

crystals >>

①  $\text{Ca}^{2+}$  pyrophosphate dihydrate  $\Rightarrow$  - pseudogout  
CPPD - chondrocalcinosis

② Basic  $\text{Ca}^{2+}$  phosphate Hydroxyapatite  $\Rightarrow$  - calcific periarthritis  
BCP - soft tissue calcinosis  
- Acute/chronic inf. arthritis

③ MonoNa urate (MSU)  $\Rightarrow$  - Acute/chronic gout  
- Renal calculi  
- Tophi

Gout

- Deposition of **Monosodium urate monohydrate** in **CT & joints**
- Age > 50
- M > F
- Hyperuricemia: urate plasma > 6.8 mg/dL

\* Urate is a metabolite of **purines** & **ionized** form of **uric acid**

\* Hyperuricemia due to renal under excretion of urate in 90% of cases  
over production in 10%.

"not all hyperuricemia is gout, BUT increases risk"

▷ **Inflammatory response to urate crystals**

↳ Urate crystal in joints **interact** with **undifferentiated** phagocytes  $\rightarrow$  induced TNF- $\alpha$  & signaling pathways & endothelial cells  $\rightarrow$  leads to neutrophil adhesion to endothelium  $\rightarrow$  influx  $\rightarrow$  amplification  $\Rightarrow$  **Neutrophilic synovitis**

↳ Activate NALP3 inflammasome  $\rightarrow$  IL-1 $\beta$   $\rightarrow$  **inf. loop**

↳ Resolution: clearance by differentiated phagocyte

coating the <sup>(+)</sup> crystal with proteins  
↓

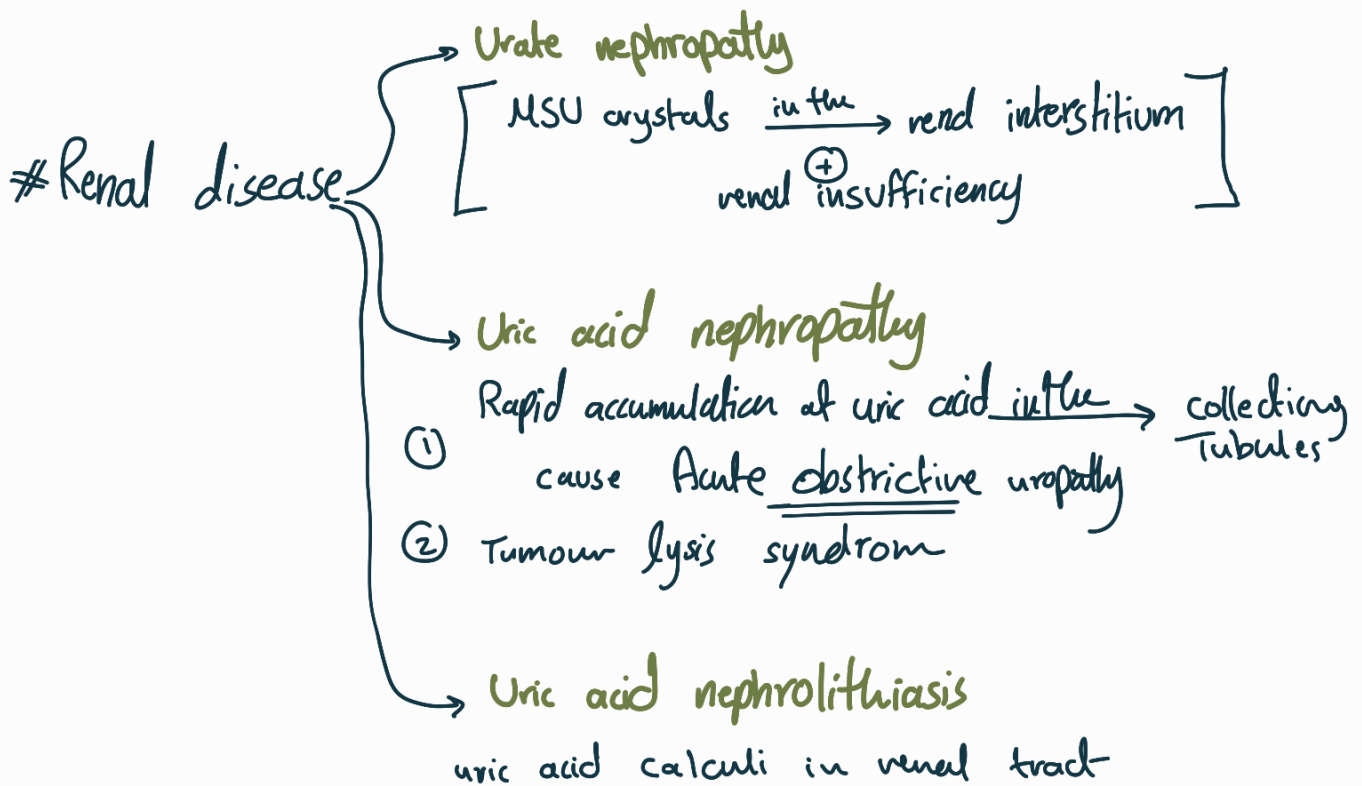
neutrophilic apoptosis &  
(inactivation of inflammatory mediators  
(self limiting within 2 weeks))

# Longstanding hyperuricemia → induce chronic inflammation

- Recurrent attacks of acute  
inflammatory arthritis  
(Big toe)

synovitis cartilage loss Bone erosions  
(By inducing chondrocytes  
to produce metalloproteinase & NO)

- Chronic arthropathy
- Accumulation of urate crystal  
in the form of tophaceous deposit.
- Uric acid nephrolithiasis



## Acute gout :-

- Acute inflammatory arthritis / associated with tenosynovitis
- M x 8 > F (uncommon before menopause)
- Age (30-60)
- Recurrent attacks of acute synovitis with (intercritical periods in between)
- Big toe / 10% polyarticular < elderly women
- Rapid & severe symptoms. < diuretics

\* labs → leucocytosis & ↑ ESR  
↑ uric acid (normal in acute attack)

Triggers ⇒ Disturbances in extracellular fluid urate conc.

① Drugs  
low dose aspirin  
loop diuretics  
Thiazide  
Allopurinol  
uricosuric

- ② fatty food
- ③ surgery / Trauma
- ④ dehydration
- ⑤ starvation

► Tophus : Draining or chalk-like subcutaneous nodule under transparent skin, overlying vascularity

sites:

(Ears / elbow / finger / Tendon) ⇒ peripheral, cold parts

↓  
over long time  
↓

Chronic, erosive arthritis, associated with peri-articular & subcutaneous urate deposit (Tophi)

↳ May cause dactylitis

Diagnosis: crystal identification / serum urate / radiology / synovial fluid + histology  
↑ MSU crystals

\* Serum urate level

- M > F
- falsely low during attack
- Aim: reduce level to  $\leq 6 \text{ mg/dl}$
- risk of gout with
  - degree
  - duration of hyperurcemia

### \* Radiographic

- normal  $\rightarrow$  early in the disease
- punched out erosions with sharp margins and overhanging edges  $\rightarrow$  few repetitive acute attacks

### \* Crystal identification (Anisotropy)

- even asymptomatic inter-critical period
  - $\hookrightarrow$  crystals are present in previous attack joints
  - in all untreated
  - 70% of treated

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\* high dose  $\rightarrow$  phenylbutazone  
 aspirin  $\Rightarrow$  protection that increase excretion

low dose  $\rightarrow$  phenyl  
 aspirin  $\Rightarrow$  decrease excretion

\* thiazide  $\downarrow$  excretion.

Treatment :-  $\rightarrow$  with NSAIDs 0.5 mg x2 / day  
\* NSAIDs / colchicine / steroids

Lindomethacin 50mg / 8hrs  
naproxen 500mg / 12hrs  
Diclofenac 50mg / 8hrs

\* Intra-articular uses  $\left\{ \begin{array}{l} \text{NSAIDs is} \\ \text{contraindicated} \\ \oplus \\ \text{1 or 2 joints are} \\ \text{inflamed} \end{array} \right.$

2<sup>nd</sup> Systemic for polyarticular disease  
 $\left\{ \begin{array}{l} \text{NSAIDs or colchicine contraindicated} \\ \text{chronic kidney D.} \\ \text{peptic ulcer} \\ \text{chronic heart failure} \end{array} \right.$

\* Allo/purinol or urico-suric drugs  $\otimes$  NOT used during attack.

# Avoid  $\left\{ \begin{array}{l} \text{Diuretics} \\ \text{weight gain} \\ \text{alcohol} \\ \text{Aspirin low doses.} \end{array} \right.$