Urinary Tract Infections

- Symptomatic and asymptomatic UTI's are a common problem
- > 10 million office visits per year
- > 1 million hospital admissions per year
- Cause of significant nosocomial morbidity
- Affects women more than men throughout life

UTI's in Adults

- Acute uncomplicated UTI in young women
- · Acute uncomplicated pyelonephritis
- Recurrent UTI's in women
- Complicated UTI's in older women
- Catheter-associated bacteriuria
- Asymptomatic bacteriuria
- Candiduria











Epidemiology of Urinary Tract Infections by Age Group

		Females		Males
Age	Prevalence	Risk Factor	Prevalence	Risk Factor
<1	1%	Anatomic or functional urologic abnormalities	1%	Anatomic or functional urologic abnormalities
1-5	4.5%	Congenital abnormalities, vesicoureteral reflux	0.5%	Congenital abnormalities uncircumcised penis
6-15	4.4%	Vesicoureteral reflux	0.5%	None
16-35	20%	Sexual activity, diaphragm use, spermacide	0.5% s	Homesexual activity, anal intercourse
36-65	35%	Gynecologic surgery, bladder prolapse	20%	BPH, obstruction, catheterization, surgery
>65	40%	All of above, incontinence, chronic catheterization	35%	All of above, incontinent chronic catheterization, condom catheters



Microbial Species Most Often Associated with Specific Types of UTI's

Organism	Acute uncomplicated cystitis	Acute uncomplicated pyelonephritis	Complicated UTI	Catheter- associated UTI
E.coli	79%	89%	32%	24%
S. saprophyticus	11%	0%	1%	0%
P. mirabilis	2%	4%	4%	6%
Klebsiella spp.	3%	4%	5%	8%
Enterococcus spp.	2%	0%	22%	7%
Ps. aeruginosa	0%	0%	20%	9%
Mixed	3%	5%	10%	11%
Other*	0%	2%	5%	10%
Candida spp.	0%	0%	1%	28%
S. epidermidis	0%	0%	15%	8%





UTI in Women - Host Factors

- Short urethra
- Vaginal colonization
- Diaphragm / vaginal spermacide
- Sexual intercourse
- Delayed post-coital voiding
- P₁ blood group upper UTI

UTI - Other Host Factors

- Extra-renal obstruction
 - posterior urethral valves
 - urethral strictures
- prostatic hypertrophy
- Neurogenic bladder
- · Vesico-ureteral reflux
- Catheterization/instrumentation

Urinary Tract Infections

- The initial pathogenic event in UTI is an encounter between bacteria and host mucosa at the tissue surface
- Attachment, binding of bacteria to mucosal cells, is the result of multiple interactions between bacterial surface ligands (adhesins) and epithelial cells (receptors).



Anti-adherence Mechanisms in the Urinary Tract

- Normal bacterial flora of vaginal, introital, and periurethral region and urethra
- Uromucoid (Tamm-Horsfall protein)
- · Urinary oligosaccharides
- Urinary immunoglobulins (IgG, IgA, S-IgA)
- Bladder mucopolysaccharide (glycosaminoglycan)
- Mechanical effects of flushing



UTI - Bacterial Factors - 1

• Attachment

- Type 1 fimbriae (MS-adhesins) attach to mannosides on urothelial cell
 P fimbriae attach to globoseries receptors on urothelial cell these strains cause pyelonephritis
 - 97% of women with recurrent pyelo are P1 blood group (+)
 - · women with pyelo due to VU reflux same prevalence of P1 as gen. pop.
- Afimbrial adhesins (AFA I, AFA III)

Toxins

- RTX hemolysins protein toxins that contain a tandem duplication of 9 amino acids (cause pores in cell membrane, lysis)
- E. coli that do not produce these toxins are less virulent
- · Phase variation
 - Type I down-regulated, Type P upregulated in strains that cause uppertract infections (PAP gene expression triggered by temperature, [glucose], concentration of certain amino acids.

UTI - Bacterial Factors - 2

Internalization

- enters bladder cells, protected from antibody, phagocytes
- intracellular persisters ??source of recurrent infection
- Doubling time
 - if <50 60 minutes, increased ability to cause cystitis
 - E. coli bowel strains that do not cause UTI's generally have slower doubling times
- · Serum-resistant capsules
- Anti-phagocytic mechanisms (e.g., P-fimbriae)
- Iron acquisition efficiency is a virulence factor
- uropathogenic strains may have multiple sequestration systems

UTI - Clinical

• Children

- < 2 years - enuresis, fever, poor weight gain

-> 3 years - dysuria, lower abdominal pain

• Adults

- urgency, frequency, dysuria, cloudy or malodorous urine, bladder or flank pain
- Pyelo: fever >101 F, chills (bacteremia), flank pain and tenderness







Urinary Catheters

- · Foreign body
- Biofilm formation
 - bacteria, bacterial glycocalyces, host proteins, urinary salts (apatite and struvite)
- · Sanctuary site for bacteria
- Condom catheters carry same risk of infection as indwelling (Foley) catheters
- 100% become infected in 7-10 days

Bacteriuria in the Catheterized Patient

- Avoid use of antimicrobials, if possible
- · Indications for treatment
 - symptomatic infection
 - suspected sepsis
 - renal transplant
 - immunocompromised patient
- pre-operative patient
- · Remove or change catheter during treatment

UTI - Diagnostic Criteria

- U/A microscopic quantitative
- Leukocyte esterase test
- Nitrate \rightarrow nitrite test
- Leukocyte esterase / nitrate test
- Gram's stain, unspun urine

UTI - Diagnostic Criteria

- Collection: clean midstream specimen or straight-catheterized specimen
- >10 WBC/ μ L in symptomatic female
- (+) Gram's stain of unspun urine
- Culture criteria
 - $->10^5$ CFU/mL = infection
 - symptomatic female: 10^2 - 10^4 CFU/mL of
 - E. coli, Proteus, S. saprophyticus are significant





Indications for Evaluating the Urinary Tract

Children

- ultrasound, IVP, VCUG
- Bacteremic pyelonephritis – ultrasound, or IVP
- Nephrolithiasis or Neurogenic Bladder – ultrasound, or IVP with post-voiding films
- Men with 1st infection - careful prostate examination
- Men with 2nd infection

 ultrasound or IVP with post-voiding films

General Principles of Treatment

- Quantitative cultures may be unnecessary before treatment of typical cases of acute uncomplicated cystitis.
- Susceptibility testing is necessary in all recurrent or complicated infections, perhaps not for uncomplicated cases.
- Identify or correct factors predisposing to infection (obstruction, calculi)
- relief of symptoms may not indicate bacteriologic cure: follow-up cultures are indicated if symptoms recur.
- Duration of therapy depends on the site and duration of the infection.
- · Classify recurrences as re-infection or relapse.





Treatment of Asymptomatic Bacteriuria

- Pregnancy
- Neurological or structural abnormality of the urinary tract with $> 10^5 \mbox{ CFU/mL}$
- Pre-op for GU (and other?) surgery

