

## Short Course Antimicrobial Therapy in Adults

# **BOTTOM LINE:** Use of short course antimicrobial therapy is effective, reduces antimicrobial overuse and cost, and reduces risk of antibiotic-associated harms to patients.

Many randomized controlled studies have shown that short course antimicrobial treatment is noninferior to longer courses of treatment (see Table 1). Short course therapy is also endorsed by various guidelines.<sup>1,4,8,12,20,25,31-33,38-40,49,56</sup>

Infection	Duration of		Comments
	Therapy	y (days)‡	4
	Short*	Long	
Cellulitis/Abscess <sup>2-4</sup>	5-6	10-12	<ul> <li>Assumes source control achieved.</li> <li>Residual inflammation after 5-6 days of antimicrobial therapy will resolve with time and does NOT require prolongation of antimicrobials.</li> </ul>
Native hand/wrist septic arthritis⁵	14	28	Assumes source control achieved.
Osteomyelitis <sup>6-8</sup>	42	84	Applies to osteomyelitis associated with diabetic foot infection and vertebral osteomyelitis.
AECOPD/Sinusitis <sup>9-20</sup>	≤ 5	≥7	Applies to mild-moderate AECOPD and acute bacterial sinusitis.
<b>CAP</b> <sup>21-26</sup>	3-5	≥ 7	<ul> <li>Excludes hospitalized patients with severe CAP.</li> <li>Treat until clinically stable: temp ≤ 37.8°C, HR &lt; 100 bpm, RR &lt; 24 breaths per minute, arterial O<sub>2</sub> sat ≥ 90% on room air (or the patient's baseline level of home O<sub>2</sub>), SBP ≥ 90mm Hg, and normal mental status; and for a minimum of 3 days.</li> </ul>
HAP/VAP <sup>27-33</sup>	7-8	14-21	• In patients infected with non-fermenting Gram negative bacilli (e.g. <i>P. aeruginosa</i> ), short course therapy may be associated with increased recurrence BUT NOT decreased clinical cure, or increased hospital or ICU days, mechanical ventilation, or mortality. <sup>34,35</sup>
Intra-abdominal infection <sup>36-40</sup>	4-8	10-15	Assumes source control achieved.
Complicated UTI¤/ Uncomplicated pyelonephritis <sup>41-49</sup>	5 or 7	10 or 14	<ul> <li>Limited data for shorter duration for non- fluoroquinolone antibiotics</li> <li>NB: For empiric therapy, do not use fluoroquinolones due to increasing resistance, rare but serious adverse effects, and significant association with CDI.**</li> </ul>
Gram negative bacteremia <sup>50-56</sup>	7	> 10	<ul> <li>Assumes source control achieved and infection not associated with a clinical syndrome requiring longer therapy, such as osteoarticular or endovascular infections.</li> <li>For <i>Pseudomonas aeruginosa</i> bacteremia, there are three retrospective studies that found short course therapy (6-10 [avg. 8] days<sup>57</sup> and 7-11 days<sup>58,59</sup>) as effective as longer courses (11-15 [avg 13] days<sup>57</sup> and 12-21 days<sup>58,59</sup>).</li> </ul>

### Table 1<sup>†</sup>. Short course duration of therapy recommendations for common infections

Prepared by: Amor-Myville Blanco, 4th year PharmD Student; an

to, 4<sup>th</sup> year PharmD Student; <u>amormyvi@ualberta.co</u> and Susan Fryters, AS/ID pharmacist, <u>Reviewed by: Dr. Isabelle Chiu, In</u>fectious Diseases physician, Edmonton Zone

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<sup>†</sup>Adapted from <u>https://www.bradspellberg.com/shorter-is-better</u> <sup>‡</sup>Duration of therapy includes both IV and oral therapy, and starts after source control, where required, is achieved.

<sup>α</sup>Complicated UTI - associated with functional and/or anatomical abnormalities of the urinary tract. See Bugs & Drugs <u>Complicated UTI/Males</u> for more details.

\*Refer to https://www.bugsanddrugs.org/ for detailed treatment recommendations.

\*\*Refer to Antimicrobial Stewardship Backgrounder April 2022: <u>Avoid Fluoroquinolones as First-line Therapy</u> for more details. AECOPD=acute exacerbation of chronic obstructive pulmonary disease, avg=average, CAP=community-acquired pneumonia, CDI=*C. difficile* infection, HAP=hospital-acquired pneumonia, VAP=ventilator-associated pneumonia, UTI=urinary tract infection

**Background**: Avoiding unnecessarily long durations of antibiotic therapy is one way to address antimicrobial overuse. Limiting antibiotic exposure prevents antibiotic resistance and reduces the likelihood of a patient developing adverse effects or *C. difficile* infection.<sup>60,61</sup> With each excess day of antibiotic use, there is a 4-5% increased odds of a patient experiencing an antibiotic-associated adverse event.<sup>60,62</sup>

#### Table 2. Duration of antimicrobial therapy – THEN & NOW

THEN	NOW
Shorter courses of antibiotics are less effective	<ul> <li>Shorter courses are non-inferior to longer duration of therapy</li> <li>54 RCTs, including ~19,000 patients, have compared the efficacy of short versus traditional, longer duration therapies for the infections listed in Table 1 and shorter treatment was shown to be non-inferior.</li> <li>Non-inferiority was measured by rates of clinical and microbiological cure, survival, and bacterial recurrence.</li> </ul>
Antibiotic courses should be completed as prescribed despite resolution of symptoms	<ul> <li>Shorter courses reduce risk of selective pressure for resistant organisms</li> <li>There is no evidence that taking antibiotics beyond symptom resolution reduces antimicrobial resistance. There is evidence however that longer durations of therapy drive resistance by increasing selective pressure.<sup>62,63</sup></li> <li>Symptoms can persist due to inflammation and not bacteria.</li> <li>Shorter courses reduce antimicrobial cost</li> <li>Drug-acquisition cost savings are estimated to be \$678–798 million CAD/year in North America if treatment was shortened from 14 days to 7 days for Gram negative bacteremia.<sup>64</sup> There would be additional cost savings with shorter duration treatments of other infections.</li> </ul>
There is no harm in taking an antimicrobial for a longer duration	<ul> <li>Shorter courses reduce patient exposure to antibiotics and risk of associated adverse events, and shorten hospital length of stay</li> <li>Longer durations of antibiotic therapy put patients at increased risk of adverse events including <i>C. difficile</i> infection, which is associated with significant morbidity and mortality.<sup>60-62</sup></li> <li>There is a 4-5% increased odds of a patient experiencing an antibiotic-associated adverse event with each excess day of antibiotic use.<sup>60,62</sup></li> </ul>

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Prepared by: Amor-Myville Blanco, 4th year PharmD Student;

o, 4<sup>th</sup> year PharmD Student; amor<mark>myvi@ualberta.cs</mark> and Susan Fryters, AS/ID pharmacist, Reviewed by: Dr. Isabelle Chiu, Infectious Diseases physician, Edmonton Zone

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Prepared by: Amor-Myville Blanco, 4<sup>th</sup> year PharmD Student; Reviewed by: Dr. Isabelle Chiu, Infectious Diseases physician, Edmonton Zone

and Susan Fryters, AS/ID pharmacist,

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Prepared by: Amor-Myville Blanco, 4th year PharmD Student;

#### o, 4<sup>th</sup> year PharmD Student; amormyvi@ualberta.ca and Susan Fryters, AS/ID pharmacist, s Reviewed by: Dr. Isabelle Chiu, Infectious Diseases physician, Edmonton Zone

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