Subclinical Bacteriuria

- The presence of bacteria in urine as determined by a positive culture from a properly collected urine specimen, in the absence of clinical evidence of infectious urinary tract disease
 - NOT just on sediment exam
- Reported in 2.1-12% of healthy dogs
 - 15-74% in some populations with predisposing factors
 - DM, obesity, parvo, IVDD, immunosuppressive therapy
- Cats reported in 1-13% of healthy cats
- No evidence of association with going on to develop urinary tract signs or alter survival without treatment

Journal of Feline Medicine and Surgery, (2019) 21, 1023–1038



Roswitha Dorsch, Svenja Teichmann-Knorrn and Heidi Sjetne Lund

Subclinical Bacteriuria

- What do the Human Doctors do?
 - They don't treat!
 - Even in high-risk groups!!
 - Even with malodorous urine or with pyuria...
 - Largely focused on antimicrobial stewardship
 - They do treat if there is urological surgery planned
- But what happens when they do?
 - They eliminate the current bacteriuria but...
 - Patients are quickly recolonized
 - Often with significantly higher recurrence rates than those not treated
 - Treated patients often have higher rates of AMR *E. coli* in subsequent UTIs

Subclinical Bacteriuria - Diagnosis

- Why are you looking?
 - No indication to culture in animals that lack lower urinary tract signs when there is no indication to treat if a positive result is found
 - So, not as part of annual wellness visit
 - Even in patients with DM of HAC
 - But indicated as part of an investigation of PU/PD, for instance
- Cell count cannot differentiate subclinical from cystitis
- Does not depend on the presence or absence of pyuria
- Retesting not recommended
 - Don't need to "see if its gone" or "if its still there" if no indication to treat!

Subclinical Bacteriuria – Treatment????

- In patients where it is unclear if there are clinical signs
 - Treat with 3-5 days of an appropriate antimicrobial
 - If no response, then a true infection is unlikely and stop treatment
- Patients that *can't* display signs
 - Requires clinical judgement
 - Paralyzed patients or those that can't mount a fever response
- If pyuria present, we USED to recommend treatment
 - In people, this is not recommended
 - No evidence in veterinary medicine that suggests we should act differently
- In the VERY RARE event that the urinary bacteria is thought to be a nidus for extra-urinary infection

Subclinical Bacteriuria – Treatment???

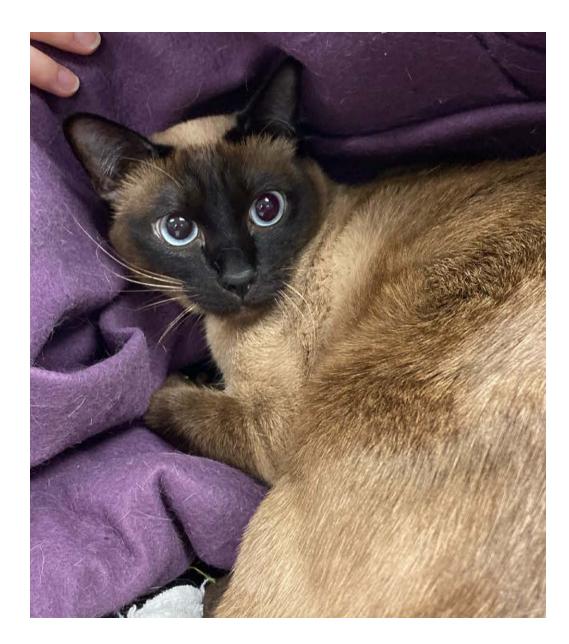
- Multidrug resistant organisms
 - This should NOT prompt treatment
 - They do not do more damage or make one more likely to develop a true infection
 - Some anecdotal evidence that, over time, these bacteria may be replaced by more susceptible organisms if treatment is withheld
- Plaque-forming and urease producing organisms
 - Can be associated with encrusting cystitis and urolith formation
 - IF those things are present, it is a reason to treat
 - Unknown if they will cause these things to happen
 - Can be difficult to treat even when we want to... (*Corynebacterium*)

Subclinical Bacteriuria – Treatment???

- What if the patient previously had subclinical bacteriuria, but is now showing clinical signs?
 - Depends on the timing since the last culture, but it is unknown how likely it is to be the same organism causing the clinical signs
 - Recommend reculture to direct therapy appropriately
- What about supplements?
 - Cranberry extracts, probiotics
 - No evidence that these will prevent or resolve subclinical bacteriuria
 - No real contraindication either

Teemo

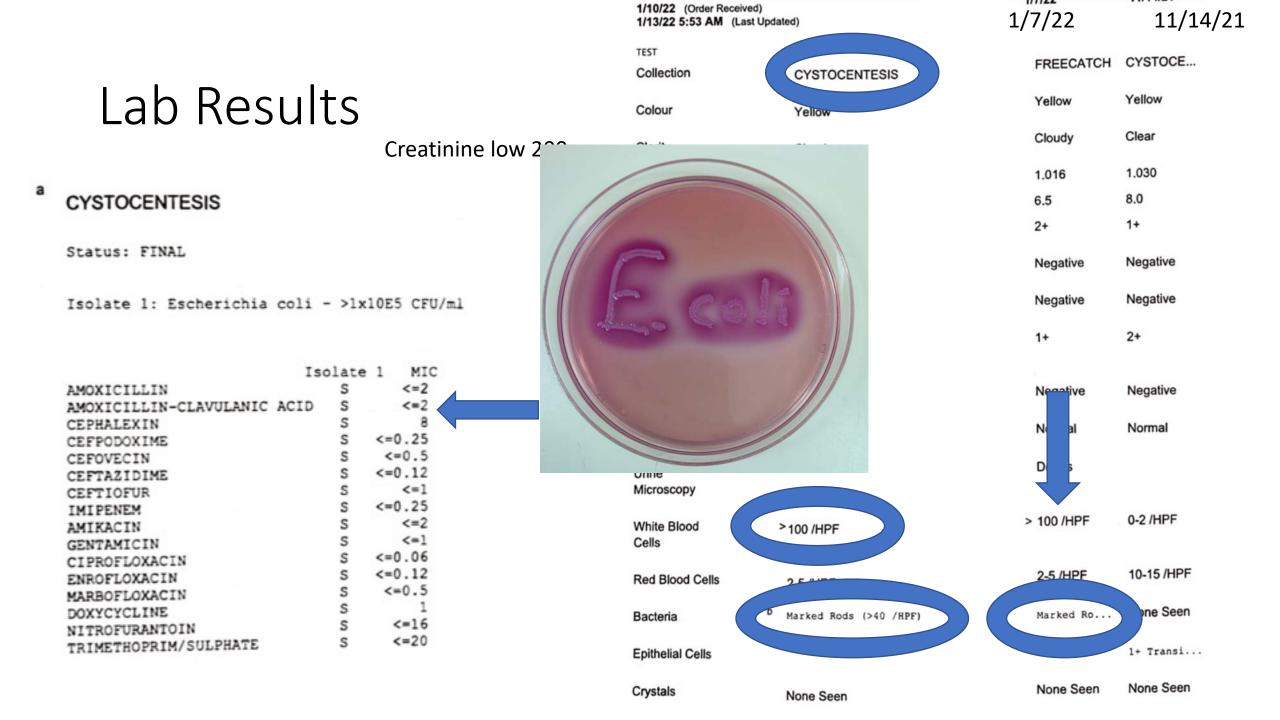
- 3-year-old MN Siamese
- Referred for recurrent UTIs
- Previous History
 - Initial episode almost 2 years ago
 - Stranguria
 - ER visit Tx supportive care, no ABs
 - Follow up
 - CBC/Chemistry mild increase in creatinine (low 200s) and SDMA
 - UA pyuria, bacteriuria



Teemo

- Recent History
 - Dec 2021
 - Owners had big event at home and Teemo developed stranguria after this
 - Seen by an ER clinic and he had small bladder, treated supportively for FIC
 - Follow up UAs and culture
 - Bacteriuria, pyuria
 - The previous elevated creatinine persisted
 - Treated with prolonged course of Clavamox for possible pyelonephritis (6 weeks)
 - Last week:
 - Saw me owners report him as being completely normal at home. Only urinary signs had been those two episodes, nothing persistent at home
- Physical Exam
 - Normal





URINALYSIS			
TEST	RESULT	REF.RAM	IGE/UN
Collection	CYSTOCENTESIS		
Colour	Yellow		
Clarity	Cloudy		
Specific Gravity	1.025		No Cl
pH	5.5		Signs
Urine Protein	Negative		
Glucose	Negative		
Ketones	Negative		
Blood / Hemoglobir	Negative		
Bilirubin	Negative		
Urobilinogen	Normal		
White Blood Cells	30-50 /HPF		
Red Blood Cells	None Seen		
Bacteria a	Marked Rods (>40 /H	HPF)	
Epithelial Cells	1+ Transitional epithe /HPF)	elial cell	s (1-2
Crystals	None Seen		

REF.RANGE/UNITS OCENTESIS

No Clinical Signs at home TEST: UCUL Source: b Site: Status : Culture Results:

Organism 1:

CYSTOCENTESIS

CYSTO

FINAL

Escherichia coli - >1x10E5 CFU/ml

	Organism 1		
	Escherichia coli		
Antibiotic	Interp.	MIC µg/mL	
Amoxicillin	S	4	
Amoxicillin / Clavulanic Acid	S	<=2	
Imipenem	S	<=0.25	
Cephalexin	S	8	
	Organism 1 Escherichia coli		
Antibiotic	Interp.	MIC µg/mL	
Cefovecin	S	<=0.5	
Cefpodoxime	S	<=0.25	
Ceftazidime	S	<=0.12	
Ceftiofur	S	<=1	
Amikacin	S	<=2	
Gentamicin	S	<=1	
Ciprofloxacin	S	<=0.06	
Enrofloxacin	S	<=0.12	
Marbofloxacin	S	<=0.5	
Doxycycline	S	1	
Nitrofurantoin	S	<=16	
Trimethoprim / Sulfa	S	<=20	

I = Intermediate R = Resistant S = Sensitive TF = To Follow N/I = Not Indicated See NOTES section for more information.

Abdominal Ultrasound

- Get images from SmartPACS
- Very mild chronic changes to the kidneys
- No ureteral changes
- Normal urinary bladder
 - No wall thickening
 - No debris in bladder
 - No uroliths
 - No mass







Key Features of Case

- Early stage CKD
 - Mild increases in SDMA and Creatinine, normal BUN
 - Poorly concentrated USG, but not isosthenuric
- Persistent bacteriuria
 - Pyuria
- Cat is asymptomatic!
 - Has only had true lower urinary tract signs 2 times
 - Episodes more consistent with FIC or FLUTD
 - Identifiable stressful causes
 - Quick resolution of clinical signs



Diagnosis?

- Subclinical bacteriuria
- Treatment recommendations
 - Do not treat with antimicrobials
 - Renal diet
 - Home management as for FIC cats
 - In the future?
 - If there are lower urinary tract signs, consider therapy
 - Also consider recurrent FIC



- Monitoring
 - Check chemistry panel and USG in 4-6 months
 - To monitor CKD
 - Should we check a UA?
 - Maybe for pH
 - What if there is pyuria/bacteriuria
 - Culture?
 - Only if you think you'd treat...

Case 2

Recurrent Bacterial Cystitis



Remember Predisposing causes/factors

- Anatomical factors
 - Ectopic ureters
 - Persistent vaginal membranes
 - Recessed/hooded vulvas
 - Prostate
 - Pelvic Bladder
- Neurological disease
 - Inability to empty bladder
 - USMI
- Inflammatory Disease
 - Polypoid cystitis
 - Proliferative urethritis

- Concurrent Disease
 - Urolithiasis
 - CKD
 - DM
 - HAC
 - General PU/PD
- Immunosuppression
 - Drugs
 - Chemotherapy

Zoey

- 6-month-old female Labrador cross
- Referred for recurrent UTIs
 - Multiple courses of treatment
 - Never cleared or immediate return
- Incontinent since adoption
 - Puddles behind her after lying/sleeping



Lab R	esults	5
URINALYSIS		
TEST	RESULT	REF.RANGE/UNITS
Collection	free catch	
Colour	Yellow	Reinell
Clarity	Cloudy	A Service
Specific Gravity	1.024	A State of the second
pH	6.5	
Urine Protein	Negative	
Glucose	Negative	
Ketones	Negative	Contraction of the local division of the loc
Blood / Hemoglobin	Negative	
Bilirubin	Negative	
Urobilinogen	Normal	
Urine Microscopy	Debris 1+ WBC clum	ps (1-2 /HPF)
White Blood Cells	30-50 /HPF	
Red Blood Cells	None Seen	
Bacteria *	Marked Rods	(>40 /HPF)
Epithelial Cells	(HPF)	epithelial cells (1-2
Crystals	Triple phosph	ate (1-5 /HPF)

MICROBIOLOGY

TEST: CUL Source: b	FREEC	ATCH		
Status : Culture Results:	FINAL			4
Organism 1 :	Klebsie	Klebsiella pneumoniae - >1x10E5 CFU/ml		
Organism 2 :	Proteu	Proteus mirabilis - 1-10x10E3CFU/ml		
	Orga	Organism 1 Organism 2		
	Klebsiella	pneumoniae	Proteu	s mirabilis
Antibiotic	Interp.	MIC µg/mL	Interp.	MIC µg/mL
Amoxicillin	R	16	S	<=2
Amoxicillin / Clavulanic Acid	5	<=2	5	<=2
Imipenem	S	<=0.25	N/I	
Cephalexin	S	<=4	1	16
Cefovecin	S	<=0.5	S	<=0.5
Cefpodoxime	S	<=0.25	S	<=0.25
Ceftazidime	5	<=0.12	S	<=0.12
Ceftiofur	5	<=1	S	<=1
Amikacin	S	<=2	5	<=2
Gentamicin	S	<=1	S	<=1
Ciprofloxacin	S	<=0,06	S	<=0.06
Enrofloxacin	S	<=0.12	S	<=0.12
Marbofloxacin	S	<=0.5	5	<=0.5
Doxycycline	S	1	R	>=16
Nitrofurantoin	1	64	R	128
Trimethoprim / Sulfa	S	<=20	S	<=20

URINALYSIS		
TEST	RESULT	REF.RANGE/UNITS
Collection	free catch	
Colour	Yellow	
Clarity	Clear	
Specific Gravity	1.028	
pH	6.0	
Urine Protein	Negative	
Glucose	Negative	
Ketones	Negative	
Blood / Hemoglobi	nNegative	
Bilirubin	Negative	
Urobilinogen	Normal	
Urine Microscopy	1+ Epithelial c 1+ WBC clum	lumps (1-2 /HPF) ps (1-2 /HPF)
White Blood Cells	20-30 /HPF	
Red Blood Cells	None Seen	
Bacteria	None Seen	
Epithelial Cells	2+ Transitiona /HPF)	al epithelial cells (3-5
Crystals	None Seen	
ORDER NOTE	S UTI	has bneen clavaseptin

-

URINALYSIS		
TEST	RESULT	REF.RANGE/UNITS
Collection	FREE CATCH	
Colour	Yellow	
Clarity	Clear	
Specific Gravity	1.029	2 -
pН	5.5	
Urine Protein	Negative	
Glucose	Negative	
Ketones	Negative	
Blood / Hemoglobi	Negative	
Bilirubin	Negative	
Urobilinogen	Normal	
Urine Microscopy	1+ WBC clumps (1-2	? /HPF)
White Blood Cells	6-10 /HPF	
Red Blood Cells	None Seen	
Bacteria	None Seen	
Epithelial Cells	1+ Squamous epithe /HPF)	lial cells (1-2
Crystals	None Seen	

PATHOLOGY

TEST: 1PATHA

Pathologist Interpretation

URINALYSIS: USG adequate. Pyuria, recommend culture.

Noted history of UTI and therapy

	1	of your		Urea (BUN)	8.4	3.2 - 11,0 mmol/L
HEMATOLOG		0		BUN: Creatinine	20	
TEST	RESULT	REF.RANGE/UNITS		Ratio		
RBC	6.5		н	Phosphorus	2.5	0.8 - 2.0 mmol/L
Hematocrit	0.44			Calcium	2.7	2.2 - 2.8 mmol/L
Hemoglobin	145	134 - 207 g/L		Sodium	146	142 - 152 mmol/L
MCV	67.7	59.0 - 76.0 fL		Potassium	4.8	4.0 - 5.4 mmol/L
MCH	22.3	21.9 - 26.1 pg		Na: K Ratio	30	28 - 37
MCHC	329.5	326.0 - 392.0 g/L	L	Chloride	110	108 - 119 mmol/L
RDW	15.5	10.0 - 19.0		TCO2	24	13 - 27 mmol/L
% Reticulocyte	1.5	%	L	(Bicarbonate)		
Reticulocytes	97.5	10.0 - 110.0 x10E3/uL		Anion Gap	17	11 - 26
Detertente	27.2	24.5 - 31.8 pg		Total Cations	151	mmol/L
Reticulocyte Hemoglobin	21.2			Total Anions	134	mmol/L
WBC	16.6	4.9 - 17.6 x10E9/L	ŀ	. Total Protein	54	55 - 75 g/L
% Neutrophils	44.9	%		Albumin	29	27 - 39 g/L
% Lymphocytes	42.8	%		Globulin	25	24 - 40 g/L
% Monocytes	5.6	%	L	Albumin: Globulin	1.2	0.7 - 1.5
% Eosinophils	6.5	%	L	Ratio	24	18 - 121 IU/L
% Basophils	0.2	%	T	ALT	19	16 - 55 IU/L
Neutrophils	7.5	2.9 - 12.7 x10E9/L	1.		171	5 - 160 IU/L
H Lymphocytes	7.1	1.1 - 5.0 x10E9/L	ľ	H ALP	5	0 - 13 IU/L
Monocytes	0.9	0.0 - 1.2 x10E9/L		GGT Bilirubin - Total	1.2	0.0 - 5.2 umol/L
Eosinophils	1.1	0.0 - 1.5 x10E9/L			7.2	3.4 - 8.9 mmol/L
Basophils	0.0	0.0 - 0.1 x10E9/L		Cholesterol		337 - 1469 IU/L
Platelets	300	143 - 448 x10E9/L		Amylase	443	0 - 250 IU/L
	ts Platelet	assessment Adequate		Lipase ^b	62	10 - 200 IU/L
Platelet Commen	Mild pla	atelet clumping observed		H Creatine Kinase	269	
				Osmolality	294	250 - 310 mmol/kg
	PPC V	VBC, and platelet		Hemolysis Index	Normal	
CBC Comment		ology normal		Icterus Index	Normal	
				Lipemia Index		
CHEMISTRY	REGULT	REF,RANGELUNITS		ENDOCRINO		OFF OANOF INTO
Glucose	5.4	3.5 - 6.3 mmol/L		TEST	RESULT 44.6	REF,RANGE/UNITS
H IDEXX SDMA *	17	0 - 14 ug/dL		Total T4 °	44.0	
Creatinine	105	44 - 133 umol/L				

URINALYSIS

TEST	RESULT	REF.RANGE/UNITS
Collection	Free Catch	
Colour	Yellow	
Clarity	Cloudy	
Specific Gravity	1.022	
pH	6.0	
Urine Protein	Negative	
Glucose	Negative	
Ketones	Negative	
Blood / Hemoglobi	nNegative	
Bilirubin	Negative	
Urobilinogen	Normal	
Urine Microscopy	1+ WBC clumps (1	-2 /HPF)
White Blood Cells	30-50 /HPF	
Red Blood Cells	None Seen	
Bacteria *	Marked Rods (>40	/HPF)
Epithelial Cells	1+ Transitional epi /HPF)	thelial cells (1-2
Crystals	None Seen	

MICROBIOLOGY

TEST: UCUL Source: ^b Site: Status : Culture Results:

Organism 1 :

CYSTOCENTESIS CYSTO FINAL

Klebsiella pneumoniae - >1x10E5 CFU/ml

	Organism 1		
	Klebsiella	a pneumoniae	
Antibiotic	Interp.	MIC µg/mL	
Amoxicillin	R	16	
Amoxicillin / Clavulanic Acid	S	<=2	
Imipenem	S	<=0.25	
Cephalexin	S	<=4	

	Organism 1		
	Klebsiella pneumoniae		
Antibiotic	Interp.	MIC µg/mL	
Cefovecin	S	<=0.5	
Cefpodoxime	S	<=0,25	
Ceftazidime	S	<=0.12	
Ceftlofur	S	<=1	
Amikacin	5	<=2	
Gentamicin	S	<=1	
Ciprofloxacin	S	<=0.06	
Enrofloxacin	S	<=0.12	
Marbofloxacin	S	<=0.5	
Doxycycline	S	1	
Nitrofurantoin	S	32	
Trimethoprim / Sulfa	S	<=20	

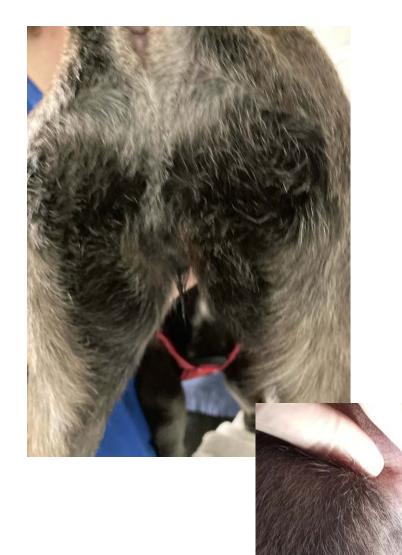
Abdominal Ultrasound

- Right kidney has mild pelvic dilation (1.4cm)
- Right ureter moderately dilated throughout
- Urinary bladder normal in appearance
 - Individual ureteral papillae not clearly visualized
 - This is often the case

An ultrasound-guided cystocentesis was obtained, when he process and complications. Diagnostic Impressions: There is moderate dilation of the right renal pelvic as well as dilation of the entire visualized right ureter, which can be traced adjacent to the urinary bladder, highly suggestive of a right-sided ectopic ureter. Consultation with a surgeon for a scope and/or contrast CT and surgical intervention is highly recommended. The urinary bladder contains a moderate intervention is highly recommended. The urinary bladder contains a moderate amount of echogenic material with some tiny sand-like debris, consistent with chronic infection. Janet A. Nieckarz-Loeven, DVM Dipl. ACVR

Physical Exam

- General exam normal
- Urinary/reproductive exam
 - Slightly recessed vulva
 - Urine leaking during exam
 - Wet fur
 - No significant erythema
- Rectal Exam
 - Normal soft urethra





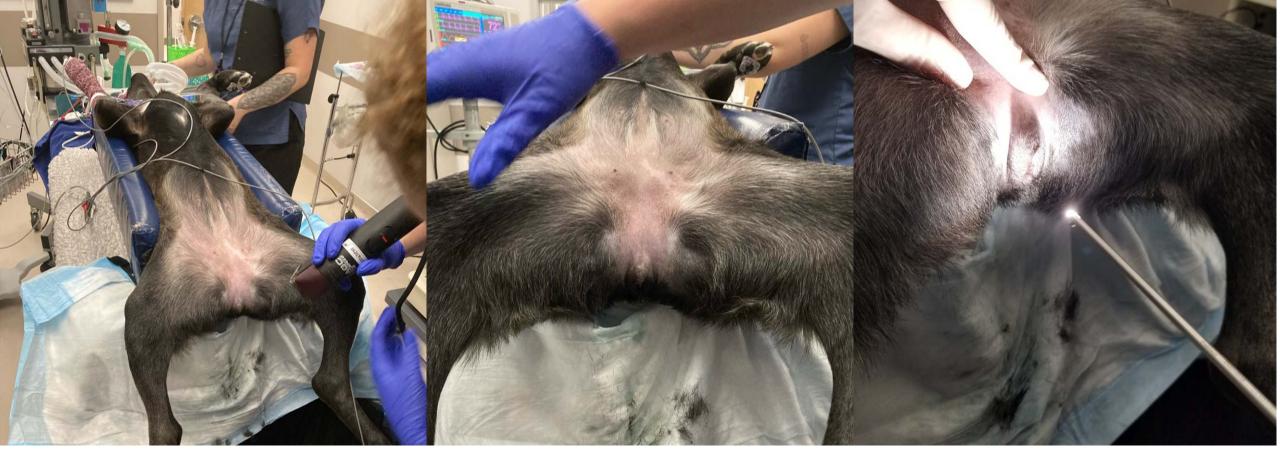
Cystoscopy Equipment





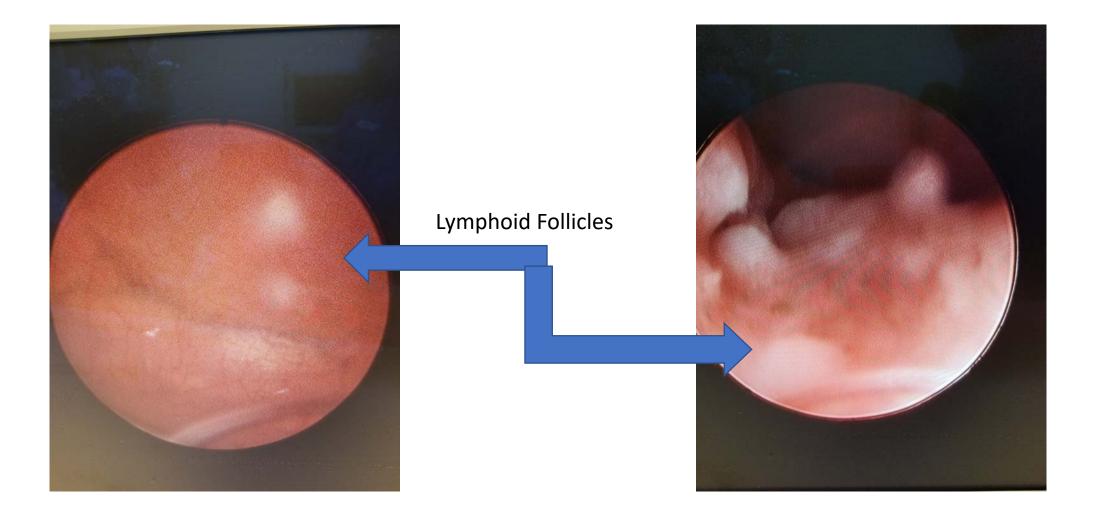
Rigid Cystoscopy



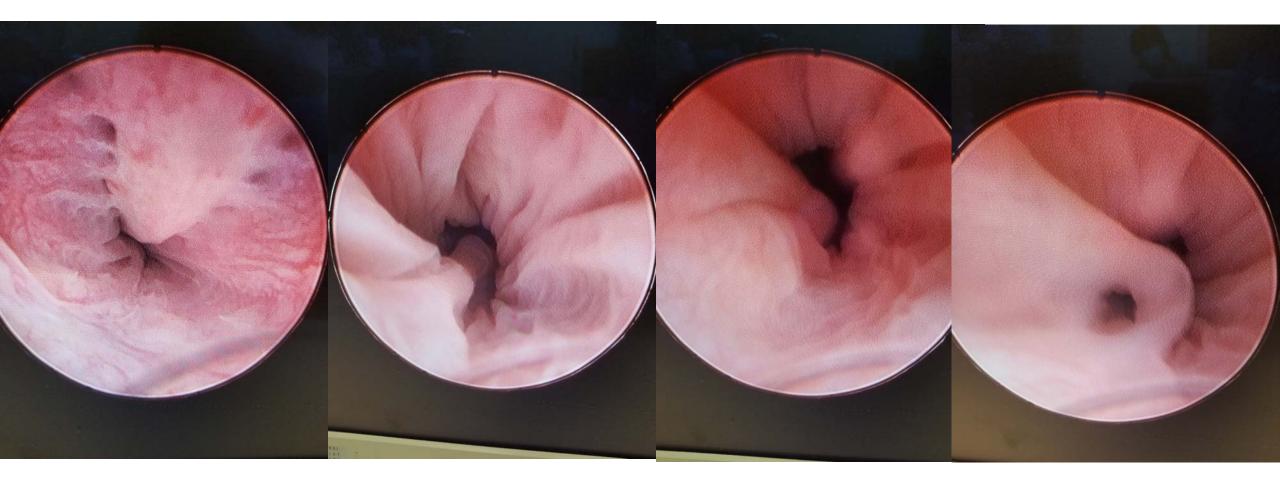


Cystoscopy Positioning

Cystoscopy - Vestibule



Cystoscopy



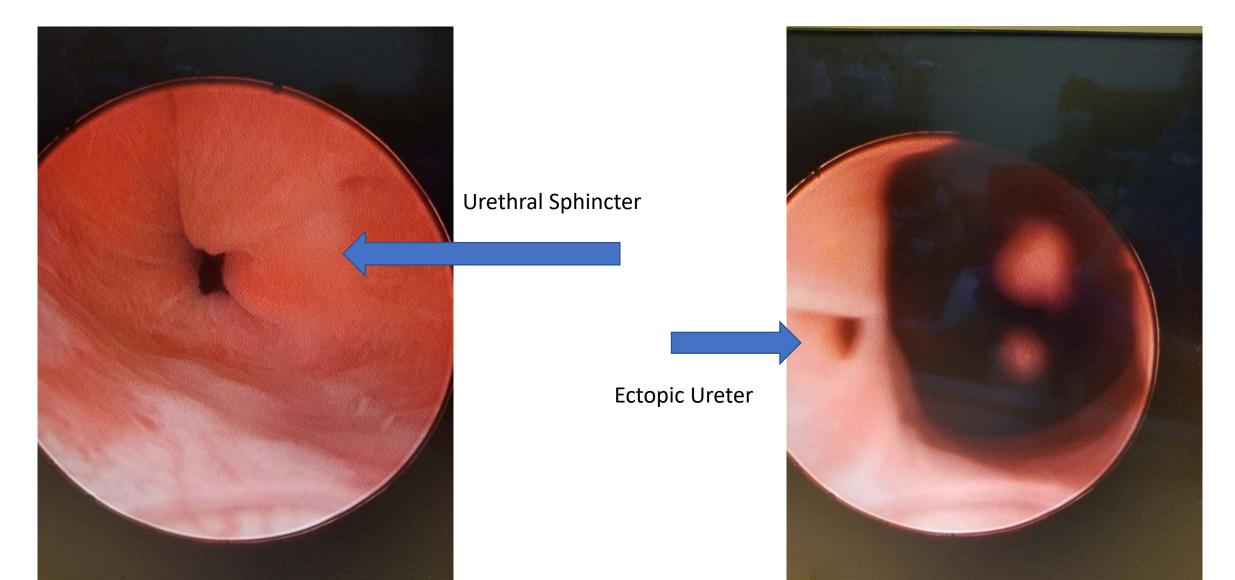
Urethral Papilla

Distal Urethra

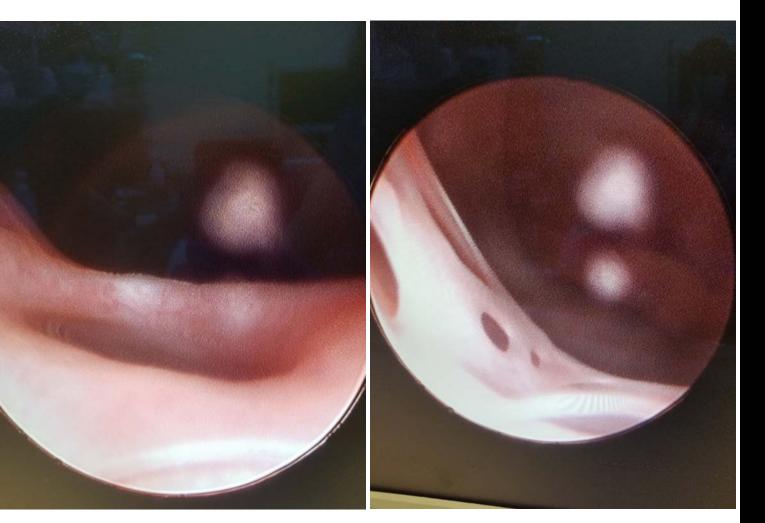
Mid Urethra

Ectopic Ureter

Cystoscopy – Urethrovesicular Junction



Too many openings!



More typical appearance of ureteral papilla

Multiple ectopic R ureteral openings



What now?

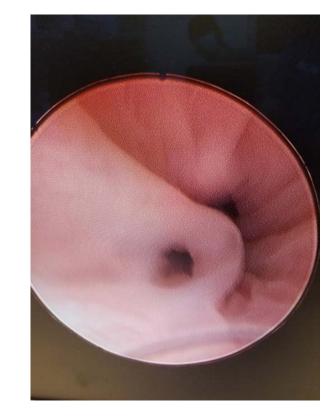
- Surgery
 - Right ureter identified (quite dilated)
 - Ligated R ureter and reimplanted in bladder
 - Effectively cuts off all the other openings
- Recovered well
- 2 days post-op still incontinent
 - Needs time to recover
 - Catheterized after surgery



- Next Steps
 - Allow for complete recovery first
- Urine culture pending
 - Antibiotic therapy for 7 more days
- If incontinence persists:
 - PPA or DES trial
 - Consider placement of hydraulic occluder

Ectopic Ureters

- Intramural (95%)
 - Open into the bladder tunnel to another location
- Extramural (5%)
 - Open into another site (urethra, vagina, uterus, vestibule)
- 32-91% are bilateral
- Signalment
 - 90% occur in young female dogs
 - Lots of predisposed breeds, but anyone can have them

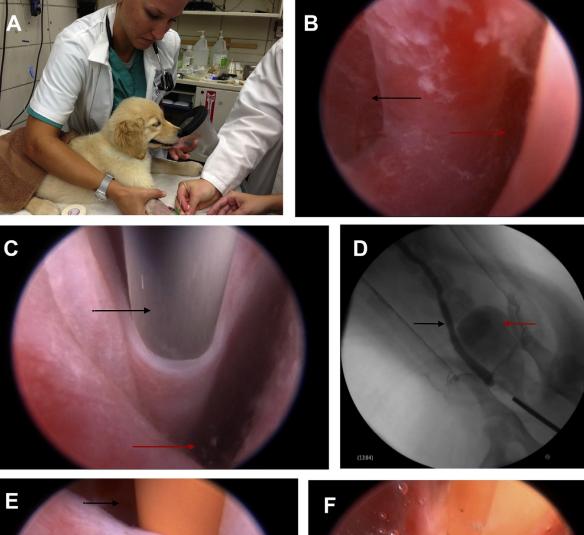


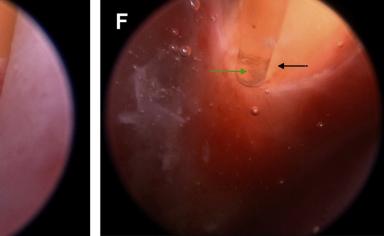
Ectopic Ureters

- Clinical signs
 - Urinary incontinence
 - Particularly when lying down/sleeping
 - Usually since birth
 - Recurrent urinary tract infections
 - Difficult to house train
- Treatment
 - Surgical reimplantation
 - Laser ablation

Canine Incontinence

Vet Clin Small Anim 49 (2019) 125–140 https://doi.org/10.1016/j.cvsm.2018.11.003





Mark J. Acierno, MBA, DVM^a, Mary Anna Labato, DVM^{b,*}

Persistent Vaginal Membranes

- Several variations
- How do they cause disease
 - Allow for urine pooling resulting in irritation
 - Ascending bacterial infection
- Diagnosis
 - Digital vaginal exam
 - Cystoscopy
- Treatment
 - Break the band of tissue
 - Often can do with your finger





Recessed/Hooded Vulvas

Transurethral cystoscopy in dogs with recurrent urinary tract infections: Retrospective study (2011-2018)

Marie Llido¹ | Catherine Vachon¹ | Melanie Dickinson² | Guy Beauchamp¹ | Marilyn Dunn¹

- JVIM 2019
 - Observed a high prevalence of hooded vulva (69%)
 - Episioplasty was performed in only 21%
- Severely affected dogs more likely to be clinically relevant
 - Episioplasty reduces UTIs in 84% to 100% of dogs
 - Obesity can be a big factor in conformation
- How does this cause disease?
 - Urine pooling, perivulvar dermatitis, fecal contamination
 - Ascending bacterial infection

Received: 13 September 2019 Accepted: 30 January 2020 DOI: 10.1111/jvim.15728 Journal of Veterinary Internal Medicine

STANDARD ARTICLE

Recessed Vulvas

- Or does it?
- 14% of dogs, most common in spayed females and overweight dogs
- Did NOT find a higher rate of UTIs

Characterization of recessed vulvas in dogs

OBJECTIVE

To determine the prevalence of vulvar recession in a large population of dogs and to compare the reproductive and physical differences between dogs with and without recessed vulvas.

ANIMALS

250 female dogs presenting to a tertiary referral institution.

PROCEDURES

Female dogs > 6 months of age presenting to a tertiary referral institution were enrolled. At enrollment, a full medical history was obtained with particular emphasis on the presence of lower urinary tract (LUT) disease in the 3 months prior to presentation. All dogs underwent a full physical examination including perivulvar cytologic examination and scoring of the degree of perivulvar skin coverage on the basis of an 8-point scale. Dogs with scores of \geq 7 were classified as having recessed vulvas. When available, urinalysis data were also included.

RESULTS

Recessed vulvas were identified in 36 of 250 (14%) dogs. Dogs with recessed vulvas had significantly higher body condition scores and body weights than unaffected dogs. In addition, recessed vulvas were more common in spayed than sexually intact dogs. Dogs spayed at ≤ 1 year of age were almost 3 times as likely to have vulvar recession, compared with dogs spayed at > 1 year of age. No significant difference was identified between affected and unaffected dogs with respect to the prevalence of LUT signs, urinary tract infections, or perivulvar dermatitis.

CONCLUSIONS AND CLINICAL RELEVANCE

Although recessed vulvas were relatively common in dogs, they did not appear to be associated with an increased risk of LUT disease or perivulvar dermatitis. (J Am Vet Med Assoc 2021;259:744–748)

Recessed or Hooded Vulvas

- Bottom line
 - I do think this is a factor in some patients with recurrent UTIs
 - UTIs that we cannot solve by other means
 - Full investigation for other underlying causes
 - PU/PD disorders
 - Endocrinopathies
 - Dermatological disease
 - Obesity
 - Weight loss prior to procedure important
 - Need to know what your doing!
 - May need to be repeated?



The Prostate Gland

- What diseases?
 - Prostatitis/prostatic abscess
 - Intact male dogs
 - Prostatic neoplasia
 - Neutered male dogs
 - Benign prostatic hypertrophy
 - Neutered male dogs
- Clinical Signs
 - Febrile
 - Rectal exam:
 - Large prostate
 - Painful
 - Variable texture

- Laboratory Changes
 - Inflammatory CBC
- Sampling
 - Ultrasound guided FNA
 - Cytology and culture
 - Risks?
 - Prostatic washing
 - Ejaculate
 - Intact male dogs

Prostatitis

- How to treat?
 - Consider the target tissue
 - Blood/prostate barrier
 - Abscess
 - Surgical debridement and omentalization
- When to neuter?
 - AFTER treatment is complete and successful
 - Otherwise, decreased prostatic blood flow will limit deliver of antibiotics

Urolithiasis

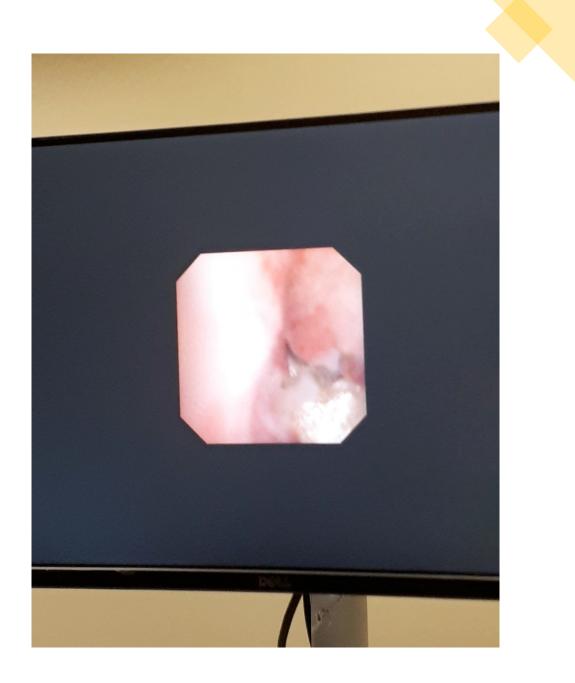
• Types

- Struvite
- Calcium oxalate
- Urate
- Cysteine
- Mixed
- Analysis
 - ALWAYS analyze a stone
 - Even if they have had them before!!
 - Culture the stone
 - Culture bladder wall

- An entire topic to itself!
 - Characteristics of each type
 - Medical vs surgical management
 - Males vs females
 - Diet
 - Dissolution vs maintenance
 - Underlying diseases
 - Hypercalcemia
 - Hyperadrenocorticism
 - Predisposed breeds

Cysteine Urolith

- Young adult (3 or 4 years old) Intact male Rottweiler
- Presenting Complaint:
 - Stranguria, hematuria of 1 month duration
 - Licking at penis
 - Lethargy, decreased appetite
- Physical exam
 - Bloody urine dripping from penis
 - Large, firm bladder on abdominal palpation
- Abdominal radiographs
 - Very large bladder
 - No visible uroliths



Urinary Incontinence

- Urethral Sphincter Mechanism Incontinence (USMI)
 - 3-20% of female dogs
 - Post spay (usually 3-4 years later)
 - Loss of estrogen decreases responsiveness to sympathetic stimulation, resulting in decreased tone
 - More than just this though!
 - GNRH and LH play a role in feedback and loss of smooth muscle contraction
 - Increased collagen in spayed females
 - Should we spay later?
 - Some studies say yes, but methodology issues make it hard to know
 - A more recent study suggests that dogs >25kg may benefit from later spay
 - Not true of small dogs
 - Congenital often present in dogs with ectopic ureters

USMI and UTIs

- How does USMI influence UTIs?
 - Chronic periurethral/perivulvar dermatitis
 - Ascending infection of normal skin organisms
 - Recurrent infections
 - Sometimes different bacteria or mixed infections
 - Repeated therapy can lead to antimicrobial resistance

USMI Diagnosis

- Rule out other causes of incontinence
 - Ectopic ureters
 - Urolithiasis
 - Bladder/urethral masses
 - Pelvic bladder positioning
 - PU/PD disorders
 - Older patients often develop mild USMI
 - If they become PU/PD, their bladder is always full, stressing the sphincter tone
 - Fix the PU/PD and we sometimes fix the incontinence
 - Neurological disease
 - Pudendal nerve damage results in decreased sphincter tone
 - Sacral nerve damage results in lack of sensation to urinate leading to overflow

USMI Treatment

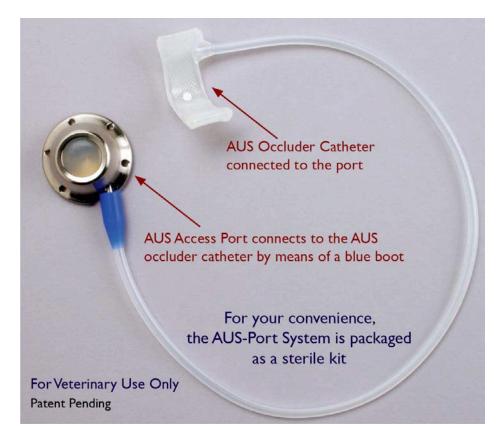
- Pharmacological Therapy
 - Phenylpropenalamine (PPA)
 - Multiple times per day (2-3)
 - Diethylstilbesterol
 - Daily for 5 days, then 1-2 times weekly
- Physical Intervention
 - Collagen injections
 - Good short term effect, likely need to be repeated
 - Hydraulic Occluder
 - Artificial urethral sphincter
 - I have had good success with my patients





USMI in Male Dogs

- Less frequent than in female dogs
- Medical Treatment less rewarding
 - Only 44% of male dogs respond to PPA
 - Only 38% respond to testosterone to become continent
 - 50% no response at all
 - Side effects: aggression and prostatomegaly
- My treatment recommendation:
 - Surgical placement of hydraulic occluder



Pelvic Bladder

- Some dogs with urinary incontinence have a bladder that is caudally located in the pelvis
- Unclear exactly why or how this related to incontinence
 - At least 50% of dogs with pelvic bladder are completely continent
- Often have a misshapen bladder
 - Fails to taper into the urethra
- Part of a bigger syndrome?
 - Shortened urethra
 - Dysfunctional detrusor muscle
 - Urethra that overrides the bladder?



PU/PD Disorders and UTIs

- Why does PU/PD lead to UTIs?
 - Dilute urine?
 - Lack of evidence that this and the resulting decreased osmolality predisposes to UTI
 - Glucosuria
 - Decreased neutrophil chemotaxis
 - Enhanced bacterial growth
 - Hypercalcemia
 - Increased mineral content and stone formation
- Clinical signs
 - Polyuria can hide pollakiuria
 - Dilute urine can make identification of hematuria and pyuria challenging

Diabetes Mellitus

- Most common reason for a well controlled diabetic to become poorly controlled
 - Major cause of DKA in dogs and cats
- Why do they get UTIs?
 - Enhanced bacterial growth due to the glucosuria
 - Dilute urine lower urine osmolality
 - Decreased neutrophil chemotaxis less pyuria
 - Due to glucosuria
 - Hard for owners to identify clinical signs PU/PD

Retrospective Evaluation of Urinary Tract Infection in 42 Dogs with Hyperadrenocorticism or Diabetes Mellitus or Both

S. Dru Forrester, Gregory C. Troy, M. Nell Dalton, Jennifer W. Huffman, and Golde Holtzman

- 42/101 dogs 41.6% had positive cultures
 - 21/46 (46%) with HAC
 - 18/49 (37%) with DM
 - 3/6 (50%) with both

Table 1.	Clinical signs in 101 dogs with hyperadrenocor-
ticism or	diabetes mellitus or both and results of quantita-
tive urine	cultures.

	Positive Culture (n = 42)	Negative Culture (n = 59)	P Value	Power
Polyuria/polydipsia	28 (67%)	45 (76%)	0.37	0.17
Stranguria	2 (4.76%)	0 (0%)	0.17	0.33
Pollakiuria	2 (4.76%)	1 (1.7%)	0.57	0.14
Discolored urine	2 (4.76%)	1 (1.7%)	0.57	0.14

Table 2. Urinalysis findings in 101 dogs with hyperadrenocorticism or diabetes mellitus or both and results of quantitative urine cultures.

	Positive Culture $(n = 42)$	Negative Culture $(n = 59)$	P Value
Specific gravity ^a	1.020 ± 0.013	1.022 ± 0.014	0.60
pH	6.13 ± 0.90	6.28 ± 0.98	0.52
Proteinuria	24 (57%)	38 (64%)	0.54
Occult blood	24 (57%)	20 (34%)	0.03
Hematuria	21 (50%)	20 (34%)	0.15
Pyuria	25 (60%)	9 (15%)	0.001
Bacteriuria	29 (69%)	4 (7%)	0.001

Immunosuppressed Conditions

- Primary
 - Impaired neutrophil chemotaxis may allow for bacterial colonization
 - Neutropenia
 - Inability to fight infection
- Secondary
 - Immunosuppressive therapy for a primary immune-mediated disease
 - Some evidence it may increase the risk of subclinical bacteriuria as opposed to true UTI
 - Do steroids or excess cortisol limit the ability of the body to express cystitis?
 - Chemotherapy
 - Is the patient suddenly ill?
 - Cyclophosphamide

Urinary Neoplasia

- Types
 - TCC/urothelial carcinoma (UC)
 - Prostatic carcinoma
 - Leiomyoma
- Locations
 - Bladder
 - TCC/UC often in the neck or trigone region (>50%)
 - Urethra
 - Prostate
 - Vaginal



Urinary Neoplasia: Diagnostic Testing

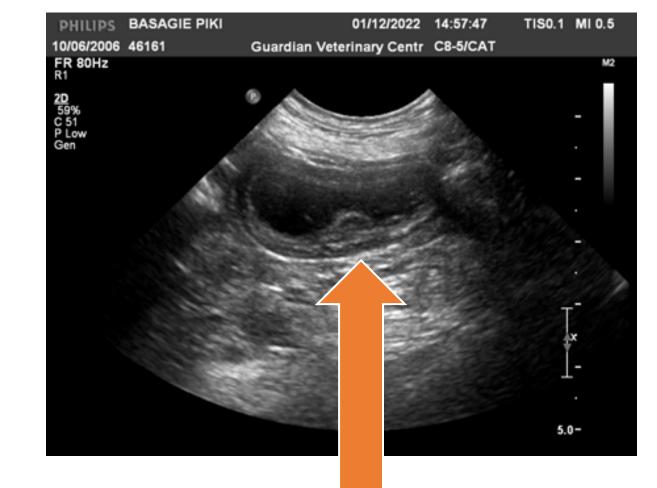
- Abdominal Ultrasound
- Traumatic Catheterization
- Cystoscopy + biopsy
- BRAF testing



The power of molecular diagnostics

CADET® BRAF evaluates urine samples from dogs for the presence of cells containing a mutation for canine bladder/prostate cancer (TCC/UC). It's cutting-edge technology that is accurate, affordable, and convenient for both veterinarian and pet owner.

Download the Test Request Form



Abdominal Ultrasound

- Excellent tool to identify masses with the bladder itself
 - Apex vs trigone
- Allows assessment of complicating factors
 - Ureteral obstruction and/or dilation
 - Regional lymph nodes
 - Prostate in male dogs
- Sampling?
 - FNA can seed the abdomen/body wall resulting in spread and rapid progression of disease
 - Guide catheter for traumatic catheterization



Traumatic Catheterization

- Materials:
 - Rigid polypropylene catheters
- Purpose:
 - Dislodge or break off small pieces of a mass lesion within the bladder/urethra
 - Differentiate between neoplastic and inflammatory disease
- Utility:
 - Easily performed under sedation
 - Often collects diagnostic samples
 - But not always
 - Can be guided by ultrasound or via rectal exam

Liquid Biopsy – CADET[®] BRAF testing

- There is a single mutation in exon 15 of the BRAF gene results in a change of an amino acid (valine to glutamic acid)
- 85% sensitivity and 99% specificity
 - 15% of TCC/UC do not have the BRAF mutation
 - 2/3 of these can be detected with BRAF Plus
 - False positive results have not been identified to date
- Sample:
 - 10mL of FREE CATCH sample
 - Specific collection container
 - CANNOT be refrigerated

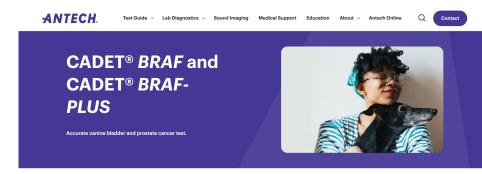
CADET BRAF and BRAF-Plus

Achieve accurate and early diagnosis of TCC/UC

CADET[®] *BRAF* is a highly sensitive test designed to monitor the b-raf mutation in TCC/UC cases during the course of their treatment, for therapeutic response and relapse. CADET[®] *BRAF* testing can be used for both the rapid, non-invasive assessment of dogs displaying clinical signs consistent with TCC/UC and for confirmed cases undergoing treatment.

How CADET[®] BRAF is used in a clinical setting

CADET[®] *BRAF* evaluates free-catch urine samples from dogs for the presence of cells harboring the BRAF mutation or specific copy number variations associated with TCC/UC. The assays identify 95% of TCC/UC cases. The extremely low limit of detection of 10 mutation-bearing cells in a urine sample allows early diagnosis of a developing TCC/UC, often several months before any advanced clinical signs associated with the cancer become evident.



The power of molecular diagnostics

CADET® BRAF evaluates urine samples from dogs for the presence of cells containing a mutation for canine bladder/prostate cancer (TCC/UC). It's cutting-edge technology that is accurate, affordable, and convenient for both veterinarian and pet owner.

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Clinical indications for when to use CADET® BRAF and BRAF-PLUS

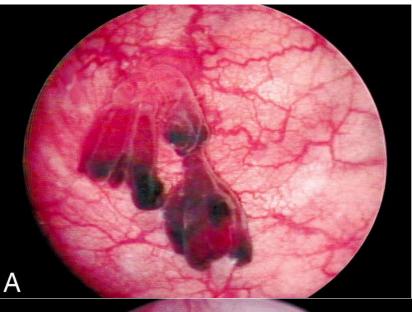
- Clinical cases presenting with hematuria, stranguria, and/or urinary incontinence with diagnostic imaging evidence of a mass in the bladder.
- Confirmation of the TCC/UC diagnosis of a bladder mass from a stained cytology slide following ultrasonography and cytological examination of a fine-needle aspirate from tumorbearing cells.
- Early diagnosis in clinical cases with recurrent, complicated, or antibiotic-resistant urinary tract infections presenting with hematuria without ultrasonographic evidence of a bladder mass.



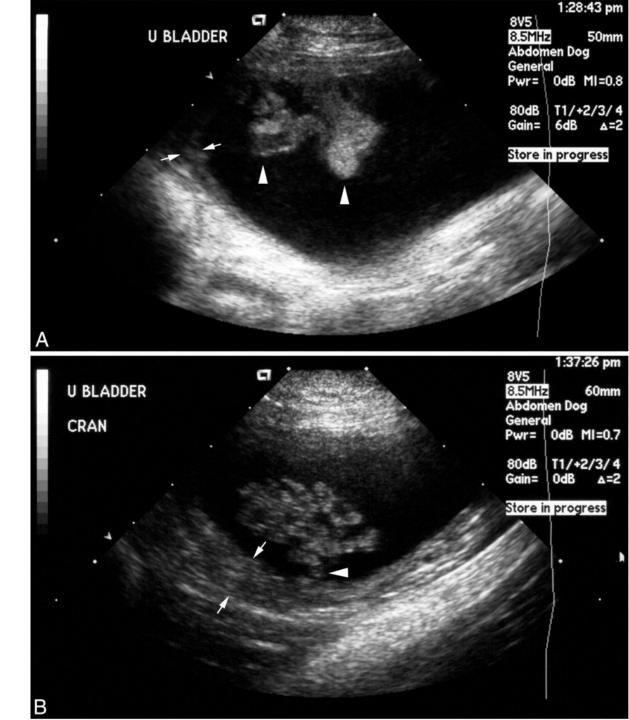
Urinary Neoplasia: Treatment

- Treatment of concurrent infection
 - Ideal drug?
 - Duration?
- Treatment of primary neoplasm
 - NSAIDs (piroxicam)
 - Chemotherapy
 - Radiation therapy
- Surgical excision
 - Maybe for TCC in the apex
 - Vaginal masses

- Non-neoplastic inflammatory condition
 - From one to many mass like lesions
 - Hyperplastic transitional cells and variable amounts of inflammation
- Lesions may be ulcerated, hemorrhagic, hard to diff. from TCC
 - Often more cranioventral as opposed to TCC (neck/trigone)
- Cytologically can be similar to TCC
 - Nuclear criteria for malignancy should not be common, but...
 - Biopsy and histopathology needed to confirm diagnosis
 - Full thickeness Sx biopsies consistently better than endoscopic bx



Polypoid Cystitis in 17 Dogs (1978-2001) J Vet Intern Med 2003;17:499-509



В

- Why does this happen?
 - No definitive answer
 - Chronic inflammation the most likely answer
 - Signs range from 2 weeks to 3 years before diagnosis in some dogs
 - People chronic catheterization
- Clinical Signs:
 - Hematuria
 - Often without UTI
 - Persistent or Recurrent UTIs
 - Some dogs asymptomatic (esp males)
 - Females >> males (88% vs 12%)

- What to do about it?
 - Treat concurrent UTI
 - This is a situation where longer duration therapy becomes appropriate
 - Abnormal mucosa makes it harder to clear infection
 - Culture bladder wall
 - Some studies find up to 18% of dogs with negative urine culture can have + mucosal culture
 - Manage concurrent urolithiasis
 - Surgery to excise the affected tissue is the best solution
 - Over ½ of dogs were normal after surgery
 - Some dogs have recurrent signs and need multiple surgeries
 - More common if the polyp debulked rather than partial cystectomy

Proliferative Urethritis

- Inflammatory, infiltrative disease of the urethra
 - Typically lymphoplasmacytic in nature
 - Causes partial or complete urethral obstruction
 - Previously called granulomatous urethritis
- Unknown cause
 - 73% of dogs have a history of UTIs
 - >50% of patients have + FisH for adhesive/invasive bacteria
 - Possibly immune-mediated reaction to infection
- Female dogs only

Proliferative Urethritis

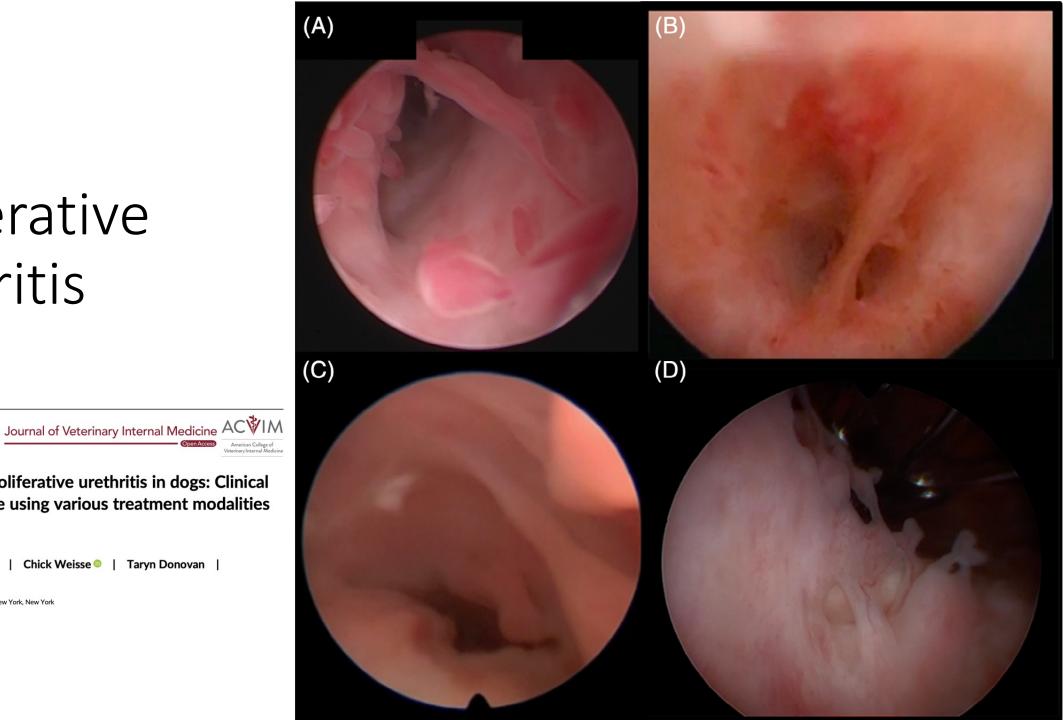
Received: 16 June 2020 Accepted: 2 December 2020 DOI: 10.1111/jvim.16007

STANDARD ARTICLE

Retrospective study of proliferative urethritis in dogs: Clinical presentation and outcome using various treatment modalities in 11 dogs

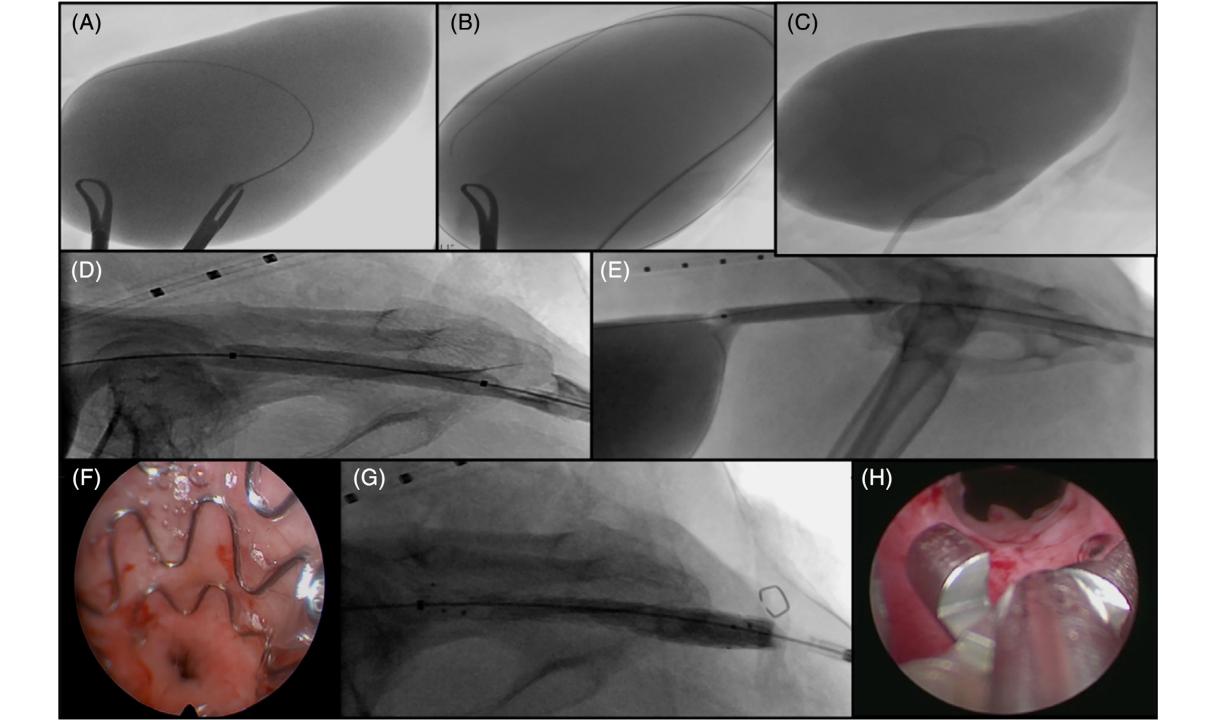
Max Emanuel 💿 Allyson C. Berent | Chick Weisse <a>[] | Taryn Donovan | Kenneth E. Lamb

Department of Interventional Radiology, Animal Medical Center, New York, New York



Proliferative Urethritis

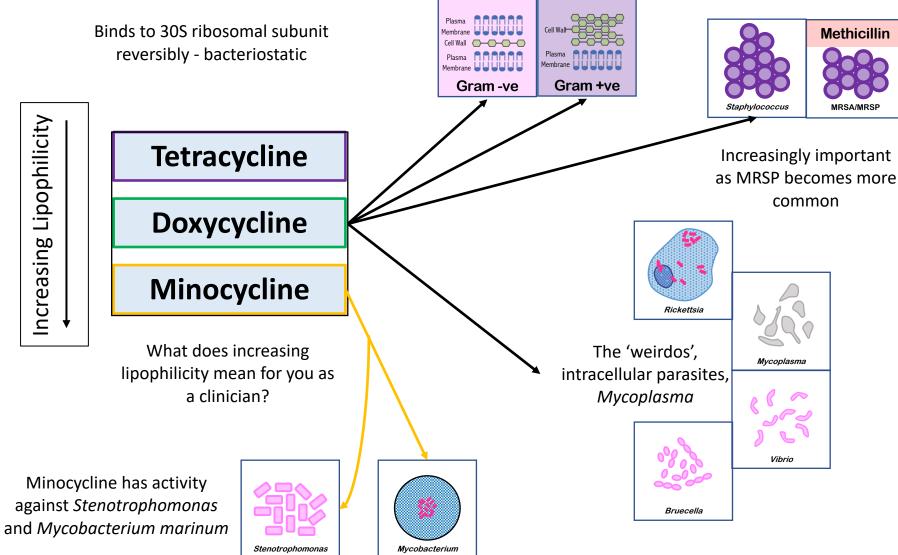
- Treatment
 - Balloon dilation and/or placement of urethral stent
 - Without this, recurrent obstruction in ~3months
 - Median remission time 687 days (range 196-1738d) and Stent placement does seem to offer better outcome
 - Complications: 67% incontinent after stent placement
 - Immunosuppressive therapy
 - Azathioprine, often in combination with other drug (i.e. pred)



More About Drugs

Tetracyclines

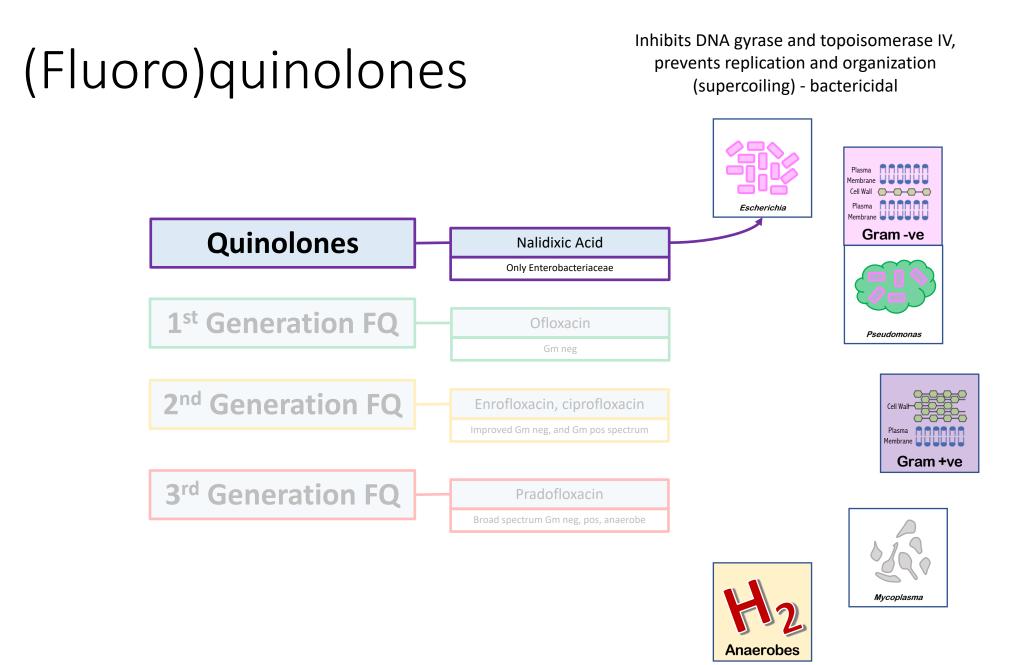
Broad spectrum agents. Gram positive activity more limited than Gram negative Resistance is common, so susceptibility testing essential

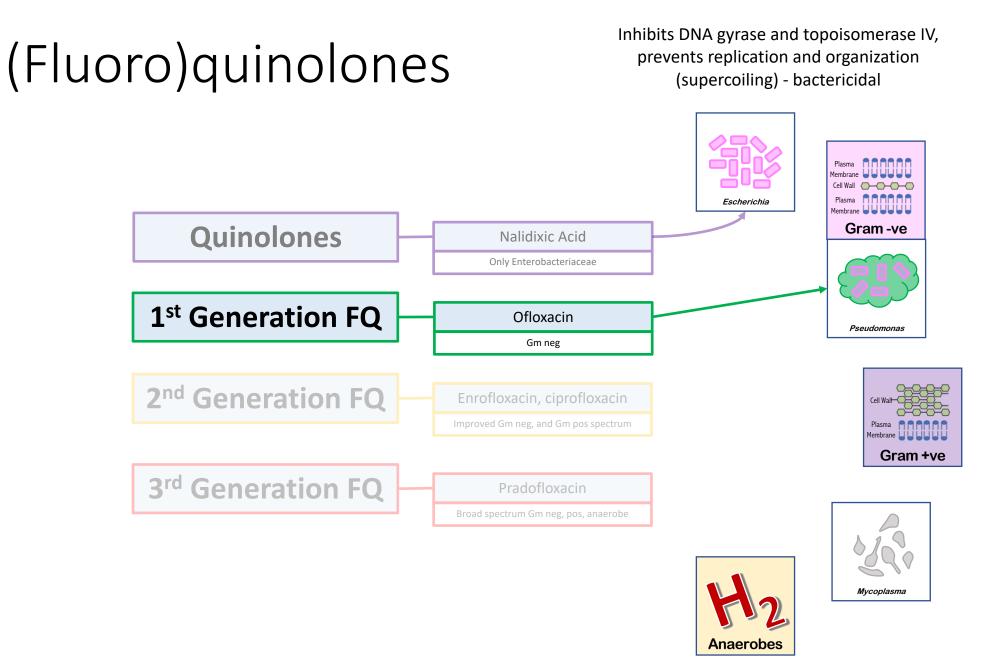


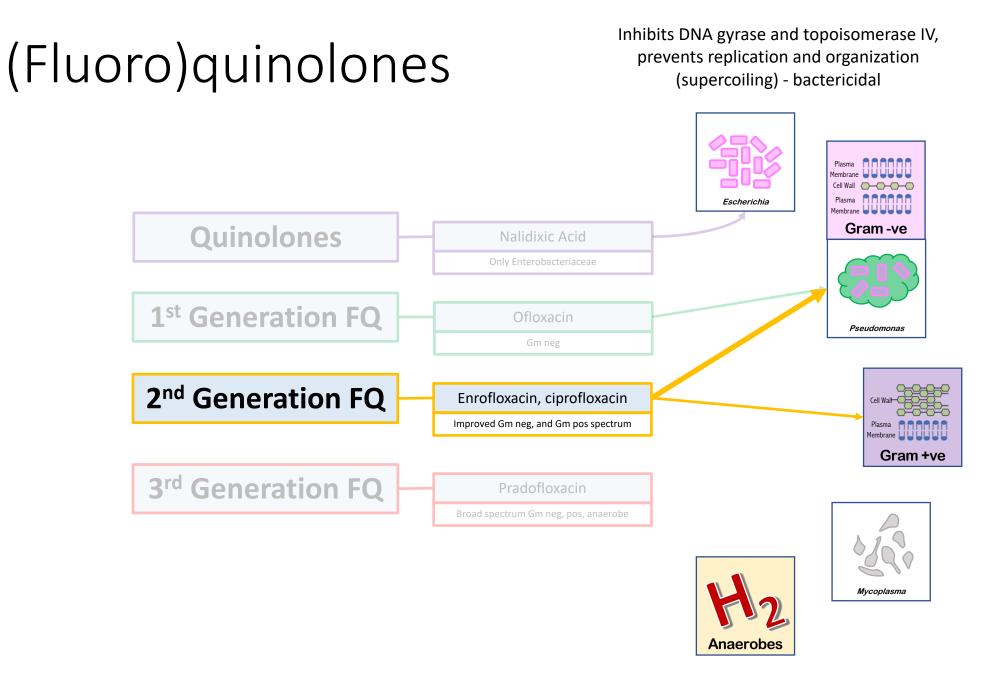
Mechanisms of Tetracycline Resistance

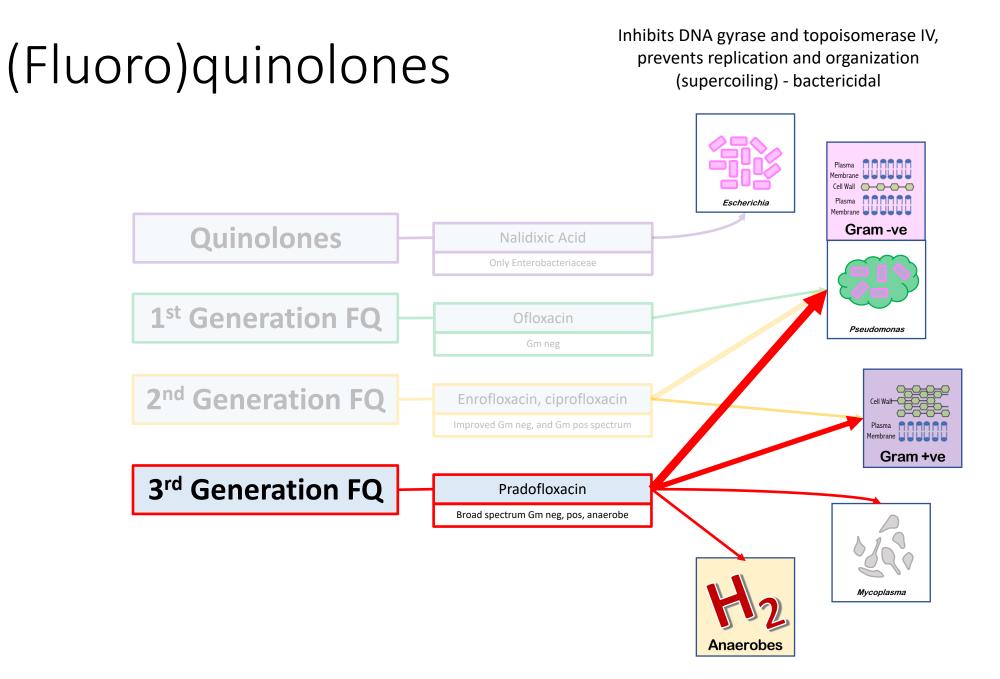
• Efflux

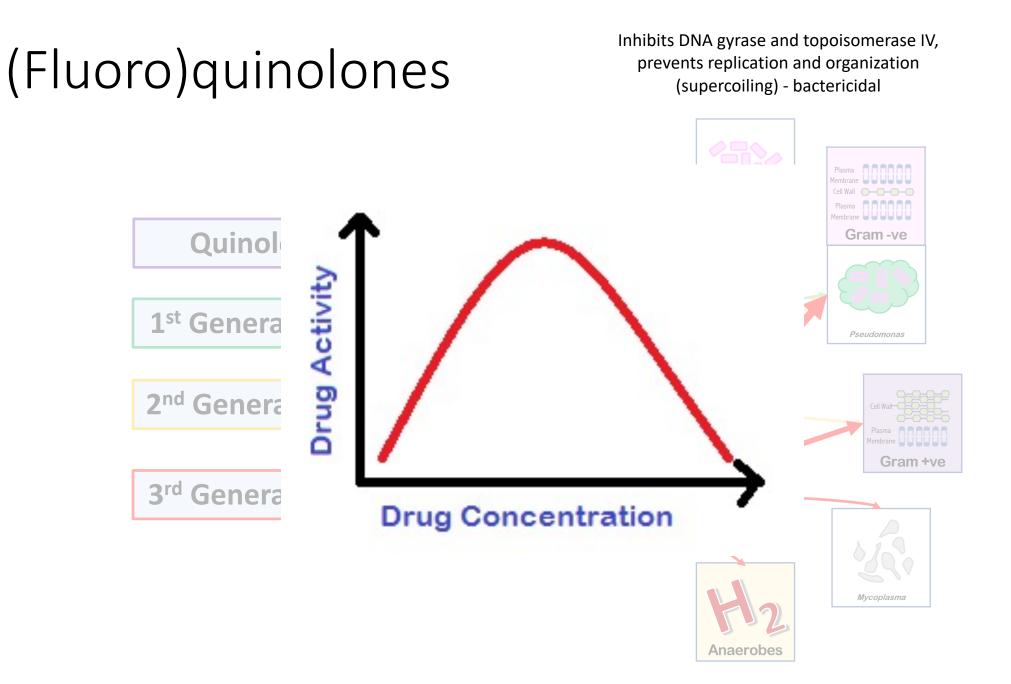
- Common in Gram positive and negative
 - Resistance not necessarily across class...
 - If you want to use a drug test it!
- Ribosomal protection
 - Very common
 - S. pseudintermedius (tetM)
 - Conformational change in tetracycline binding site on 30S subunit of ribosome
- Ribosomal mutations, enzymatic inactivation also occur



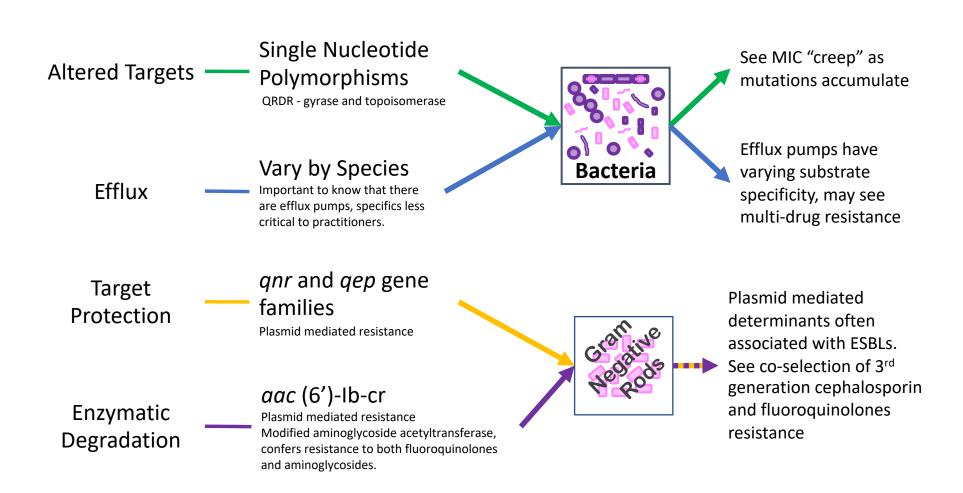


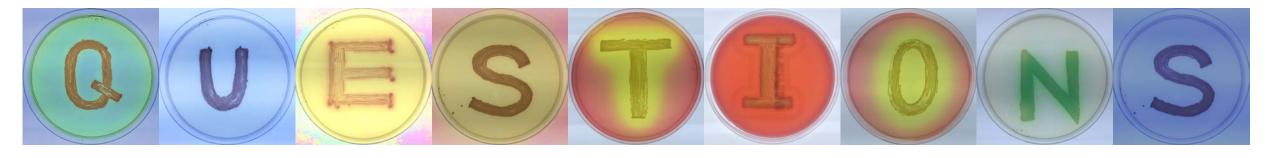






Mechanisms of Resistance









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