## Nephrolithiasis cases

### Primary Care Internal Medicine October 2015

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A Teaching Affiliate of Harvard Medical School



 45 year old male, otherwise healthy, acute onset right flank pain + nausea presents to your office for evaluation.

PMH: none

- PE: T 98.9, BP 130/80, Pulse 90
- abdomen soft, non-tender. Mild right CVA tenderness





#### Labs: WBC 12.5, Cr 1.2

## Urinalysis: 2+ RBCs, 2+ WBCs, negative nitrites, negative bacteria

#### Imaging studies: KUB? RUS? CT?







 CT scan shows
 6 mm right upper ureteral stone with hydronephrosis.





MASSACHUSETTS GENERAL HOSPITAL



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# What determines the next steps?





- A 6 mm stone has a 50-60% chance of passing spontaneously
  - Alpha blocker significantly increases passage rate
- My practice:
  - 2-3 weeks alpha blocker (e.g. flomax) if patient meets following criteria:
    - Pain controlled on PO regimen
    - No evidence of infection (fever, nitrites, leukocytosis) WBCs in the urine are okay (reflect inflammation from stone)
  - KUB and renal ultrasound in 2-3 weeks to assess stone passage



FRAL HOSPITAL







 EMERGENCY – if signs of pyelonephritis or systemic infection, this may be a LIFE-THREATENING SITUATION

- Decompression of the urinary system must be done emergently
  - Ureteral stent or nephrostomy tube
  - Choices vary by urologist
  - My practice: stent when possible to avoid external drainage of nephrostomy tube
  - Treat stone after patient has stabilized
    - 2-3 weeks of abx





- Shockwave lithotripsy (SWL)
- Truly minimally invasive shockwaves delivered externally with imaging to confirm
- Sedation
- ~50-60% success rate
- Limited by BMI
- Cannot be done in patients on anticoagulation

- Ureteroscopy (URS)
- Flexible or semirigid endoscope passed into urethra and stones fragmented w/ laser and extracted
- General anesthesia
- 90-95% success rate for stone < 1 cm</li>
- Often requires ureteral stent for ~ 1 week
- No issues w/ BMI
- Safe in patients on anti-coagulation





- Shockwave lithotripsy (SWL)
  - Schematic of an electromagnetic lithotriptor







#### Ureteroscopy (URS)

- Flouroscopic images from laser fragmetation of stone

Flexible ureteroscope in lower pole of kidney

Retrograde pyelogram





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- Distal stones pass more frequently than proximal stones
- Most common place for stone to lodge is distal ureter or ureterovesical junction (~60%)
- Alpha blockers recommended to aid passage for all stones < 1 cm in diameter
- Passage rate is not increased after a single 2 week course of alpha blocker



- MY PRACTICE
- Pain management
  - Acute: IV ketorolac (toradol), IV narcotics
  - For Discharge: diclofenac, percocet or vicodin, colace, alpha blocker
  - Others
    - Antibiotics? if cystitis or infection, should have stent placed; if no infection, no abx
    - Pyridium for dysuria
    - Anti-cholinergics (ditropan) for frequency





- IMAGING
- CT is gold standard
  - Most efficient and accurate
  - Risks = ionizing radiation (but remember, non-contrast CT scan is far lower dose than contrast enhanced)
  - CT more costly than other modalities
- KUB (plain radiography) + ultrasound (RUS)
  - Will pick up a majority of stones
  - Sufficient in many cases (maybe not for obese patients)
  - Less detail than CT
  - Less radiation than CT
  - Less \$\$\$ than CT







## KUB showing right renal stone



#### RUS showing right hydronephrosis and left renal stone





 53 year old female. First stone age 44. She has passed 4 stones (1 every other year) and has had 2 procedures (ureteroscopy) by local urologist. "I need a drug to make these go away!!!!"

PMH: nephrolithaisis (CaOx), DM, HTN, hypothyroid

PE: BMI 33, otherwise unremarkable





Nephrolithiasis – 10% lifetime prevalence worldwide

 Men > women, but female #s are increasing at faster pace

50% recurrence within 10 years of first stone event





#### Stone composition

- Calcium oxalate (most common); CaP (increasing);
  Calcium based stones ~ 80%
- Uric acid (diabetic, overweight) ~ 5-10%
- Cystine (genetic disorder, may present in childhood) ~1%
- Struvite (recurrent infection) ~3%
- Miscelaneous others (protease inhibitors, other rx)



#### Absolute indications for metabolic evaluation

- mutiple/recurrent stones
- solitary kidney
- GU anatomy abnormalities which may make surgery difficulty
  - diversion, horseshoe kidney

#### Relative indications for metabolic evaluation

- young age
- single stone, particularly difficult episode for patient (i.e. sepsis, surgery)
- patient preference



#### Workup

- chem 7, calcium, PTH, uric acid (if PTH is elevated, check vitamin D!!!)
- 24-hour urine
  - 2 collections standard, but many obtain only 1
- Maintenance/monitoring
  - Recent guidelines published by the AUA
  - My practice recheck 24-hour urine 3-4 months after starting a new therapy (e.g. thiazide, allopurinol, k citrate) then yearly x 1-2 years
  - My practice stop checking yearly 24-hour urines once patient's stone disease is stablized





Pharmacotherapy for calcium oxalate stones – the basics

- Hypercalciuria thiazide diuretic (level 1 evidence)
- Hyperoxaluria no standard of care
  - Magnesium oxide, magnesium hydroxide, calcium carbonate with meals

- Hypocitraturia potassium citrate (level 1 evidence)
- Hyperuricosuria allopurinol (level 1 evidence)





Pharmacotherapy for uric acid stones – the basics

- Potassium citrate or sodium bicarbonate
- Can dissolve stones at higher doses
- Lower doses for maintenance
- Usually <u>do not need allopurinol</u> unless potassium citrate fails or if patient has hyperuricemia





 21 year old male, passed 1 stone this year. KUB shows a 5 mm stone in left kidney.

PMH: nephrolithiasis

PE: BMI 30, otherwise unremarkable

"How can I change my diet??? I don't want a pill."





## FLUID INTAKE

- Level 1 evidence randomized trial of fluid intake of 2.5 L/day versus 1.0 L/day
  - Recurrences in 5 years decreased by > 50%
  - Time to recurrence increased from 25 to 38 months





## ANIMAL PROTEIN

 Theoretical risk of increased urine calcium and decreased urine citrate with increased protein consumption and portion size

- Data equivocal but several studies demonstrate association between protein consumption and risk of nephrolithiasis
- My practice "Don't eat anything bigger than your hand (palm) in a single serving."





## SODIUM

 Sodium and calcium co-transported in renal proximal tubule; increased dietary sodium → increased urine sodium → increased urine calcium

- Data equivocal but several studies demonstrate association between dietary sodium and risk of nephrolithaisis
- My practice "Moderate sodium intake."





## OXALATE

- Area of major patient concern
  - Patients can control this part of the diet

 Green vegetables, spinach, potatoes, cereal, oranges, carrots, tea, coffee, nuts, beans, chocolate, strawberries, vitamin C supplements

Urine oxalate = ~30% from diet, 70% from endogenous production (liver)

Taylor and Curhan (2007)





## OXALATE (continued)

- Epidemiologic studies demonstrate that dietary oxalate
  - Is not related to kidney stone risk in younger women
  - Is "modestly" related to kidney stone risk in older women
- The only foods with enough oxalate to change urine oxalate are
   Spinach, beets, rhubarb, vitamin C (> 1000 mg/day)



## PHYTATE

 Common dietary sources include cold cereals, bread, beans

 Epidemiologic evidence demonstrates that phytate is protective against incident stone disease in women





## LEMON JUICE/LEMONADE

- Citrate is potent inhibitor of stone diseae
- Potassium-citrate shown to prevent stone recurrence
- Lemon juice and homemade lemonade preparations have been shown in some retrospective studies to significantly raise urine citrate
- My practice: Mix ½ cup lemon juice (concentrate) w/ 7 ½ cups water and drink daily

Aras et al (2008), Seltzer et al (1996)





## VITAMIN C

 Can be metabolized to oxalate and therefore can theoretically increase risk of calcium oxalate stone formation

- Modest increase in stone risk for consumption
- > 1000 mg/day





- My general dietary recommendations:
- > 2 L fluid/day
- Watch salt intake (< 2g), animal protein (no portion larger than palm of your hand)
- Limit spinach/beets/rhubarb to 1 serving/week
- Vitamin C < 1000 mg/day</p>
- ½ cup lemon juice mixed in 7 ½ cups of water, sweeten w/ artificial sweetener as needed



 67 year old female, recent admission to hospital for hip fracture. Subsequent workup reveals osteoporosis w/ osteopenia.

 PMH: osteoporosis, osteopemia, HTN, history of kidney stones (passed 4 stones since age 55)

PE: BMI 24, otherwise unremarkable





## **Knowing most stones** are made of calcium, is it safe to **Supplement calcium? Supplement vitamin D?** Give bisphosphonates?







Calcium plus Vitamin D Supplementation and the Risk of Fractures

- 36,000 women randomized to calcium carbonate + vitamin D versus control
  - Mean f/u 7 years

• 17% increased risk in kidney stone formation in treatment arm

Real #s: <u>2.6%</u> of treatment versus <u>2.3%</u> of placebo







#### <u>Dietary calcium</u> is PROTECTIVE against stone disease (i.e. reduces stone risk)

Binds oxalate in GI tract

#### <u>Supplemental calcium</u> data is equivocal

 Metanalysis of 12 calcium supplementation trials for osteoporosis shows that calcium supplementation likely is protective against stone disease (up to 2000 mg/day)



 Vitamin D supplementation – may increase GI absorption and therefore urinary excretion of calcium

 No evidence of increased urine calcium in stone formers given 50,000 IU ergocalciferol/week x 8 weeks

 No evidence of relationship between serum vitamin D and urine calcium in stone formers





#### <u>Bisphosphonates</u> - decrease fasting calciuria

- Evidence that bisphosphonates decrease kidney stone formation in bed-ridden patients
- Risedronate protective against stone formation
- -? Ibandronate may increase stone formation





My practice/recommendations

- Maximize DIETARY calcium
  - Take with meals
- Calcium supplementation is fine
  - Take with meals
  - Use calcium citrate
- Vitamin D supplementation no issues
- Bisphosphonates no issues
  - ? Ibandronate caveat





- Thank you!!!!!!!!!
- Questions:





