



New UTI Guidelines: Making Men Less Complicated Since 2025

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Disclosures

- I have no relevant financial relationships to disclose

Learning Objectives

1. Identify patient populations who are no longer considered "complicated" when treating UTI's
2. Be able to recognize a patient with a UTI as complicated vs uncomplicated
3. Assess first line, empiric options in the treatment of cUTI's

Abbreviations

- cUTI: Complicated urinary tract infection
- CAUTI: Catheter associated urinary tract infection
- IEAT: Inappropriate empiric antimicrobial therapy
- AEAT: Appropriate empiric antimicrobial therapy
- S/S: Signs and symptoms
- FQ: Fluoroquinolone
- SMP-TMX: Sulfamethoxazole-trimethoprim
- IDSA: Infectious Disease Society of America

New Classifications

New classifications of uUTI and cUTI

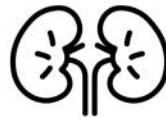
Old Classifications

Uncomplicated UTI:

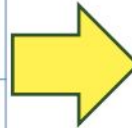
Acute cystitis in afebrile nonpregnant premenopausal women with no diabetes and no urologic abnormalities



Acute Pyelonephritis: Acute kidney infection in women otherwise meeting the definition of uncomplicated UTI above



Complicated UTI: All other UTIs



New Classifications

Uncomplicated UTI: Infection confined to the bladder in afebrile women or men

Complicated UTI: infection beyond the bladder in women or men

- Pyelonephritis
- Febrile or bacteremic UTI
- Catheter-associated (CAUTI)
- Prostatitis* (*not covered by these guidelines)

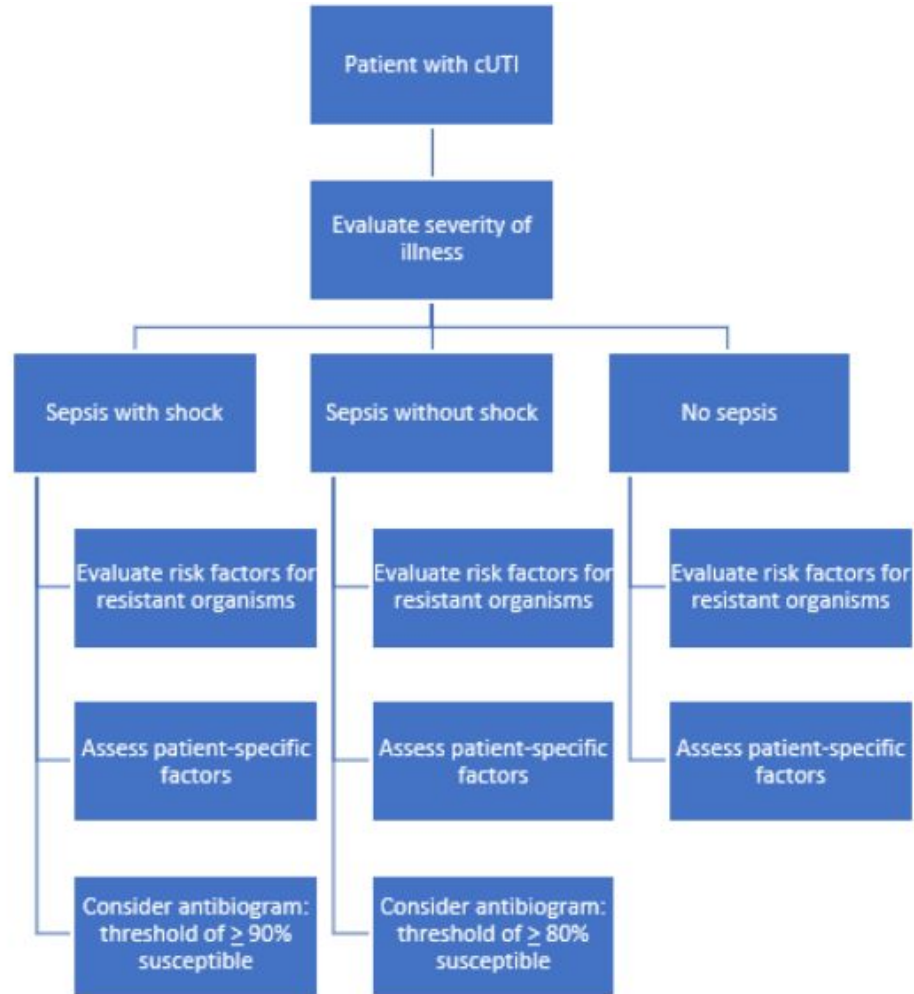


Complicated vs. Uncomplicated

Uncomplicated UTI classifications

- Clinical presentation:
 - Local bladder signs and symptoms such as dysuria, urgency, frequency, and suprapubic pain
 - Presumed to be confined to the bladder and **absence** of:
 - Fever, chills, rigors, or unstable vital signs (unless explained by a non-UTI cause)
 - Flank pain
 - Costovertebral angle tenderness
- Populations: (important update)
 - Females or **males**
 - **Underlying urologic abnormalities**
 - **Immunocompromised**
 - **Diabetes**
 - **Recurrent UTI**

Four-step Approach for Therapy



Assess Severity of Illness

Step 1: Severity of illness

- Sepsis with shock vs. Sepsis without shock vs. No sepsis
- What factors helped determine severity levels?
- Impact of inappropriate empiric antibiotic therapy (IEAT)
 - 10.8% mortality rate (ranged from **10-50%**)
 - 9.0% mortality rate in AEAT group

Optimization of Coverage

Step 2: Assess patient-specific risk factors for resistant bacteria

- Recommendation: avoid empiric coverage for resistant pathogens from previous urine cultures greater than 90 days
- Suggest avoiding fluoroquinolone use if patient has received in the past 12 months

Good Practice Statement

Step 3: Remember the patient

- Allergies
- Contraindications
- Drug-drug interactions
- Cost

Antibiogram

Step 4: Utilizing antibiogram for antibiotic selection

- Panel suggests using antibiogram for patients WITH sepsis
- Needs to be local and relevant
- Thresholds:
 - $\geq 90\%$ susceptibility in septic shock
 - $\geq 80\%$ in sepsis without shock
- Avoid routine use of broad-spectrum agents in suspected cUTI without sepsis

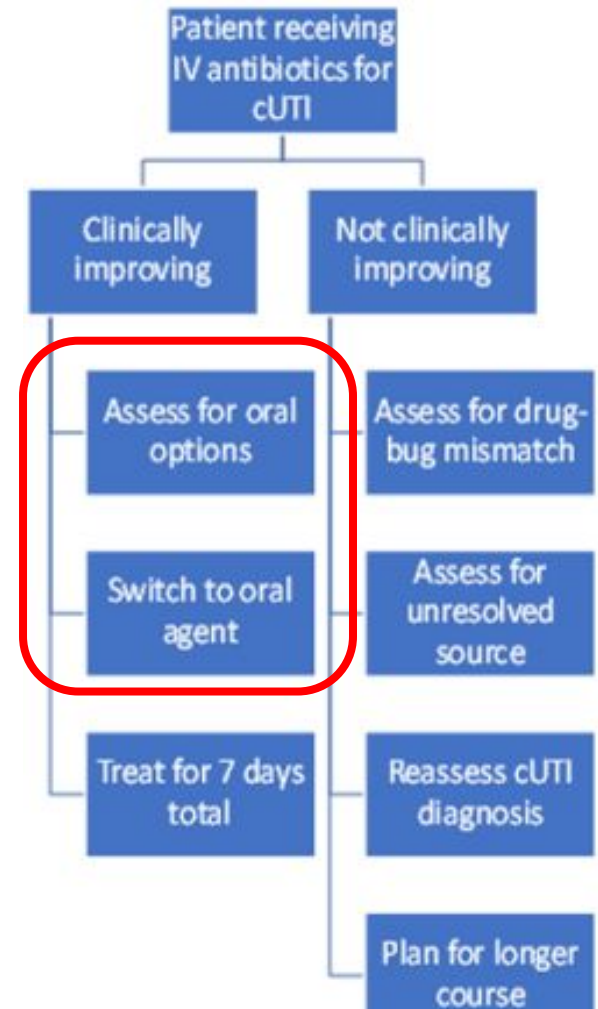
Empiric Recommendations

Condition of the Patient	Preferred	Alternative
Sepsis with or without shock	3rd or 4th generation cephalosporins, <u>carbapenems</u> , piperacillin-tazobactam, fluoroquinolones	Newer beta lactam/lactamase inhibitors, cefiderocol, or older aminoglycosides
Without sepsis- IV	3rd or 4th generation cephalosporins, piperacillin-tazobactam, fluoroquinolones	<u>Carbapenems</u> , newer beta lactam/lactamase inhibitors, cefiderocol, or older aminoglycosides
Without sepsis- PO	Fluoroquinolones or sulfamethoxazole-trimethoprim	Amoxicillin-clavulanate or cephalosporins

Transition Timing

Recommend switching from
IV to PO in patients

- Who are clinically improving on IV therapy
 - Can tolerate PO medications
 - And have an appropriate PO option



Optimal Oral Dosing

Drugs	Oral absorption (%)	Urinary excretion (%)	Dose with normal renal function
Amoxicillin-clavulanate	80/ variable	50-70/ 25-40	875mg-125mg every 8 to 12 hours Other regimens may be more effective*
Cefixime	50	50	400mg QD
Cefpodoxime	50	80	200-400mg Q12H
Ceftibuten*	75-90	73	400mg QD or 200mg Q12H (adult) 8mg/kg/day (children)
Cefuroxime	52	90	500mg Q12H
Cephalexin	90	90	500-1000mg Q6H Other regimens may be more effective*
Ciprofloxacin	70	40-50	500-750mg Q12H
Levofloxacin	99	64-100	500-750mg QD
Sulfamethoxazole-trimethoprim	70-90	84/66	800/160mg (DS) Q12H

-Other oral beta-lactams (amoxicillin, cefadroxil, cefaclor, cefdinir) found to be less efficacious.

Duration

Overall recommendation: patients who are clinically improving should be treated with a shorter 5-7 day course, rather than a longer course of 10-14 days

5-7 days of a fluoroquinolone

7 days of a non-fluoroquinolone

Duration of Treatment Reasoning

- Inclusion criteria
 - Patient population
 - Intervention vs comparator group
- 10 RCT's totaling
 - 909 males (33.9%)
 - 1772 females (66.1%)
- Majority of studies excluded chronic conditions

Certainty assessment							№ of patients		Effect		Certainty	Importance
№ of studies	Study design	Risk of bias	Inconsistency	Indirectness	Imprecision	Other considerations	Shorter duration of Abx (5 to 7 days)	Prolonged duration of Abx (10 to 14 days)	Relative (95% CI)	Absolute (95% CI)		
Clinical cure (at Test-of-Cure (TOC))												
10 ¹⁻¹⁰	randomised trials	serious ^a	not serious ^b	not serious	not serious ^c	none	903/1014 (89.1%)	962/1096 (87.8%)	RR 1.00 (0.97 to 1.04)	0 fewer per 1,000 (from 26 fewer to 35 more)	⊕⊕⊕○ Moderate	CRITICAL
Microbiological cure (at Test-of-Cure (TOC))												
10 ¹⁻¹⁰	randomised trials	serious ^d	not serious ^b	serious ^e	not serious ^c	none	778/915 (85.0%)	824/975 (84.5%)	RR 0.99 (0.94 to 1.05)	8 fewer per 1,000 (from 51 fewer to 42 more)	⊕⊕○○ Low	IMPORTANT
Recurrence of Infection (up to 180 days)												
6 ^{1,3,5,7,9,10}	randomised trials	serious ^a	not serious ^f	not serious	not serious ^c	none	41/535 (7.7%)	38/548 (6.9%)	RR 1.07 (0.69 to 1.65)	5 more per 1,000 (from 21 fewer to 45 more)	⊕⊕⊕○ Moderate	CRITICAL

Fluoroquinolone Duration

Certainty assessment							No of patients		Effect		Certainty	Importance
No of studies	Study design	Risk of bias	Inconsistency	Indirectness	Imprecision	Other considerations	Shorter duration of FQ (5 to 7 days)	Prolonged duration of FQ (10 to 14 days)	Relative (95% CI)	Absolute (95% CI)		

Clinical cure (at Test-of-Cure (TOC))

7 ¹⁻⁷	randomised trials	serious ^a	not serious	not serious	not serious ^b	none	744/851 (87.4%)	820/935 (87.7%)	RR 0.98 (0.96 to 1.01)	18 fewer per 1,000 (from 35 fewer to 9 more)	⊕⊕⊕○ Moderate	CRITICAL
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Microbiological cure (at TOC)

7 ¹⁻⁷	randomised trials	serious ^c	not serious	serious ^d	not serious ^b	none	625/752 (83.1%)	689/824 (83.6%)	RR 0.98 (0.93 to 1.03)	17 fewer per 1,000 (from 59 fewer to 25 more)	⊕⊕○○ Low	IMPORTANT
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Recurrence of Infection (up to 90 days)

4 ^{1,3,5,7}	randomised trials	serious ^a	not serious	not serious	not serious ^b	none	32/480 (6.7%)	34/494 (6.9%)	RR 0.94 (0.59 to 1.51)	4 fewer per 1,000 (from 28 fewer to 35 more)	⊕⊕⊕○ Moderate	CRITICAL
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- Shorter fluoroquinolone course likely does not
 - Reduce clinical cure
 - Reduce microbiological cure
 - Increase recurrence of infection

Pyelonephritis

Follow overall recommendation:

5-7 days of a fluoroquinolone

7 days of a non-fluoroquinolone

Recommendation derived from GRADE evidence profile

Because all studies incorporated pyelonephritis as a primary characteristic, stratification was not required.

Febrile or Bacteremic UTI

Febrile UTI

- 5-7 days of fluoroquinolone
- 7 days of non-fluoroquinolone

Bacteremic UTI

- 7 days of an effective antibiotic
 - Needs to achieve therapeutic levels in the blood, urine, and relevant tissue

Certainty assessment							№ of patients		Effect		Certainty	Importance
№ of studies	Study design	Risk of bias	Inconsistency	Indirectness	Imprecision	Other considerations	Shorter duration of Abx (≤7 days)	Prolonged duration of Abx (> 7 days)	Relative (95% CI)	Absolute (95% CI)		
Relapse of bacteremia (at 30 days)												
3 ¹⁻³	RCTs	serious ^a	not serious	not serious	serious ^b	none	13/391 (3.3%)	9/367 (2.5%)	RR 1.31 (0.57 to 3.02)	8 more per 1,000 (from 11 fewer to 50 more)	⊕⊕○○ Low	CRITICAL
Mortality (at 30 days)												
3 ¹⁻³	RCTs	serious ^a	not serious	not serious	serious ^c	none	14/390 (3.6%)	14/367 (3.8%)	RR 0.93 (0.30 to 2.91)	3 fewer per 1,000 (from 27 fewer to 73 more)	⊕⊕○○ Low	IMPORTANT
Mortality (at 90 days)												
3 ¹⁻³	RCTs	serious ^a	serious ^d	not serious	serious ^e	none	36/390 (9.2%)	36/367 (9.8%)	RR 0.94 (0.37 to 2.37)	6 fewer per 1,000 (from 62 fewer to 134 more)	⊕○○○ Very Low	IMPORTANT

Now, What About Catheters.....

- Following the new classifications, a catheter-associated UTI (CAUTI) is considered complicated
- Guidelines also acknowledge that a subset of patients with CAUTI **without** systemic symptoms may be treated as simple cystitis.
- Treatment duration
 - Uncomplicated (local S/S): 3-5 days
 - Complicated (systemic S/S): 5-7 days

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Prostatitis

- Men with febrile UTI in whom acute bacterial prostatitis is suspected may benefit from a longer treatment duration (i.e., 10-14 days), although evidence to guide the optimal duration in this subgroup is lacking.

Management of prostatitis not included in updated guidelines

Expert opinion still recommends to treat for 4-6 weeks

Study (Lead author, Year of publication, Name of trial, Countries)	Males included (No, %)	Exclusion / Inclusion criteria-based on presence/ absence of involvement of prostate/ epididymis	Stratified analysis for male with/without prostatitis	Relative estimate of clinical cure in the whole population	Relative estimate of clinical cure in men	Relative estimate of clinical cure in men with suspected acute bacterial prostatitis
Peterson 2008 USA (multicentric)	427 (39%)	Excluded if presence of acute bacterial prostatitis or epididymitis	NR	RR 1.05 (0.97-1.14)	NA	NA
Rudrabhatla 2018 India	24 (41%)	Excluded if evidence of prostatitis or prostatic abscess	NR	RR 1.00 (0.92 to 1.09)	NA	NA
Darouiche 2014 USA	52 (95%)	NR	NR	RR 1.00 (0.93 to 1.07)	Likely very similar to the whole population	NA
Ren 2017 China (multicentric)	40 (15%)	NR	NR	RR 1.01 (0.93 to 1.08)	NA	NA
Wagenlehner 2018 Germany and Poland	40 (18%)	NR	NR	RR 1.09 (0.96 to 1.23)	NA	NA
Lafaurie 2023 PROSTA-SHORT France (multicentric)	240 (100%)	Males with acute prostatitis included. Acute prostatitis was diagnosed based on pain on rectal examination, which was not systematically performed	Post-hoc analysis presence / absence of pain on rectal examination	RR 0.96 (0.92 to 1.00)	RR 0.96 (0.92 to 1.00)	In a subset of 27 men with pain on rectal examination, RR 0.77 (0.49 to 1.20) *
van Nieuwkoop 2017 FUTIRST Netherlands (multicentric)	86 (43%)	Males with acute prostatitis included	Randomized stratification for gender	RR 0.95 (0.88 to 1.03)	RR 0.88 (0.78 to 1.00)	NR

Bactrim's Role

- Majority of trials addressed FQ use
- Updated guidelines recommend 7 days of Bactrim (1 DS tab BID) for cUTI
 - Recommendation comes from lack of high-quality RCT's that specifically looked at a shorter durations for treatment
- Several RCT's exist for uncomplicated UTI's comparing Bactrim, FQ's, and oral beta-lactams

Final Remarks

- Men are not complicated
- Future outlook
 - PO options
 - Prostatitis
 - Epididymitis/orchitis

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Thank You

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