 hs therefore 10 day duration, high-dose due	L e to patient risk factors for <i>S. pneumoniae</i> resistance e.g	davcare)
		0 days (90mg/kg/day; patient weight 14kg		
		spension 210mg po BID-TID AND		
		210mg po <b>BID</b> -TID I <b>LIN</b> and <b>plain amoxicillin</b> together for eac	ch doca two to three times daily	
Canhalosporins			s. Bactericidal. Demonstrates time-dependent killing	
			amoxicillin allergy de-labelling is not appropriate/fe	
	<mark>am Allergy Chart</mark> page 6, Saskatchewan Health Au	thority (SHA) <u>Firstline</u> for further informati	ion and de-labelling algorithms/tools.	
Cephalexin M	suspension: 25mg/mL, 50mg/mL 🕸 😊	<ul> <li>-1st-generation cephalosporin.</li> <li>-Alternative for GAS pharyngitis. Both</li> </ul>	Peds: 25 to 50mg/kg/day po ÷ BID <u>or</u> QID (max 1g/day)	\$17-39
KEFLEX, g 🥕	(orange-banana) tablet: 250mg, 500mg	BID & QID dosing approved. 10 HealthCDN	Adult: 250mg po QID or 500mg po BID	\$12
Cefadroxil PL	no commercially available suspension	-1st-generation cephalosporin.	Peds: 30mg/kg/day po once daily or ÷ BID	\$15 (500mg
Cefadroxil DURICEF, g	capsule: 500mg	-Alternative for <b>GAS pharyngitis</b> . Both	(max 1g/day)	cap once daily)
DURICEF, g	capsa.c. seeg	daily & BID dosing approved. 11 HealthCDN	Adult: 500mg BID or 1000mg po once daily	\$20
Cofemanian a	25	-2nd-generation cephalosporin.	Peds: 20 to 30mg/kg/day po ÷ BID cc (max 500mg/d)	\$20-30
Cefuroxime II	suspension: 25mg/mL 🕸 🕲 (tutti-fruiti)	-Preferred to take with food 个	Adults: 250mg po BID cc or	\$20-30
axetil 🤰	tablet: 250, 500mg	absorption (~52% vs ~37% fasting).12	500mg po BID cc	\$28
CEFTIN, g		-Alternative for <b>AOM</b> or <b>acute sinusitis</b> .		4
Cefprozil	suspension: 25mg/mL, 50mg/mL 🕸 😊	<ul><li>-2nd-generation cephalosporin.</li><li>-Alternative for <b>AOM</b> or acute sinusitis.</li></ul>	Peds: 15 to 30mg/kg/day po ÷ BID (max 500mg/d) Adults: 250mg po BID or	\$14-21 \$29
CEFZIL, g	(bubblegum)	-suspension ~\$5-10 less vs cefuroxime.	500mg po BID	\$32
	tablet: 250, 500mg		Peds: 50mg/kg IM/IV daily AOM (x3 days), sinusitis (x10 days)	
Ceftriaxone III	vials for injection: 1g, 2g	<ul><li>-3rd-generation cephalosporin.</li><li>-Alt for tx failure AOM / acute sinusitis.</li></ul>	If IM, diluted/reconstituted with lidocaine 1% to $\downarrow$ pain. Expert	\$75
ROCEPHIN, g x ▼	o fourth and with a servery posicilling allower. Do	·	AS a barrage 13.14 Data last and Allegrape C. C.	IA Finatina
	ve for those with a <b>severe penicillin allergy</b> - <b>Do</b> no commercially available suspension	-Greater absorption on empty stomach	AS pharyngitis <sup>resistance 13,14</sup> . Beta-lactam Allergy pg 6, Sl Peds:** 2 to 5mg/kg/day po divided BID	HA <u>FIRSTIINE</u> .
Doxycycline DOXYCIN, g	(~\$60 if compounded, full benefit NIHB & SPDP <15yrs - ≤\$25)	$(\uparrow 20\%)$ , but may take with food to $\uparrow$	Adult: 200mg stat, then 100mg BID or	\$16
DOTTCHN, g	cap: 100mg (may open & mix with applesauce)15	tolerability.16 **<8yrs: lack of tooth	200mg stat, then 100mg once daily	\$14
	tab: 100mg (may crush, see Geri-RxFiles for tips)15	staining, may use ≤21d. 17 Ravindra'23, 18 AAP'21		
Clindamycin 🕕	solution: 15mg/mL (cherry) DO NOT REFRIGERATE (solution will thicken, but	-Take cap with a full glass of water to	Peds: 20 to 30mg/kg/day po ÷ TID (max 900mg/day)	\$24-37
DALACIN C, g	capsule: 150mg, 300mg stable 2wk room temp) <sup>19</sup>	↓ esophageal irritation risk. <sup>20</sup>	Adult: 300mg po TID	\$18
			ith time-dependence (AUC <sub>24</sub> :MIC, maximize drug co	
			p A Streptococcus (GAS) ≥18% in Saskatchewan (SK) itis. See RxFiles Beta-lactam Allergy Chart page 6, SHA F	
Azithromycin	suspension: 20mg/mL, 40mg/mL (cherry)	-Azithromycin: half-life ~72hr → 5 day	Peds: 10mg/kg day 1; 5mg/kg days 2-5 AOM, sinusitis	\$18-24
ZITHROMAX, g	After reconstitution, suspension may be stored at room	treatment ≈ 10 days therapeutic	12mg/kg/day x5d or 20mg/kg/day x3d GAS pharygitis	
	temperature but palatability may be improved if in fridge.	levels. <sup>20</sup> Appears more likely to lead to	<b>Adult</b> : 500mg po Day 1; 250mg po Days 2 to 5 <u>or</u>	\$16
Claulthan	tablet: 250mg	resistance than clarithromycin, as long half-life results in prolonged sub-	500mg po once daily x 3 days  Peds: 15mg/kg/day po ÷ BID (max 500mg/day)	\$16
Clarithromycin BIAXIN, g	suspension: 25mg/mL, 50mg/mL (fruity) DO NOT REFRIGERATE	inhibitory levels at end of therapy. <sup>21</sup>	Adult: 250mg po BID or 500mg po BID	\$18-26 \$15-20
BIAXIN. g 🎏 📙				, ,
	tablet: 250mg, 500mg, 500mg XL	-Clarithromycin more DI, but often not an issue in pediatrics. <sup>22</sup>	500mg to 1000mg XL po once daily cc	\$18-24

# **Tools to Support Antimicrobial Stewardship** all resources available @ www.RxFiles.ca/ABX



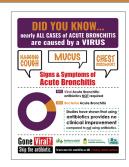
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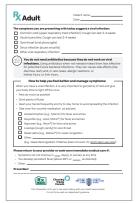






# ACUTE OTITIS MEDIA

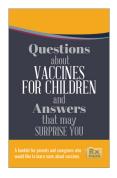
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<u>Acute Pharyngitis Chart References</u>: see RxFiles Chart <a href="https://www.rxfiles.ca/RxFiles/uploads/documents/ABX-Pharyngitis.pdf">https://www.rxfiles.ca/RxFiles/uploads/documents/ABX-Pharyngitis.pdf</a>.

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**Beta-Lactam Allergies Chart References**: see RxFiles Chart

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# **Antibiotic Formulations for AOM & Other Common Respiratory Tract Infections**

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