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1/¶Hello #medtwitter

This month's (Aug) @ASPNeph Radiology webinar was all about vesicoureteral reflux (VUR)
Here are some clinical pearls!

What's new!?

#Medtweetorial #nephtwitter #ASPNeph

#vesicoureteralreflux #vur

VUR - Against the flow!



2/ Let's get started with a poll!Which one of these is true about VUR?

3/ Ans - All the above

PMID: 32022517

Let's quickly define vesicoureteral reflux
Retrograde flow of urine from the bladder into the upper
urinary tract due to dysfunctional vesicoureteric junction
(VUJ)

- 4/ Some interesting history facts:
- Galen & Winci: UVJ is a mediator of unidirectional flow of urine from kidneys to bladder
- Hutch(1952): Relationship between VUR & Description
 chronic pyelonephritis in paraplegic pts
- ◆ Hodson(1959): UTI & tenal scarring → high likelihood of VUR



Claudius Galen, from Prints Collection, Reynolds Historical Library, the University of Alabama at Birmingham

5/ What are the causes of Vesicoureteral reflux?

- Primary VUR
 - Lack of VUJ maturation
 - Most common
- Secondary VUR
 - Organic obstruction or neurologic dysfunction

PRIMARY VESICOURETERAL REFLUX	SECONDARY VESICOURETERAL REFLUX
Lack of maturation of VUJ => Incompetent or inadequate closure => reflux	Defect in preventive reflux mechanism due organic obstruction and/or neurological dysfunction
Most common reflux	
Congenital anomalies of VUJ - Short intramural/intravesical ureter	Bladder outlet obstruction (anatomic/functional) High pressure within the bladder → reflux of urine through the UVJ Anatomic: - Posterior urethral valves - Urethral meatal stenosis Functional: - Bladder and bowel dysfunction - Neurogenic bladder - Cystitis
	Congenital anomalies of ureter: - Ureteral duplication (most common renal abnormality) - Ectopic ureter - Ureterocele latrogenic
	Ref: Rensing, Adam and Austin, Paul, ,"The diagnosis and treatment of vesicoureteral reflux: An update." Open Urology and Nephrology Journal.8, Suppl 3: M3. (2015).

6/ VUR can be induced by multiple birth defects affecting urinary tract development:

- Ureteric budding
- Ureter differentiation & Deristalsis
- ◆ UVJ formation
- ◆ Bladder & amp; urethra development

*genes implicated:

RET

PAX₂

ROBO₂

KAL₁

HOXA₁₃

NIPBL

7/ VUR

- Usually asymptomatic
- often presents with nonspecific signs & Damp; symptoms like failure to thrive, fever when UTI, lethargy, abdominal pain, voiding symptoms etc, especially with high grade VUR
- 8/ What are the Clinical consequences of VUR?

 It was previously thought that VUR → Recurrent UTIs

 Renal scarring → CKD
- ★Now it is increasingly recognized that children are born with dysplastic kidneys to begin with, which is associated with reflux-VUR, recurrent UTIs and CKD

9/Poll

Which of these is the definitive method of diagnosing VUR and defining its severity?

10/ Ans - VCUG/MCUG

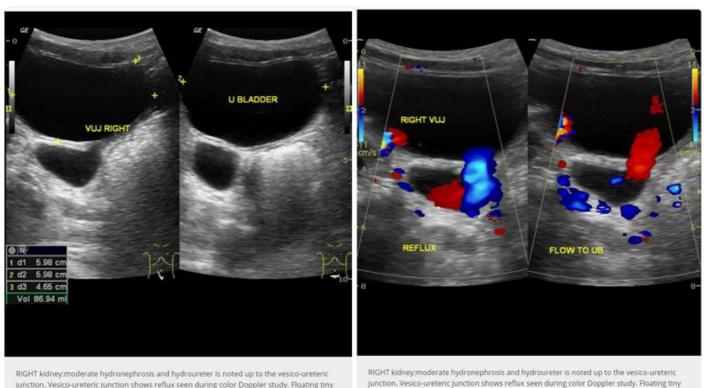
Let's look at how to proceed with imaging in a child with febrile UTI

Renal Bladder Ultrasound-RBUS

Initial imaging choice for fUTI & amp; as a follow up scan to monitor disease progression

Detects hydronephrosis & Detects hydronephrosis & Detects hydronephrosis amp; dilated ureter

Normal USG doesn't rule out VUR



junction. Vesico-ureteric junction shows reflux seen during color Doppler study. Floating tiny echoes are noted in urinary bladder

junction. Vesico-ureteric junction shows reflux seen during color Doppler study. Floating tiny echoes are noted in urinary bladder

11A/ VCUG/MCUG Voiding/Micturating cystourethrogram

Indications

♦ <6m

Fever ≥ 39°C & amp; atypical UTI

Renal anomaly on RBUS

♦6m-3y

Recurrent UTIs

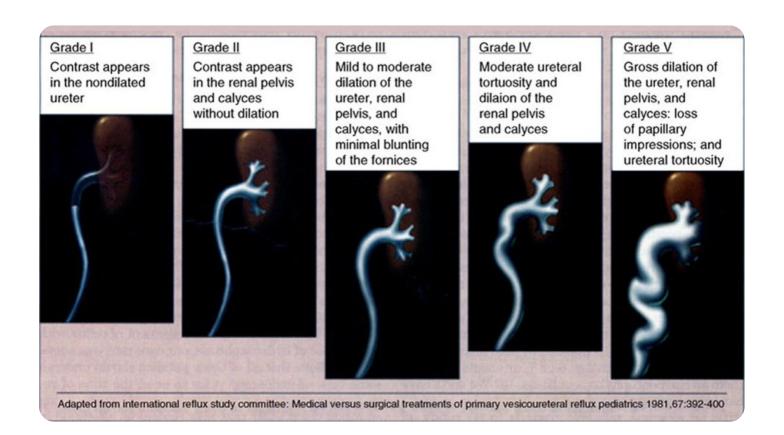
Male infants with prenatally diagnosed b/l hydronephrosis

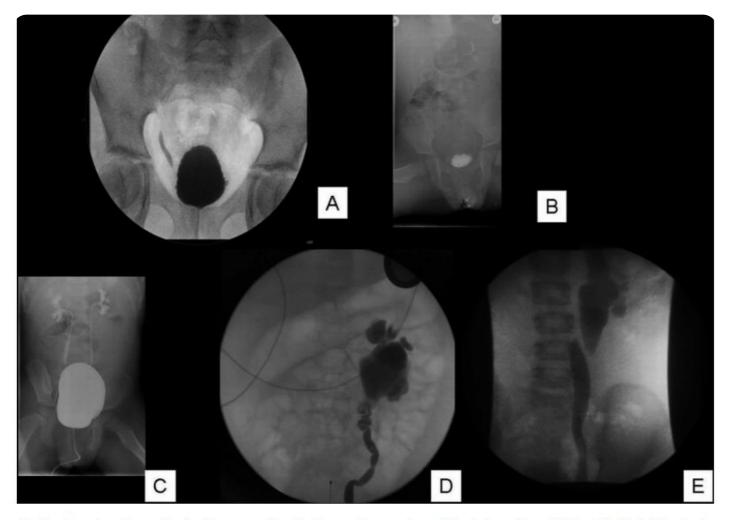
♦ >3y

Usually not indicated unless there is strong suspicion or Hypertension

11B/ VCUG findings and grading

- Findings: Retrograde reflux of the contrast into the ureters during micturition is diagnostic of VUR.
- → Grading: Divided in to 5 grades based on the severity
 of VUR on VCUG





14. Grades of vesicoureteral reflux according to the grading system of the International Reflux Study. (a) Grade 1 ureter only (b), Grade 2 into undilated pelvis (c), Grade 3 into mildly dilated pelvis, (d) Grade 4 into dilated calyces, and (e) Grade 5 into very dilated pelvis and blunting of all calyces.

12/ DMSA Renal Scan

- Nuclear imaging method using radioactive DMSA (dimercapto succinic acid)
- Assess cortical tissue, renal function and scarring
- Indicated for treatment monitoring and as a follow up scan to look for worsening renal function

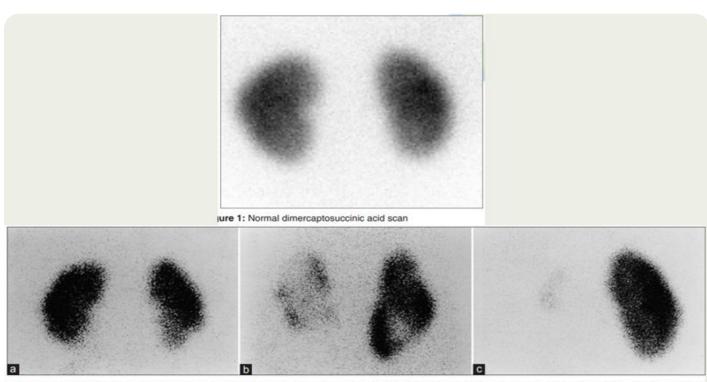


Figure 2: Examples of the renal scarring scoring system. (a) Normal left kidney, grade 1 scarring (lower pole of the right kidney). (b) Grade 2 (right kidney) and grade 3 (left kidney) scarring. (c) Grade 4 scarring (left kidney). Image adopted from Howard et al.[18]

13/ MAG3 Renal Scintigraphy (radionuclear cystourethrography)

- Nuclear medicine scan using MAG3 (mercaptoacetyltriglycine)
- Detects UVJ and UPJ obstruction
- Measures differential renal function

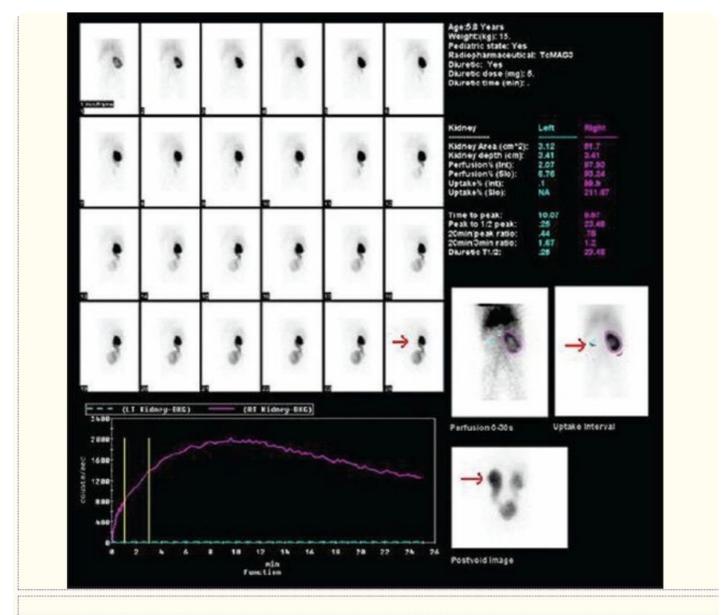


Figure 3

^{99m}Tc MAG3 renal scintigraphy showing fair functioning hydronephrotic right kidney with partial outflow clearance whereas left kidney is not visualized in dynamic images. Post-void image showing appearance of left kidney and ureter indicative of vesicoureteral reflux

14/ Treatment options 💊 📋

Over the last few years Individualized, risk-based approach to the management of VUR with an overall shift to more conservative management has been noticed

TREATMENT OPTIONS FOR VESICOURETERAL REFLUX

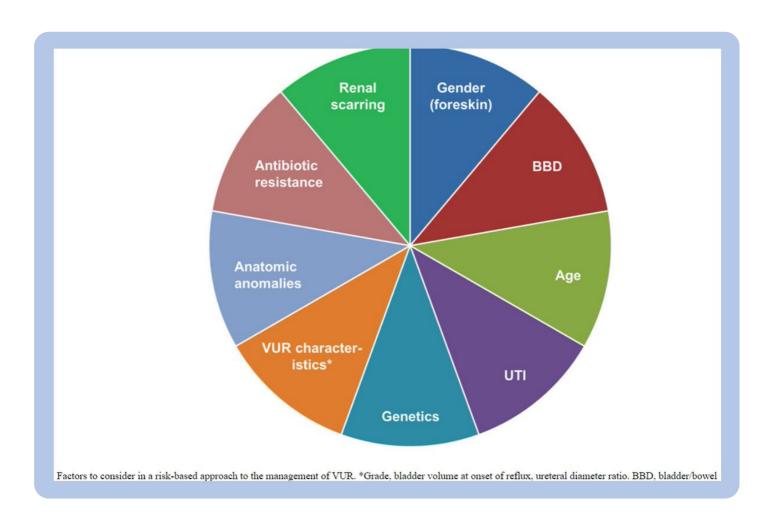
Conservative Treatment

- For VUR Grade I—III in children < 5 years</p>
- Close surveillance with urinalysis & renal imaging
- ➤ Long-term Antibiotic prophylaxis
 - Children > 3m: Trimethoprim
 - Infants < 3m: amoxicillir
- Correction of voiding dysfunction
 - Behaviour modification (timed voiding, Prevention of constipation)

Surgical Treatment

- Indications
 - VUR ≥ grade IV
 - Bilateral grade III VUR in children > 6 years
 - Worsening renal function/breakthrough urinary tract infections despite prophylactic antibiotics
- Subureteric Transurethral Injection (STING procedure)
- > Surgery: Ureteral Reimplantation

Läckgren G, Cooper CS, Neveus T, Kirsch AJ. Management of Vesicoureteral Reflux: What Have We Learned Over the Last 20 Years?. Front Pediatr. 2021;9:650326. Published 2021 Mar 31. doi:10.3389/fped.2021.650326



15/ Here is a table summarizing @AmerAcadPeds and NICE guidelines for the diagnosis and management of UTI in children

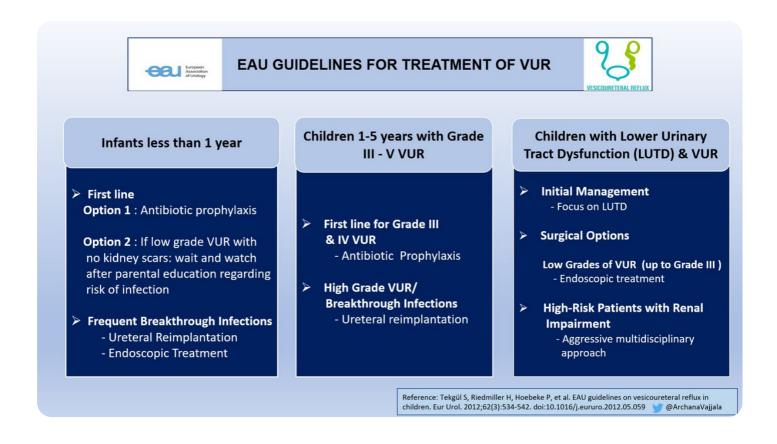


https://www.emjreviews.com/microbiologyinfectious-diseases/article/urinary-tractinfection-in-children-a-review-of-theestablished-practice-guidelines/

Parameters	AAP guidelines (2011)	NICE guidelines (2007)
Diagnosis of UTI		
- Using dipstick urinalysis		- Positive leucocyte esterase or nitrite test*
- Using microscopic urinalysis and urine culture	- Bacteriuria with or without pyuria - ≥5x10 ⁴ CFU/mL (from SPA and catheter urine specimens)	 Colony count of any Gram negative bacilli or >10³ CFU/mL of a Gram- positive coccus (from SPA urine specimen)
		- >10s CFU/mL (from catheter urine specimen)
		- ≥10 ^s CFU/mL (from 'clean-catch' or midstream urine specimen)
Radiological investigation of UTI+		
- RBUS	- Recommended in febrile infants with first UTI	- Recommended in atypical or recurrent UTI in children aged <6 months
- DMSA renal scan	- Not recommended as routine investigation for first febrile UTI	- Recommended in children aged 6 months to 3 years with atypical or recurrent UTI1
- VCUG (or MCUG)	- Not recommended as routine investigation for first febrile UTI	- Not recommended in children aged 6 months to 3 years with atypical or recurrent UTI##
Treatment and follow-up of UTI		
- Route of antibiotics/duration	- Parenteral route for 48 hours (for critically ill patients) and switch to oral route if clinical improvement occurs. 7-10 days as the total duration of therapy	- For children aged <3 months: parenteral route for 2-3 days before a switch to oral route if clinical improvement occurs. 10 days as the total duration of therapy - For children aged >3 months with upper UTI: oral route using antibiotics with low-resistance pattern. 7-14-day duration of therapy§ - For children aged >3 months with lower UTI: oral route for 3 days
- Follow-up routine urine culture	- Not recommended**	- Not recommended
- Follow-up antibiotic prophylaxis	- Not recommended	- Not recommended

16/ Treatment - EAU guidelines

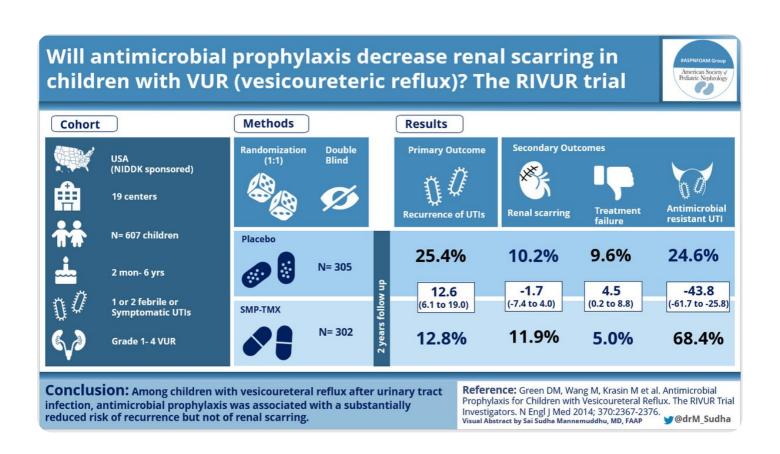
The European Association of Urology guidelines prominently state that "there is no consensus on the optimal management of VUR or on its diagnostic procedures, treatment options, or most effective timing of treatment"



2-year, multisite, RCT
607 children with VUR
Did antibiotic prophylaxis prevent recurrent UTIs
(primary) & Did amp; scarring in kidneys (secondary)?
Conclusion: Prophylaxis prevented recurrent UTI but not scarring

VA by @drM_sudha

https://www.nejm.org/doi/full/10.1056/nejmoa1401811



18A/ @AmerMedicalAssn & amp; @AmerAcadPeds recommendations:

AMA - Recommends annual evaluation of BP, height, weight, urine analysis, and RBUS until the abnormality resolved or is no longer clinically significant

18B/ AAP - All infants with fUTIs undergo renal & Damp; bladder ultrasonography. VCUG is indicated if RBUS reveals hydronephrosis, scarring, high-grade VUR, obstructive uropathy & Damp; in some atypical/complex clinical situations. Not recommended routinely after a first fUTI

PMID:20650499

19/ Prognosis:

Cystic dysplasia and VUR can lead to -

- Recurrent UTIs
- Scarring in the kidneys
- Hypertension
- CKD and eventually ESKD

20/ That's all folks!

For a case-based clinical discussion with radiology expert login to @ASPNeph website, August webinar. Answer questions to get #MOC2credits #Membereducation #ASPNFOAMgroup

21/ Special thanks to @SwastiThinks @drM_sudha @priti899 for your help in publishing my first tweetorial!

Until next time....



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