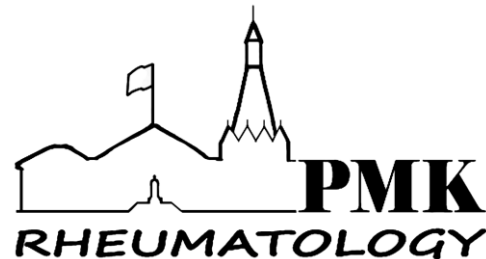


**Review in Internal Medicine 2026**

# **Essentials of Rheumatology**

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Division of Rheumatology  
Phramongkutklo Hospital**



# Case 1

- 45M SLE/LN
- Presented with gout (first attack), no tophi
- CKD (eGFR 25 mL/min/1.73 m<sup>2</sup>)
- Meds: HCQ, azathioprine, diuretic
- SUA 10 mg/dL, HLA-B\*58:01 positive
- X-ray: erosions.
- Renal US: no stone.

**Best management ?**

- A. Allopurinol
- B. Febuxostat
- C. Dotinurad
- D. Benzbromarone
- E. Lifestyle modification and stop diuretic only

# Indications for pharmacologic urate-lowering therapy (ULT)

ACR 2020 <sup>1</sup>	EULAR 2016 <sup>2</sup>	TRA 2026
<p><u>All gout with</u></p> <ul style="list-style-type: none"> <li>- Frequent attacks (≥ 2/year )</li> <li>- Radiographic joint damage</li> <li>- Tophus or tophi                             <ul style="list-style-type: none"> <li>- Recurrent flare</li> <li>- First flare with</li> </ul> </li> </ul> <ul style="list-style-type: none"> <li>• CKD stage ≥ 3</li> <li>• SUA &gt; 9 mg/dl</li> <li>• Urolithiasis</li> </ul>	<p><u>All gout with</u></p> <ul style="list-style-type: none"> <li>- Recurrent flares                             <ul style="list-style-type: none"> <li>- Tophi</li> </ul> </li> <li>- Urate arthropathy                             <ul style="list-style-type: none"> <li>- Renal stones</li> <li>- First flare with</li> </ul> </li> </ul> <ul style="list-style-type: none"> <li>• &lt; 40 year</li> <li>• SUA &gt; 8 mg/dl</li> <li>• Comorbidities: CKD, HT, IHD</li> </ul>	<p><u>All gout with</u></p> <ul style="list-style-type: none"> <li>- Frequent attacks (≥ 2/year )                             <ul style="list-style-type: none"> <li>- Tophi</li> </ul> </li> <li>- Joint damage</li> <li>- First flare with</li> </ul> <ul style="list-style-type: none"> <li>• CKD stage ≥ 3</li> <li>• SUA &gt; 9 mg/dl</li> <li>• Urolithiasis</li> </ul>

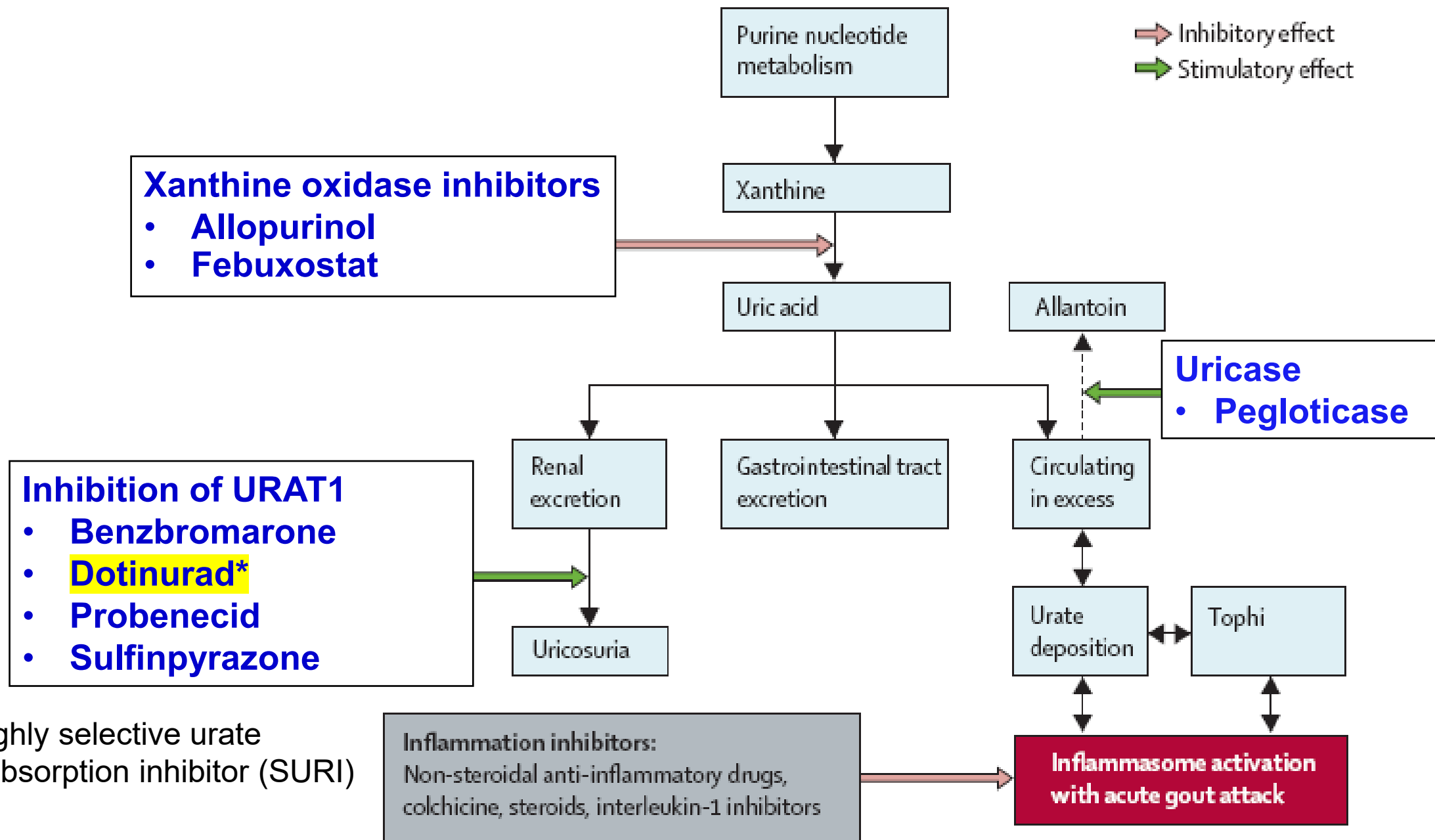
1. John D. FitzGerald, et al. Arthritis Care & Research 2020

2. Richette P, et al. Ann Rheum Dis 2016

# Long term management of Gout: “treat to target strategy”

<b>Guidelines</b>	<b>ACR 2020<sup>1</sup></b>	<b>EULAR 2016<sup>2</sup></b>	<b>TRA 2026</b>
<b>General target</b>	< 6 mg/dL	< 6 mg/dL	< 6 mg/dL
<b>Target for severe gout</b>	NA	< 5 mg/dL	NA
<b>Long-term target</b>	< 6 mg/dL	< 6 mg/dL	< 6 mg/dL

1. John D. FitzGerald, et al. Arthritis Care & Research 2020
2. Richette P, et al. Ann Rheum Dis 2016



\*highly selective urate reabsorption inhibitor (SURI)

# Clinical differences between allopurinol and febuxostat

Characteristics	Allopurinol	Febuxostat
<b>Chemical structure and activity</b>	Purine, Reversible, non-selective inhibitor of xanthine oxidase (reduced form)	Non-purine, Non-reversible, selective inhibitor of xanthine oxidase (oxidized/reduced form)
<b>Excretion</b>	Mainly kidney	Mainly intestine
<b>Daily dose</b>	100-800 mg	40-120 mg
<b>Dosing in renal insufficiency</b>	Dosage adjustment required (starting dose)	No dosage adjustment required in GFR > 30)
<b>SCARs</b>	1.6/1000 (Thais)	0.2/1000
<b>Special consideration</b>	Drug interaction (eg, azathioprine)	Drug interaction (eg, azathioprine) Cautious use in CVD or high CV risk

1. Yu KH. Recent Pat Inflamm Allergy Drug Discov 2007;1:69–75.

4. Hu M, et al. Ther Clin Risk Manag 2008;4:1209–1220.

7. Chen CH, et al. Clin Pharmacol Ther. 2019 Aug;106(2):391-401.

2. Edwards NL. Rheumatology (Oxford) 2009;48.

5. Teng GG, et al. Drugs 2006;66(12),1547-1563

3. Reinders MK, et al. Clin Interv Aging 2010;5:7–18.

6. Limkobpaiboon S et al., Chula Med J 54(5):467–478.

# Risk factors for allopurinol-induced severe cutaneous adverse reactions (SCARs)

- Genetic risk
    - HLA-B\*5801 allele **TRA 2025 - “Conditionally recommend testing”**
      - Thai, HR = 348.3 (prevalent 14-19 %); Han Chinese, HR= 580; Japan, HR= 65.6
    - Other HLA-A\*2402, DRB1\*1302
  - Non-genetic risk
    - Diuretics
    - CKD
    - Higher initial dose of allopurinol therapy
- 
- 90% AHS occurred within 180 days after initiation
  - **SCARs** = acute generalised exanthematous pustulosis (AGEP), drug reaction with eosinophilia and systemic symptoms (DRESS), Stevens–Johnson syndrome (SJS), toxic epidermal necrolysis (TEN), and Stevens–Johnson/toxic epidermal necrolysis overlap syndrome (SJS/TEN).
  - Pooled prevalence of HLA-B\*58:01 genotyping in the allopurinol-tolerant group = 10% in Asian

# Current uricosuric agent

Drug	Starting-maximum dose (mg/day)	Efficacy/safety data in eGFR (mL/min/1.73m <sup>2</sup> )	Remark
Probenecid	500-2000	> 50	High drug interaction (ASA, MTX, beta-lactam)
<b>Benzbromarone</b>	<b>50-200</b>	<b>&gt; 20</b>	Strong CYP2C9 inhibition High drug interaction (ASA, Warfarin, Dilantin)
Sulfinpyrazone	100-800	> 30	
			Blood dyscrasias if dose 600-800 mg/day Phenylbultazone allergy
Dotinurad	1-4	> 30	

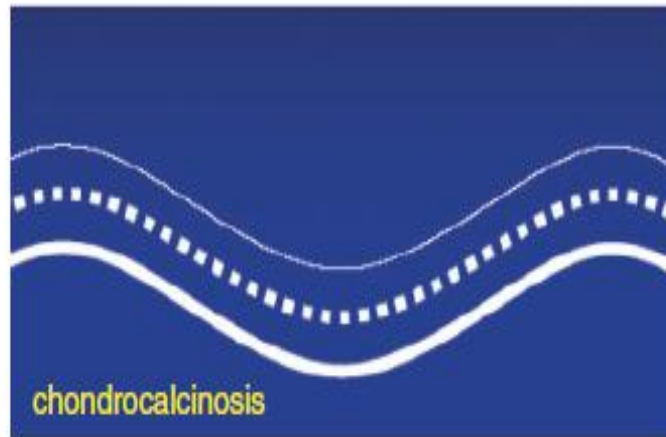
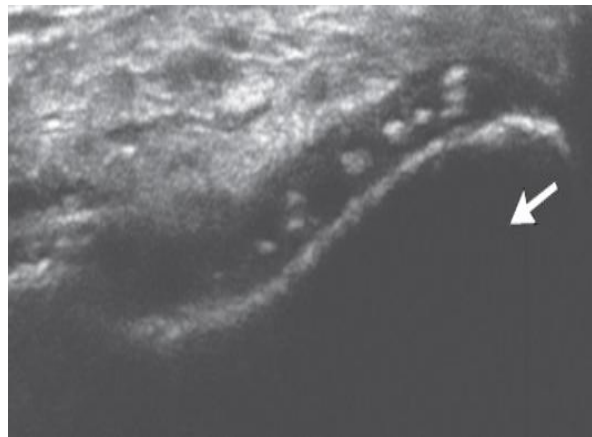
# Case 2

- 67M, ESKD s/p kidney transplant, hyperparathyroidism, T2DM
- OPD: Acute monoarthritis (knee). No fever.
- Synovial fluid: rhomboid crystals, WBC 24,000, Gram stain negative.
- eGFR 50 mL/min/1.73 m<sup>2</sup>.
- Meds: Tacrolimus, MMF, Fluconazole.

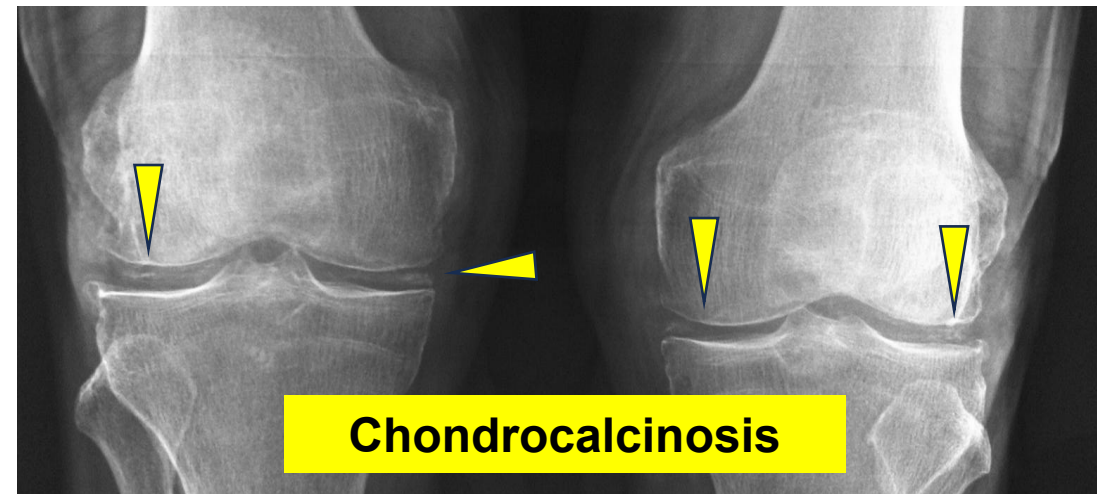
## Best treatment?

- A. Colchicine 0.6 mg BID
- B. Intra-articular steroid
- C. Prednisolone 30 mg daily
- D. Naproxen 250 mg BID
- E. ACTH

# CPPD-associated arthropathies



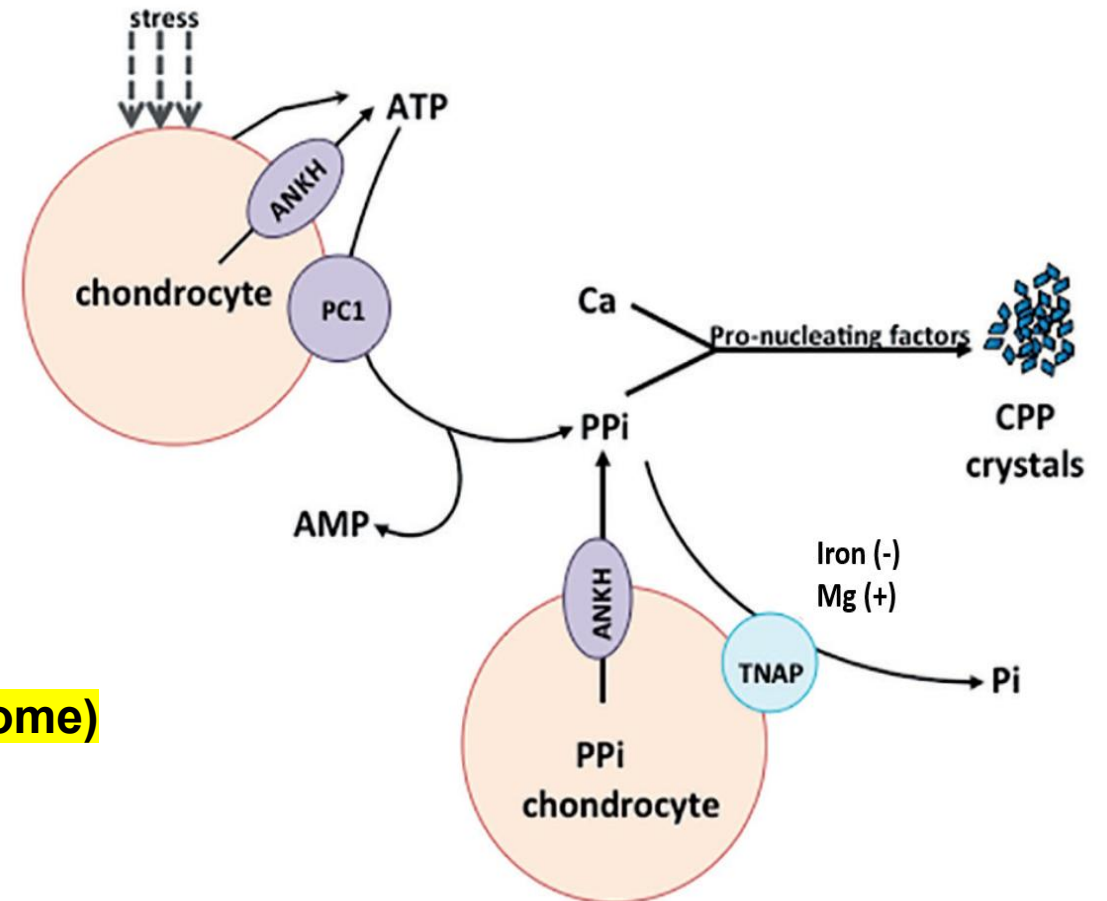
US: Hyperechoic linear in midzone of cartilage



# CPPD-associated arthropathies

CPP crystal nucleation = Extracellular PPI (ePPI)/Pi + Calcium + Cartilage matrix

- Aging > 70 years
- OA
- Common: knee, wrist, shoulder
- **Onset < 55 years**
  - JIA
  - Meniscectomy
  - Joint trauma
  - **Systemic disease (4H)**
    - Hypophosphatasia (ALP)
    - Hyperparathyroid
    - Hemochromatosis (hereditary)
    - Hypomagnesemia (Gitelman-Bartter's syndrome)
    - Hypothyroidism
    - Acromegaly
    - Wilson's disease



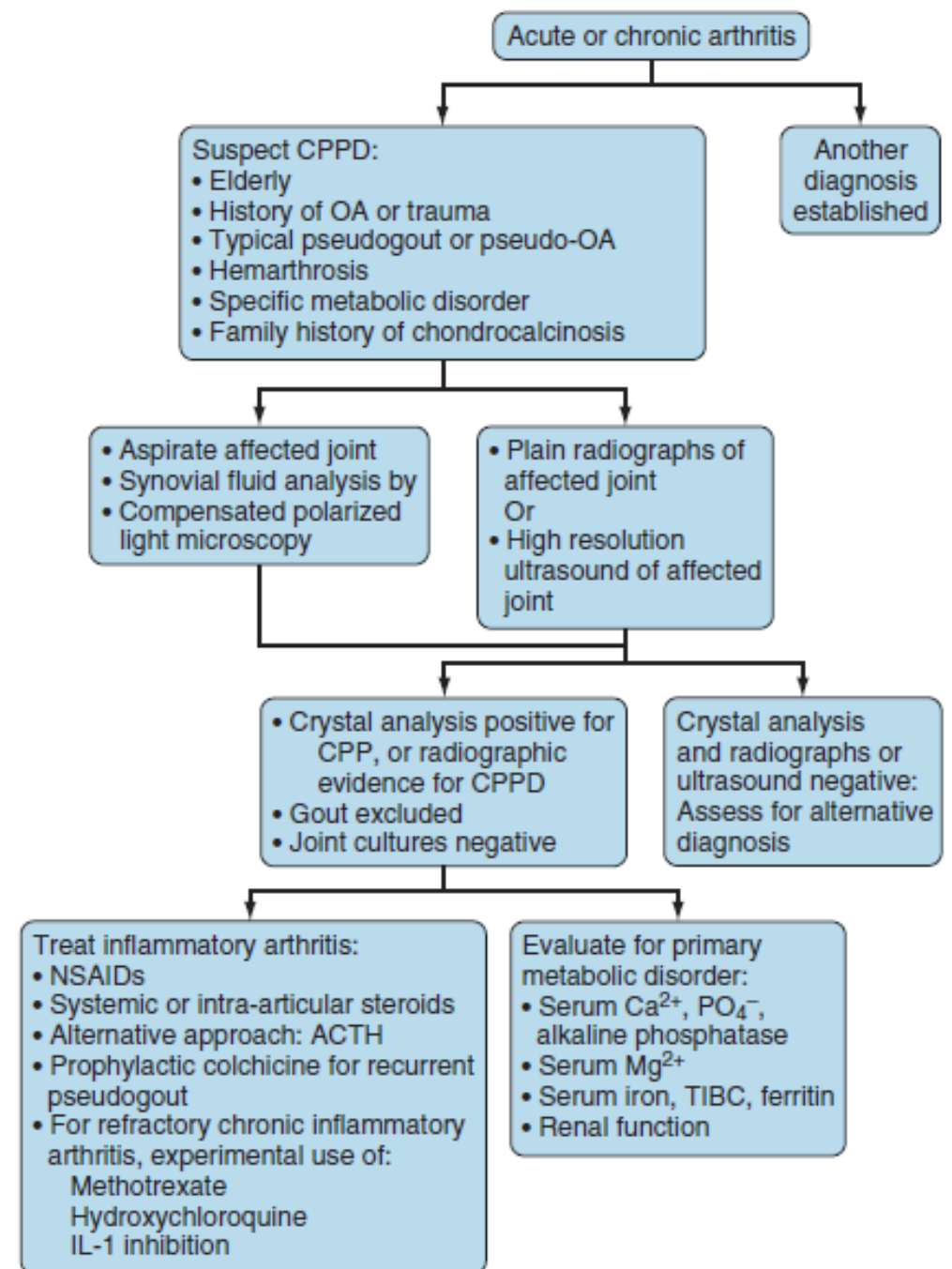
# Treatment of CPPD-associated arthropathies

## Treat inflammatory arthritis:

- NSAIDs
- Systemic or intra-articular steroids
- Alternative approach: ACTH
- Prophylactic colchicine for recurrent pseudogout
- For refractory chronic inflammatory arthritis, experimental use of:
  - Methotrexate
  - Hydroxychloroquine
  - IL-1 inhibition

## Evaluate for primary metabolic disorder:

- Serum  $\text{Ca}^{2+}$ ,  $\text{PO}_4^-$ , alkaline phosphatase
- Serum  $\text{Mg}^{2+}$
- Serum iron, TIBC, ferritin
- Renal function



# Acute treatment of crystal induced arthritis in CKD

CKD Stage  $\leq 3$

Prednisone 30mg taper over 5 days<sup>a</sup>

OR

Loading dose: Colchicine 1.2mg orally,  
and 0.6mg one hour later<sup>b</sup>  
May continue as 0.6mg once or twice daily  
after day 1, until resolution  
Consider changing treatment if no  
response after 36 hours<sup>7</sup>

OR

Intra-articular steroids if 1-2 joints affected

OR

Anakinra or canakinumab if above  
treatment fails or is contraindicated

CKD Stage  $> 3$

Prednisone 30mg taper over 5 days<sup>a</sup>

OR

Intra-articular steroids if 1-2 joints affected

OR

Anakinra or canakinumab if above  
treatment fails or is contraindicated

Avoid colchicine if alternatives are  
available  
May use as much as the loading dose of  
colchicine once with close monitoring, with  
no additional colchicine for 2 weeks<sup>b</sup>

Colchicine should be avoided  
in severe renal impairment  
(CLcr  $< 30$  ml/min)

\*NSAIDs are contraindicated in CKD.

### ตารางที่ 3 ขนาด รูปแบบ และระยะเวลาการบริหารยารักษาเกาต์กำเริบเฉียบพลัน

ชื่อยา	ขนาดยา	รูปแบบยา	ระยะเวลาการใช้ยา
Colchicine	1.2 มก.ทันที	รับประทาน	วันแรก
	0.6 มก. 1 ชม. ถัดไป <sup>§</sup>		
	1.2 มก. ทันที		
Colchicine	0.6 มก. ชม. ที่ 1 และ 12 <sup>^</sup>	รับประทาน	วันแรก
	0.5-0.6 มก. ทุก 8-12 ชม. <sup>%,^</sup>		
NSAIDs	ตามขนาดแนะนำของยาแต่ละตัว	รับประทาน	จนกว่าจะหาย
Prednisolone	30-35 มก./วัน	รับประทาน	4-5 วัน
ACTH	100 หน่วยสากล <sup>#</sup>	ฉีดเข้ากล้ามเนื้อ	ครั้งเดียว
Betamethasone	7 มก.	ฉีดเข้ากล้ามเนื้อ	ครั้งเดียว
Triamcinolone acetonide	40 มก.	ฉีดเข้ากล้ามเนื้อ	ครั้งเดียว
	20-40 มก.*	ฉีดเข้าข้อ	ครั้งเดียว

ACTH; adrenocorticotrop hormone, ชม.; ชั่วโมง, มก.; มิลลิกรัม

\*ขึ้นอยู่กับขนาดของข้อ, <sup>#</sup> international unit, <sup>§</sup> Terkeltaub RA, et al Arthritis Rheum 2010,<sup>^</sup> Laosuksri P, et al. IJRD 2023, <sup>%</sup> Roddy E, et al. Ann Rheum Dis 2020

## Colchicine prophylaxis dose

GFR (ml/min)	Colchicine oral dosage (mg/day)
≥ 50	0.6 mg bid
35-49	0.6 mg OD
10-34	0.6 mg Q 2-3 days
<10 or on hemodialysis or significant hepatic or biliary dysfunction	Avoid colchicine
Aged ≥ 70 years	50% of usual maintenance dose

# Guidelines for colchicine dosing in the presence of CYP3A4/P-glycoprotein inhibitors

FDA classification	Concomitant drug	Dosing recommendations	
		Acute gout flare	Prophylaxis of gout flares
Strong P-gp inhibitor	<b>Cyclosporine (Tacrolimus)</b>	<b>0.6 mg then repeated no earlier than 3 days</b>	<b>0.3 mg once/day or 0.3 mg every other day</b>
Strong CYP3A4 inhibitors	<b>Clarithromycin Ketoconazole Itraconazole Ritonavir</b>	0.6 mg then repeated no earlier than 3 days	0.3 mg once/day or 0.3 mg every other day
Moderate CYP3A4 inhibitors	<b>Diltiazem Verapamil Erythromycin Fluconazole Grapefruit juice</b>	1.2 mg then repeated no earlier than 3 days	0.6 mg once/day or 0.3 mg once/day
Weak CYP3A4 inhibitor	<b>Azithromycin</b>	<b>No dose reduction</b>	No dose reduction
Other potentially significant drug-drug interactions	<b>Statin , fibrate, or digoxin</b>	<b>No dose reduction</b> Should be monitored for any signs or symptoms of muscle	No dose reduction Should be monitored for any signs or symptoms of muscle

## Case 3

- A 65F with RA on MTX 10 mg/week + PRED 5 mg/day for 3 month
- Today: improvement (PGA 8→6/10), but still has functional limitation and tender joints, without swelling.
- ESR improved but elevated (50 mm/hr, last 80 mm/hr). ALT normal.
- X-ray: erosions, Anti-CPP/RF positive

### Next step?

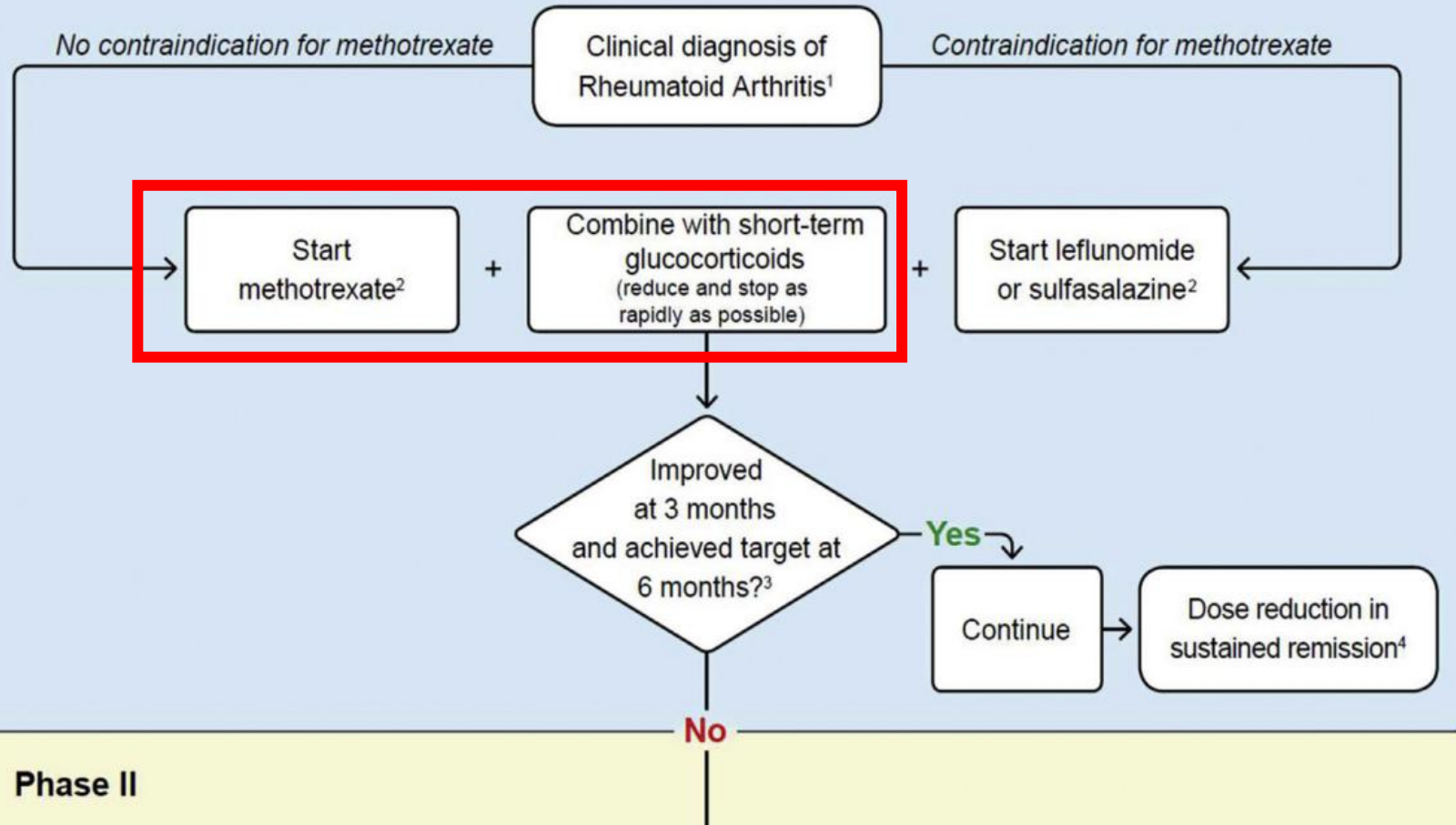
- A. Observe 3 more months
- B. Add 2 gm of sulfasalazine
- C. Reduce prednisolone to 2.5 mg/day
- D. Increase prednisolone to 7.5 mg/day
- E. Increase methotrexate to 15 mg/week

# 2021 ACR Recommendation of RA

Recommendations	Certainty of evidence
<p>Initiation of treatment in DMARD-naive patients with moderate-to-high disease activity</p> <p>Methotrexate monotherapy is <b>strongly</b> recommended over:</p> <ul style="list-style-type: none"> <li>Hydroxychloroquine or sulfasalazine</li> <li>bDMARD or tsDMARD monotherapy</li> <li>Combination of methotrexate plus a non-TNF inhibitor bDMARD or tsDMARD¶</li> </ul> <p>Methotrexate monotherapy is <b>conditionally</b> recommended over:</p> <ul style="list-style-type: none"> <li>Leflunomide</li> <li>Dual or triple csDMARD therapy¶</li> <li>Combination of methotrexate plus a TNF inhibitor¶</li> </ul>	<p>Very low/low‡</p> <p>Very low/moderate</p> <p>Low/very low</p> <p>Low</p> <p>Moderate</p> <p>Low</p> <p>Very low</p>
<p>Initiation of a csDMARD without short-term (&lt;3 months) glucocorticoids is <b>conditionally</b> recommended over initiation of a csDMARD with short-term glucocorticoids.</p> <p>Initiation of a csDMARD without longer-term (≥3 months) glucocorticoids is <b>strongly</b> recommended over initiation of a csDMARD with longer-term glucocorticoids.</p>	<p>Moderate</p>
<p>Initiation of treatment in DMARD-naive patients with low disease activity</p> <ul style="list-style-type: none"> <li>Hydroxychloroquine is <b>conditionally</b> recommended over other csDMARDs.</li> <li>Sulfasalazine is <b>conditionally</b> recommended over methotrexate.</li> <li>Methotrexate is <b>conditionally</b> recommended over leflunomide.</li> </ul>	<p>Very low</p> <p>Very low</p> <p>Very low</p>
<p>Initiation of treatment in csDMARD-treated, but methotrexate-naive, patients with moderate-to-high disease activity#</p> <p>Methotrexate monotherapy is <b>conditionally</b> recommended over the combination of methotrexate plus a bDMARD or tsDMARD.**</p>	<p>Moderate/very low</p>

# EULAR recommendations for the management of RA 2026

## Phase I



- **Target:** Remission (or low disease activity if remission unlikely)
- **Timeline:** Aim within 6 months
- **Adjust therapy** if inadequate response by 3 months (~50% improvement expected)
- **Sustained remission:** ≥6 months (ACR/EULAR criteria)

**“GC-bridging”** should be considered when initiating or changing csDMARDs, in different dose regimens and routes of administration, **but should be tapered and discontinued as rapidly as clinically feasible.**

# Standard target dose of csDMARDs

csDMARD	Dose	Maximum
<b>Methotrexate</b>	<b>0.3 mg/kg/week</b>	<b>Max 15 mg/week</b>
Hydroxychloroquine	5 mg/kg/day	Max 400 mg/day
Sulfasalazine	40 mg/kg/day	Max 3 g/day
Leflunomide	20 mg/day	
Azathioprine	2 mg/kg/day	

# Poor prognostic factors

1. High acute phase reactant levels
2. High swollen joint count
3. Presence of RF and/or ACPA, especially at high levels
4. Presence of early erosions
5. Failure of 2 or more csDMARDs
6. Persistently moderate or high disease activity (after csDMARD therapy) according to composite measures including joint counts despite csDMARD therapy

May be consider early combination of csDMARDs (MTX, SSZ, HCQ) or initiation of biologic therapy

# Case 4

- **55F with RA**
- **Meds: MTX (weekly), adalimumab (q2w), prednisolone 10 mg/day**
- **Planning elective TKA in 1 month.**
- **Stable disease. No neck pain, hoarseness, or dyspnea.**
- **HTN well-controlled; no active cardiac conditions.**

**Best perioperative medication management?**

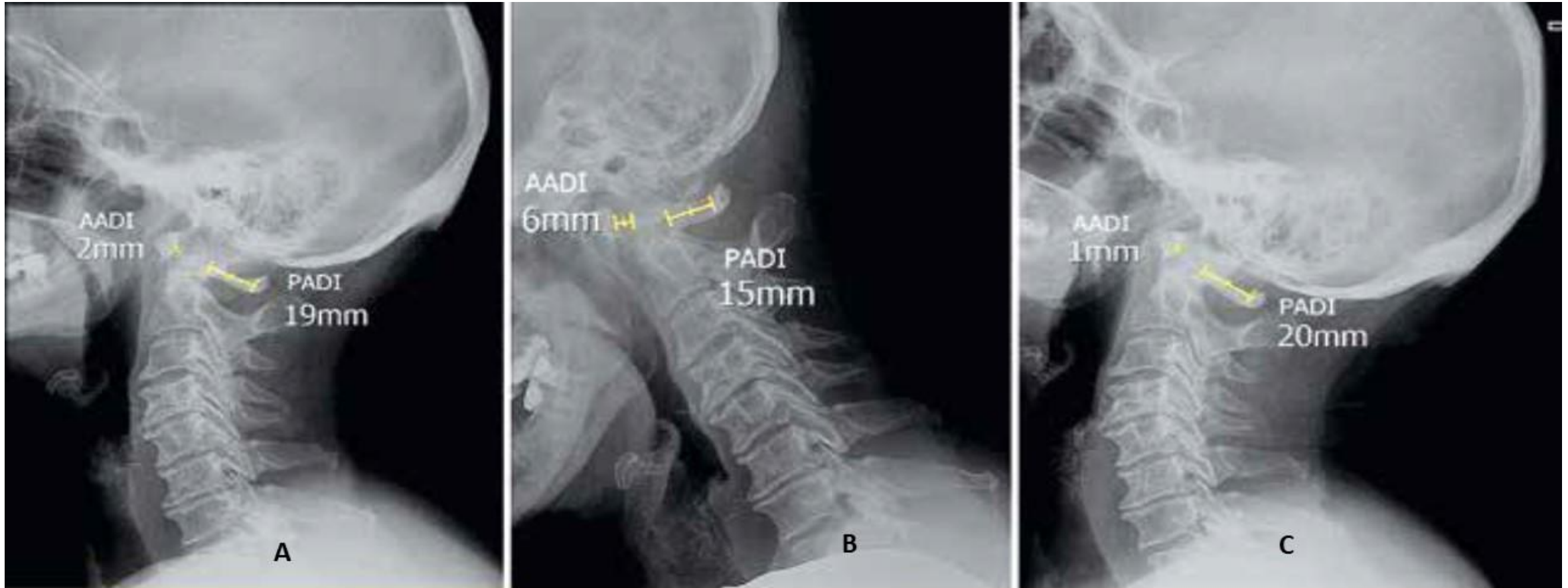
- A. Continue all**
- B. Reduce prednisolone**
- C. Hold MTX 1 week pre-op**
- D. Hold adalimumab; schedule surgery ~3 weeks after last dose**
- E. C+D**

# Pre-operative evaluation in patient with RA

## Pulmonary assessment

1. The upper airway - arthritis of the cricoarytenoid joints and temporomandibular joints
2. Interstitial lung disease

Cervical spine instability		
<b>Atlanto axial subluxation</b> <b>(AAS)</b>	Anterior Atlanto-Dental Interval (AADI)	> 5-8 mm.
	Posterior Atlanto-Dental Interval (PADI) or Space Available for the Cord (SAC)	< 14 mm.
<b>Subaxial subluxation</b> <b>(SAS)</b>	Subluxation occurring in joints on C3–C7 level	> 3.5 mm
<b>Cranial settling</b> <b>(CrS)</b>	Superior migration of the odontoid into foramen magnum	



## Anterior Atlanto-Dental Interval (AADI), Posterior Atlanto-Dental Interval (PADI)

lateral neutral (A), flexion (B), extension (C)

# 2022 ACR/AAHKS Guideline (1)

## ยาที่สามารถใช้ต่อได้ในช่วงผ่าตัด

	Dosing interval	Recommended timing of surgery since last medication dose
Medications to continue through surgery		
DMARDs: continue these medications through surgery (all patients)		
Methotrexate	Weekly	Anytime
Sulfasalazine	Once or twice daily	Anytime
Hydroxychloroquine	Once or twice daily	Anytime
Leflunomide (Arava)	Daily	Anytime
Doxycycline	Daily	Anytime
Apremilast (Otezla)	Twice daily†	Anytime†
Severe SLE-specific medications: continue these medications in the perioperative period in consultation with the treating rheumatologist‡		
Mycophenolate mofetil	Twice daily	Anytime
Azathioprine	Daily or twice daily	Anytime
Cyclosporine	Twice daily	Anytime
Tacrolimus	Twice daily (IV and PO)	Anytime
Rituximab (Rituxan)	IV every 4–6 months†	Month 4–6†
Belimumab SC (Benlysta)	Weekly†	Anytime†
Belimumab IV (Benlysta)	Monthly†	Week 4†
Anifrolumab (Saphnelo)§	IV every 4 weeks†	Week 4†
Voclosporin (Lupkynis)§	Twice daily†	Continue†

# 2022 ACR/AAHKS Guideline (2)

ยาที่แนะนำให้หยุดช่วงผ่าตัด	Dosing interval	Recommended timing of surgery since last medication dose
Medications to withhold prior to surgery¶		
Biologics: withhold these medications through surgery		
Infliximab (Remicade)	Every 4, 6, or 8 weeks	Week 5, 7, or 9
Adalimumab (Humira)	Every 2 weeks	Week 3
Etanercept (Enbrel)	Every week	Week 2
Golimumab (Simponi)**	Every 4 weeks (SQ) or every 8 weeks (IV)	Week 5 Week 9
Abatacept (Orencia)	Monthly (IV) or weekly (SC)	Week 5; week 2
Certolizumab (Cimzia)	Every 2 or 4 weeks	Week 3 or 5
Rituximab (Rituxan)	2 doses 2 weeks apart every 4–6 months	Month 7
Tocilizumab (Actemra)	Every week (SC) or every 4 weeks (IV)	Week 2; week 5
Anakinra (Kineret)	Daily	Day 2
IL-17 secukinumab (Cosentyx)	Every 4 weeks	Week 5
Ustekinumab (Stelara)	Every 12 weeks	Week 13
Ixekizumab (Taltz)§	Every 4 weeks†	Week 5†
IL-23 guselkumab (Tremfya)§	Every 8 weeks†	Week 9†
JAK inhibitors: withhold this medication 3 days prior to surgery#		
Tofacitinib (Xeljanz)	Daily or twice daily†	Day 4†
Baricitinib (Olumiant)§	Daily†	Day 4†
Upadacitinib (Rinvog)§	Daily†	Day 4†
Not severe SLE: withhold these medications 1 week prior to surgery		
Mycophenolate mofetil	Twice daily	1 week after last dose†
Azathioprine	Daily or twice daily	1 week after last dose
Cyclosporine	Twice daily	1 week after last dose†
Tacrolimus	Twice daily (IV and PO)	1 week after last dose†
Rituximab (Rituxan)	Every 4–6 months	Month 7
Belimumab IV (Benlysta)	Monthly†	Week 5†
Belimumab SC (Benlysta)	Weekly†	Week 2†

# Glucocorticoid Management

- Use of more than 10 mg of glucocorticoids per day (vs. no glucocorticoid use) resulted in a predicted risk for hospitalized infection of 13.25% and a predicted 1-year cumulative incidence of PJI of 3.83%.
- A lower cumulative GC dose was not associated with hypotension.
- 2022 ACR/AAHKS conditionally recommended patients with rheumatic disease undergoing THA or TKA who are receiving GCs, continuing their current daily dose of GCs rather than administering supraphysiologic doses of GCs on the day of surgery.

# Case 5

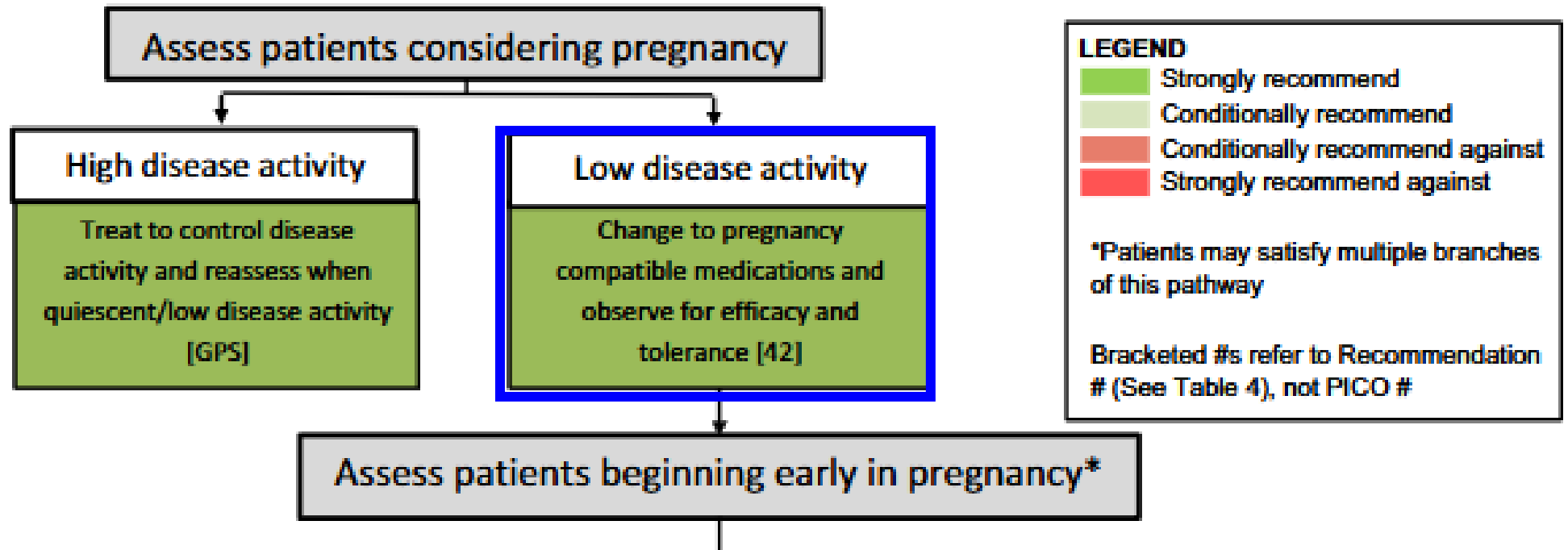
- 29F with SLE in remission ×12 months
- Planning pregnancy.
- On prednisolone 5 mg/day, MMF, HCQ.

**Best recommendation?**

- A. Continue all**
- B. Stop MMF  $\geq 6$  weeks before conception; continue others**
- C. Stop MMF → switch to azathioprine; observe 6 months before conception**
- D. Stop prednisolone**
- E. Continue only HCQ**

# 2020 ACR Guideline for the Management of Reproductive Health in Rheumatic and Musculoskeletal Diseases

Counsel RMD patients regarding improved maternal and pregnancy outcomes when disease is quiescent/low activity before pregnancy [GPS]. Co-management with rheumatology and other specialists preferred [GPS.]



# 2020 ACR Guideline for the Management of Reproductive Health in Rheumatic and Musculoskeletal Diseases

Medication	Pre-conception	During pregnancy	Breastfeeding
<b>Conventional medications</b>			
Hydroxychloroquine	++	++	++
Sulfasalazine	++	++ Folic 5 mg/day	++
Colchicine	++	++	++
Azathioprine, 6-mercaptopurine	++	++	+ Low transfer
Prednisone	+ Taper to <20 mg/day by adding pregnancy-compatible immunosuppressants	+ Taper to <20 mg/day by adding pregnancy-compatible immunosuppressants	+ After a dose of >20 mg, delay breastfeeding for 4 hours
Cyclosporine, tacrolimus	+ Monitor blood pressure	+ Monitor blood pressure	+ Low transfer
Nonsteroidal antiinflammatory drugs (cyclooxygenase 2 inhibitors not preferred)	+ Discontinue if the woman is having difficulty conceiving	+ Continue in first and second trimesters; discontinue in third trimester	+ Ibuprofen preferred

# 2020 ACR Guideline for the Management of Reproductive Health in Rheumatic and Musculoskeletal Diseases

**Table 3.** Maternal medication use: overview of medication use before and during pregnancy, and during breastfeeding

Medication	Pre-conception	During pregnancy	Breastfeeding
<b>Not compatible with pregnancy</b>			
Methotrexate	XX Stop 1–3 months prior to conception	XX Stop and give folic acid 5 mg/day	X Limited data suggest low transfer
Leflunomide	XX Cholestyramine washout if detectable levels	XX Stop and give cholestyramine washout	XX
Mycophenolate mofetil and mycophenolic acid	XX Stop >6 weeks prior to conception to assess disease stability	XX	XX
Cyclophosphamide	XX Stop 3 months prior to conception	+ Life-/organ-threatening disease in second and third trimesters	XX
Thalidomide	XX Stop 1–3 months prior to conception	XX	XX
Tofacitinib, apremilast, baricitinib	Unable to determine due to lack of data; small molecular size suggests transfer across the placenta and into breast milk		



# 2024 EULAR recommendations for use of antirheumatic drugs (2)

Drugs compatible with breastfeeding				Drugs may be considered in breastfeeding if no alternative medication can be used		Drugs to be avoided in breastfeeding due to insufficient data	
Drug	LoE/GoR	Drug	LoE/GoR	Drug	LoE/GoR	Drug	LoE/GoR
NSAID: preferential use of ibuprofen (IBU) and celecoxib (CEL)	2a/B IBU 4/C CEL	IL12/23i: ustekinumab	2a/B	bosentan sildenafil	4/C	apremilast	5/D
prednisone, prednisolone IV methylprednisolone	2a/B	IL-6i: tocilizumab, sarilumab	4/C	methotrexate ≤25 mg weekly	4/C	avacopan	5/D
hydroxychloroquine (HCQ) chloroquine (CQ)	2a/B HCQ 4/C CQ	IL-1i: anakinra, canakinumab	2a/B			cyclophosphamide	4/D
sulfasalazine	2a/C	IL17i: secukinumab ixekizumab	5/D			etoricoxib	5/D
azathioprine 6-mercaptopurine	2a/B	IL-23i: guselkumab risankizumab	5/D			iloprost	5/D
cyclosporine tacrolimus	2a/B	IFNAR1i: anifrolumab	5/D			JAKi: tofacitinib (TOF), baricitinib, upadacitinib, filgotinib	4/D (TOF), others 5/D
IVIG	2a/B	IL-5i: mepolizumab	5/D			leflunomide	5/D
colchicine	2a/B	C5i: eculizumab	5/D			mycophenolate	5/D
TNFi: adalimumab certolizumab etanercept golimumab infliximab	2a/B	anti-T cell: abatacept	4/C			voclosporin	5/D
		anti-B cell: rituximab (RTX) belimumab (BEL)	2a/B RTX 4/C BEL				

Counsel RMD patients regarding improved maternal and pregnancy outcomes when disease is quiescent/low activity before pregnancy [GPS]. Co-management with rheumatology and other specialists preferred [GPS.]

Assess patients considering pregnancy

**High disease activity**  
 Treat to control disease activity and reassess when quiescent/low disease activity [GPS]

**Low disease activity**  
 Change to pregnancy compatible medications and observe for efficacy and tolerance [42]

**LEGEND**

- Strongly recommend
- Conditionally recommend
- Conditionally recommend against
- Strongly recommend against

\*Patients may satisfy multiple branches of this pathway

Bracketed #s refer to Recommendation # (See Table 4), not PICO #

Assess patients beginning early in pregnancy\*

**SLE**

- Continue HCQ (if on) [57]
- Start HCQ (if not on and no contraindication) [58]
- Low dose aspirin [56]
- Laboratory assessment of disease activity at least once per trimester [64]

**Anti-Ro/La (+)**

**No history NLE**

- HCQ [69]
- Serial fetal echo week 16-26 [67]

**History of NLE**

- HCQ [70]
- Weekly fetal echo week 16-26 [68]

**Abnormal fetal echocardiogram**

- Brief course of dexamethasone if 1<sup>st</sup> or 2<sup>nd</sup> degree heart block [71,72]
- Against dexamethasone if 3<sup>rd</sup> degree (complete) heart block [73]

**Positive aPL Test**

**No APS**

- Low dose aspirin [45]
- Against prophylactic heparin or HCQ [46, 44A]

**OB APS**

- Low dose aspirin + prophylactic heparin until 6-12 weeks post-partum [48,84]
- HCQ [44B]
- Against therapeutic heparin or IVIG [49-50]
- Against prednisone [51]

**Thrombotic APS**

- Low dose aspirin + therapeutic heparin [52]
- HCQ [44B]

# Case 6

- 28F with SLE
- Active LN and thrombotic APS
- Meds: HCQ, MMF, prednisolone, warfarin.
- LAB: LA persistently positive.

**Best contraceptive method?**

- A. Copper IUD
- B. Progestin IUD
- C. Progestin implant
- D. DMPA+ condom
- E. Progestin-only pill

# Safety and efficacy of various contraceptive methods in RMD

Method	Safety in women with RMD	1-year failure rate, %†
<b>Highly effective (LARC)</b>		
Copper IUD	Safe in all women with RMD; may increase menstrual bleeding	<1
Progestin IUD	Safe in all women with RMD; may decrease menstrual bleeding	<1
Progestin implant	Limited data, but likely safe in all women with RMD	<1
<b>Effective</b>		
Progestin-only pill (daily)	Safe in all women with RMD; higher rate of breakthrough bleeding than with combined contraceptives; must take same time every day for efficacy	5–8
DMPA (IM injection every 12 weeks)	Safe in most women with RMD; <u>exceptions: positive aPL, at high risk for OP</u>	3
Combined estrogen and progesterone pill (daily)	Safe in most women with RMD; <u>exceptions: positive aPL, very active SLE</u>	5–8
Transdermal patch (weekly)	Safe in most women with RMD; <u>exceptions: positive aPL, SLE; serum estrogen levels higher than with pill or vaginal ring</u>	5–8
Vaginal ring (monthly)	Safe in most women with RMD; <u>exceptions: positive aPL, very active SLE</u>	5–8
<b>Less effective</b>		
Diaphragm	Safe in all women with RMD	12
Condom	Safe in all women with RMD; only form to prevent STD	18
Fertility awareness-based methods‡	Safe in all women with RMD; limited efficacy, especially if menses are irregular	24
Spermicide	Safe in all women with RMD; use with condoms or diaphragm to improve efficacy	28

\* RMD = rheumatic and musculoskeletal disease; LARC = long-acting reversible contraception; IUD = intrauterine device; DMPA = depot medroxyprogesterone acetate; IM = intramuscular; aPL = antiphospholipid antibody; OP = osteoporosis; SLE = systemic lupus erythematosus; STD = sexually transmitted disease.

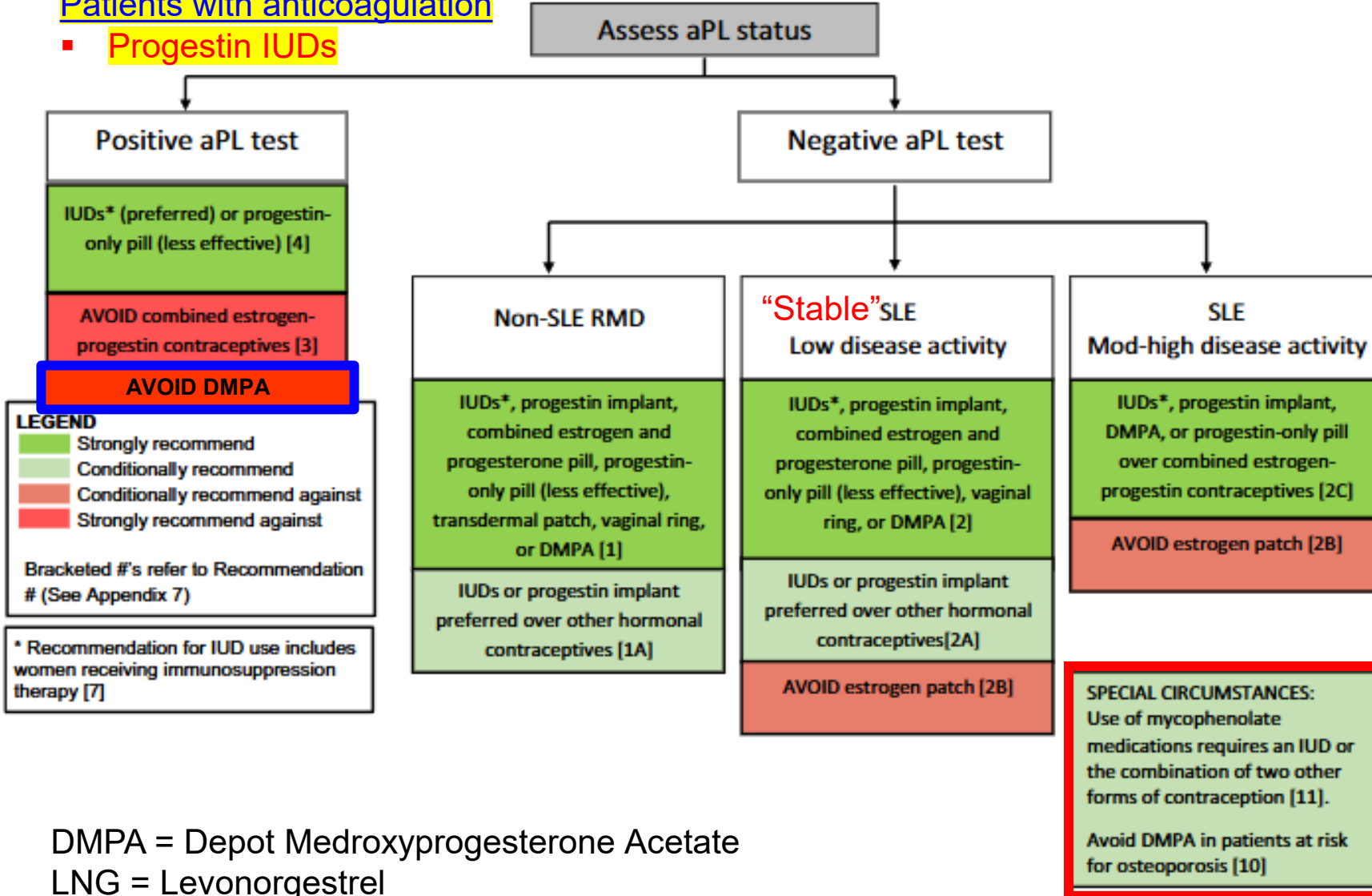
† Percent of women who will become pregnant within the first year of typical use.

‡ Methods based on the timing of the menstrual cycle.

Discuss contraception and pregnancy planning at initial or early visit with women of reproductive age and counsel regarding efficacy and safety [GPS]. Recommend barrier methods if more effective methods are contraindicated [GPS]. Recommend emergency (post-coital) contraception when necessary [6].

**Patients with anticoagulation**

**Progestin IUDs**



**LEGEND**  
 Strongly recommend  
 Conditionally recommend  
 Conditionally recommend against  
 Strongly recommend against  
 Bracketed #'s refer to Recommendation # (See Appendix 7)

\* Recommendation for IUD use includes women receiving immunosuppression therapy [7]

DMPA = Depot Medroxyprogesterone Acetate  
 LNG = Levonorgestrel

**aPL test** should be performed

- SLE or SLE-like disease
- Patients with suggestive histories or physical findings

**Positive aPL**

**Persistent and mod-high titer**

- 2 positive test results at least 12 weeks apart *plus*
- Moderate-high-titer aCL/anti-β2GPI (≥40 units)
- Positive LAC

**SPECIAL CIRCUMSTANCES:**  
 Use of mycophenolate medications requires an IUD or the combination of two other forms of contraception [11].  
 Avoid DMPA in patients at risk for osteoporosis [10]

# Case 7

- 25F with SLE → active lupus nephritis
- BP 150/90 mmHg
- Cr 2.3 mg/dL (eGFR 25 cc/min), albumin 2.4 mg/dL
- Proteinuria 3.2 g/day, RBC casts.
- Biopsy: Class IV with crescents and fibrinoid necrosis, no thrombotic microangiopathy
- Already given IV methylprednisolone.

**Next best treatment?**

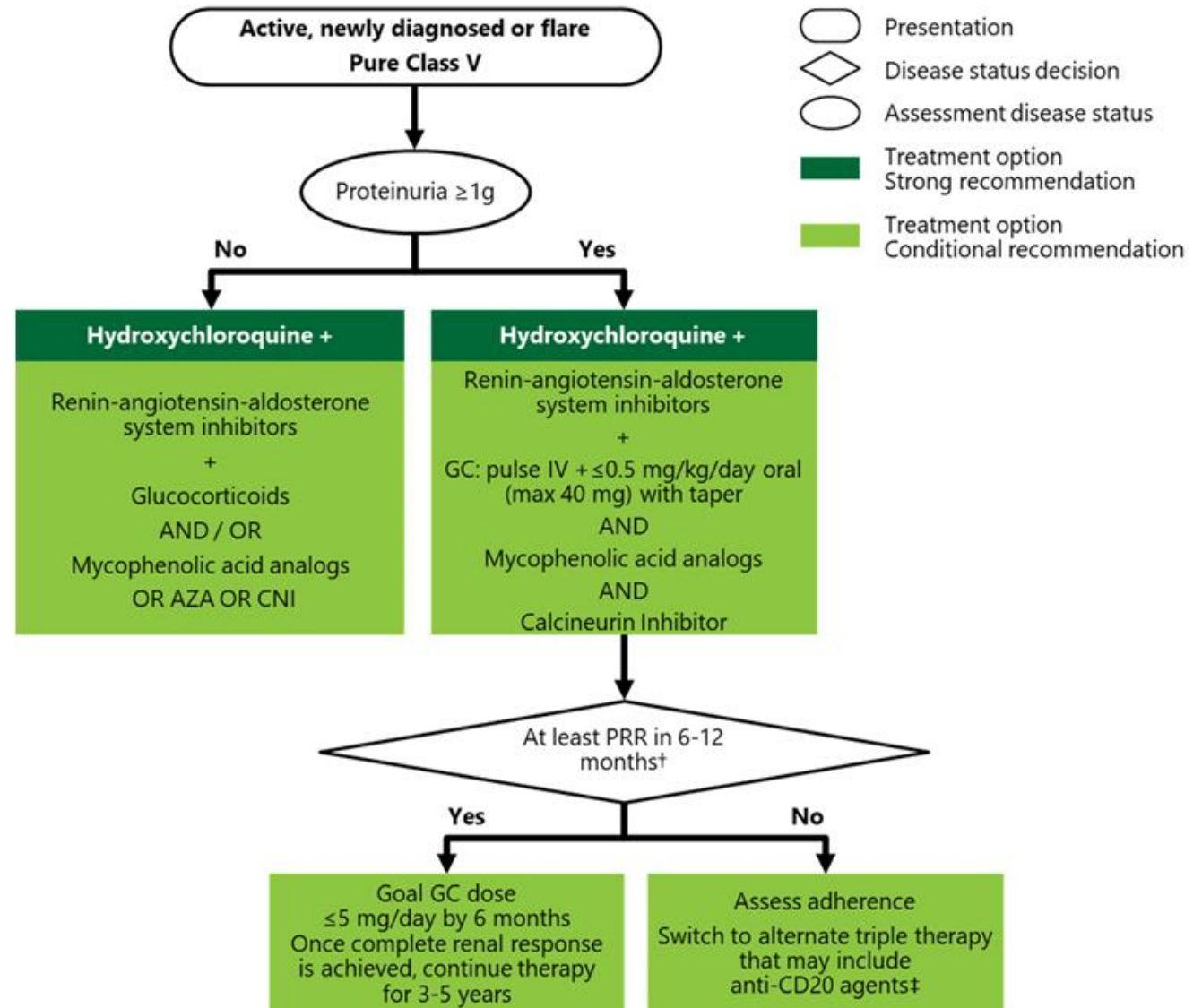
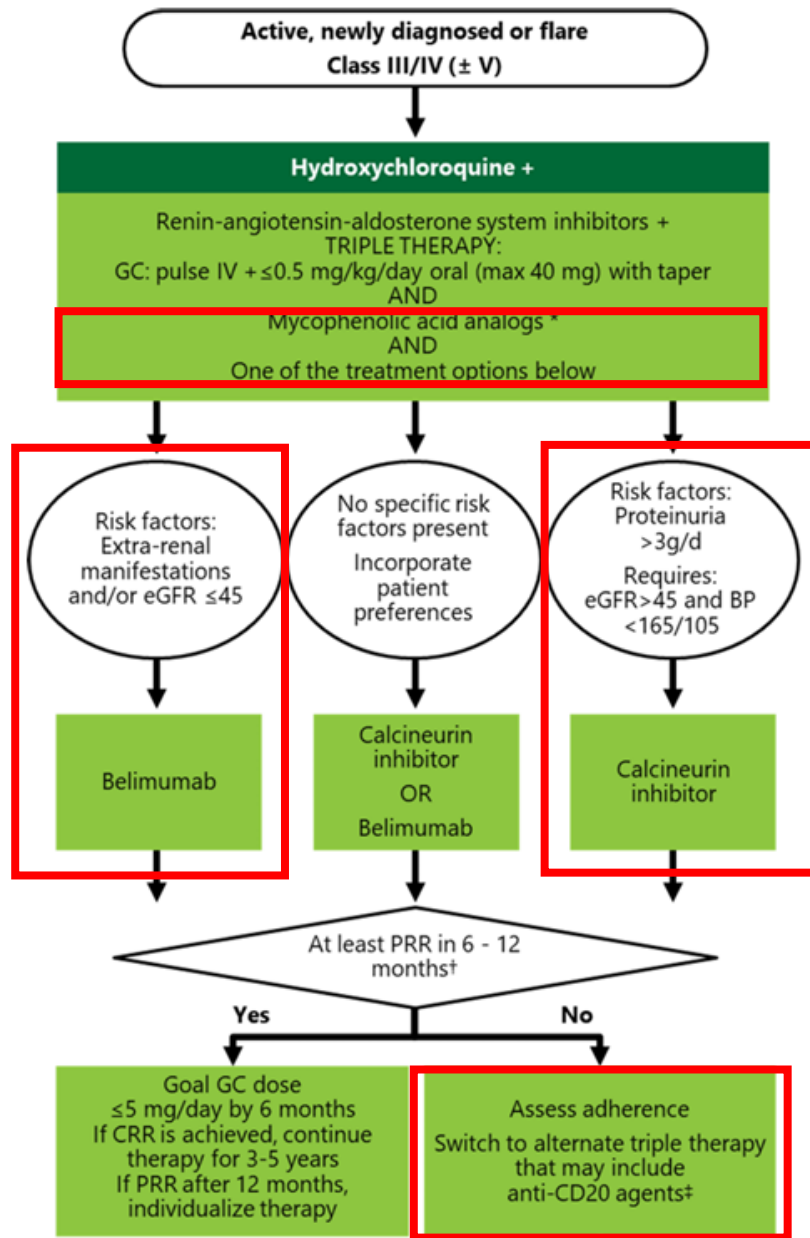
- A. MMF
- B. IV cyclophosphamide (Euro-Lupus)
- C. IV cyclophosphamide (NIH)
- D. MMF + tacrolimus
- E. MMF + Rituximab

# Current treatment of lupus nephritis

- Choice of treatment is directly based on **renal pathology and severity**.

		Double Immunosuppressive Regimen			Triple Immunosuppressive Regimen			
		<b>Cyclophosphamide (CYC)</b>			<b>Calcineurin Inhibitors + MPAA</b>		<b>Belimumab + MPAA/CYC</b>	
		<b>Modified NIH Regimen</b>	<b>ELNT Regimen</b>	<b>Oral Regimen</b>	<b>Mycophenolic Acid Analogs (MPAA)</b>	<b>Tacrolimus + MPAA</b>	<b>Voclosporin + MPAA</b>	
	<b>Dosage</b>	i.v. 0.5-1 g/m <sup>2</sup> monthly x 6 or i.v. 7.5-15 mg/kg monthly x 6	i.v. 500 mg q2wk x 6	p.o. 1.0-1.5 mg/kg/day	MMF p.o. 1.0 - 1.5 g b.i.d. or MPA p.o. 0.72 - 1.08 g b.i.d.	Tacrolimus (through level approx. 4-6 ng/ml) + reduced-dose MPAA (1-2g/d)	Voclosporin 23.7 mg b.i.d. + MPAA (2-3 g/d)	Belimumab (i.v. 10 mg/kg q2wk x 3 doses, then q4wk) + MPAA (2-3 g/d) or i.v. CYC (500 mg q2wk x 6 doses)
	<b>Length</b>	6 months	3 months	3-6 months	At least 6 months	At least 6 months Up to 24 months	Up to 36 months	Up to 2.5 years
		<b>Guideline Recommendations</b>						
<b>KDIGO</b>		Active Proliferative LN Difficulty adhering to oral regimen	Active Proliferative LN Difficulty adhering to oral regimen White patients	Active Proliferative LN	Active proliferative LN High risk of infertility, such as patients who have a moderate-to-high prior cyclophosphamide exposure	Relatively preserved kidney function (eGFR ≥ 45 mL/min) Proteinuria > 3 g/day Cannot tolerate standard dose MPAA Unfit for or will not use CYC-based regimens	Relatively preserved kidney function (eGFR ≥ 45 mL/min) Proteinuria > 3 g/day Cannot tolerate standard dose MPAA Unfit for or will not use CYC-based regimens	Patients with repeated kidney flares or at high risk for progression to kidney failure due to severe chronic kidney disease.

# 2025 ACR Guideline for LN



# 2025 EULAR Recommendation for LN



## Diagnosis / Targets

- **Kidney biopsy**  
Indispensable. Repeat if clinical uncertainty
- **Target – Prevention of Chronic kidney disease Flares**
- **Milestones**  
Kidney function.  
Preservation or improvement by 3 months  
Proteinuria.  
Reduction by 25% at 3 months  
Reduction by 50% at 6 months  
uPCR <700 mg/g by 12 months



## Immune Treatment

### “Quadruple Regimen”

- **Early combination tx**  
Hydroxychloroquine and Glucocorticoids with Immunosuppressive and CNI or biologic
- **Glucocorticoids**  
Start with pulses  
Continue with 0.3-0.7 mg/kg/d prednisone  
Taper to ≤5mg/day by 4-6 mo and withdraw when possible
- **Immunosuppressives**  
MMF, low-dose CYC
- **CNI**  
Voclosporin or TAC (CsA)
- **Biologics**  
Belimumab, Obinutuzumab



## Non-immune Treatment

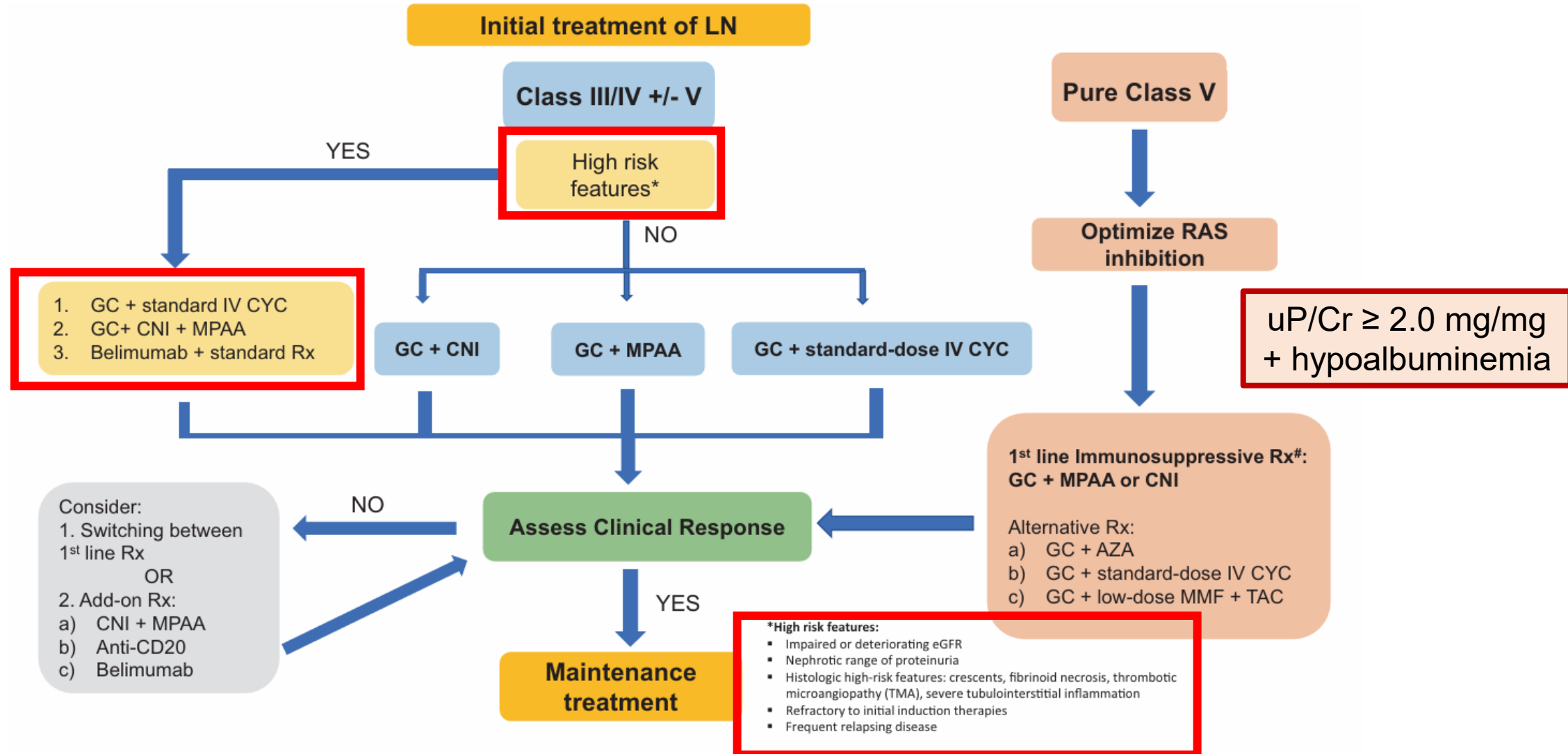
- **Nephroprotection**  
Low salt (less than 5g/day)  
Control BP (RAASi 1st choice)  
SGLT2 inhibitors (in stable ds. if residual proteinuria after 12m)
- **Dyslipidemia**
- **Vaccinations**  
Influenza, COVID-19, HZV, S. pneumoniae
- **Bone Health**



## Severe or Refractory

- **Severe**  
Consider high-dose i.v. CYC + Pulse i.v. MP
- **Refractory**  
Combination i.v.-CYC and B-cell depletion  
Addition of a CNI if heavy proteinuria  
Experimental therapies in the context of a protocol
- **Thrombotic Microangiopathy**  
Plasma exchange  
Complement inhibitors  
Anti-vWf (caplacizumab)

# 2025 APLAR Guideline for LN



**FIGURE 1** | Algorithm for initial treatment of lupus nephritis. LN, lupus nephritis; GC, glucocorticoid; IV, intravenous; CYC, cyclophosphamide; CNI, calcineurin inhibitor; MPAA, mycophenolic acid analogue; RAS, renin angiotensin system; Rx, treatment; AZA, azathioprine; TAC, tacrolimus.

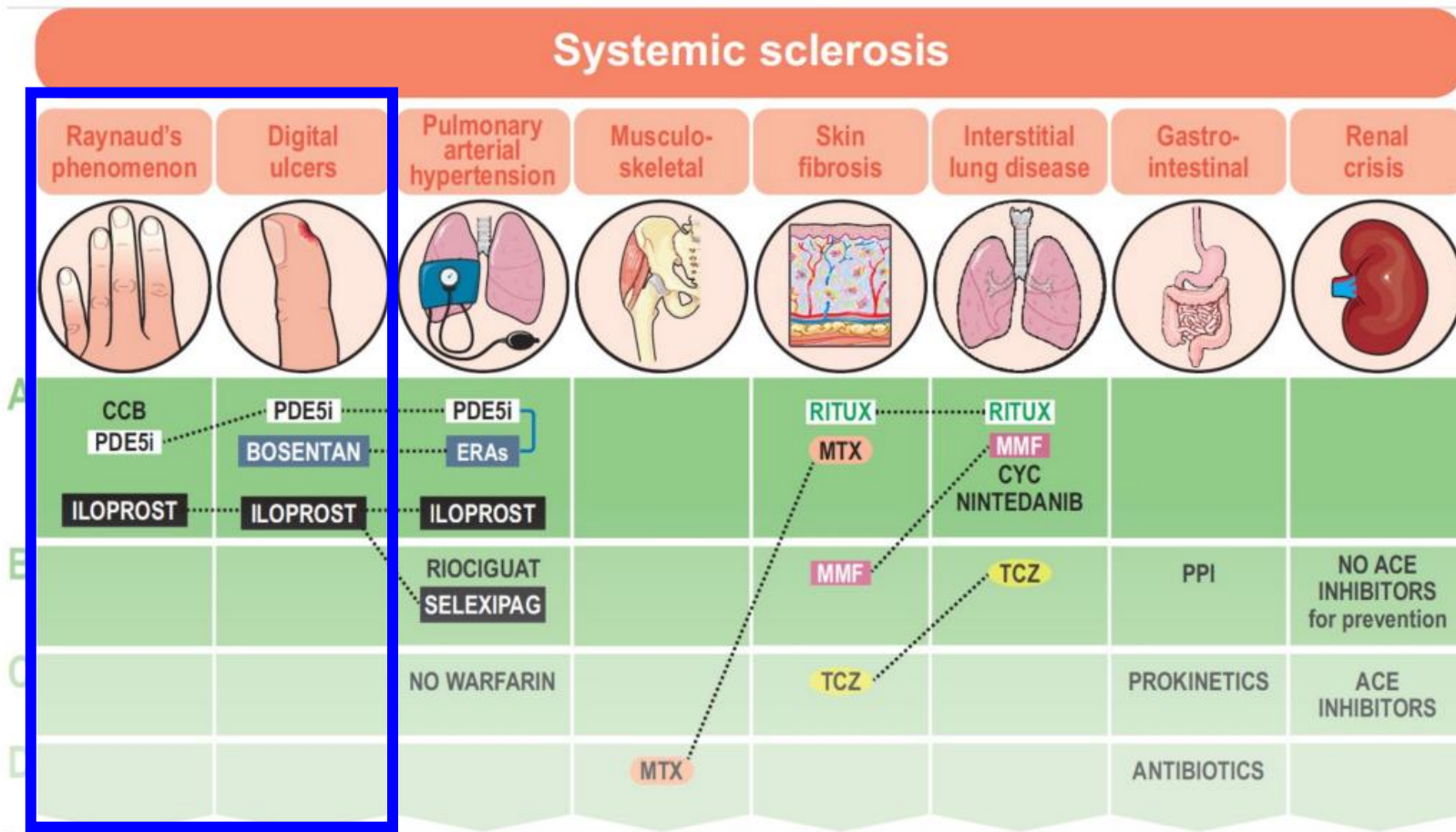
# Case 8

- 38M with systemic sclerosis with reflux esophagitis
- Frequent Raynaud's + new 2 digital ulcers (2 wks)
- PE: digital pitted scars

## Most appropriated management

- A. Long-acting nifedipine
- B. Sildenafil
- C. Bosentan
- D. Iloprost
- E. Low dose aspirin

# EULAR recommendations for the treatment of systemic sclerosis: 2023 Updated



**Figure 1** Schematic representation of the eight clinical domains covered by the 2023 recommendations. Note that severe prognosis is not represented. The different shades of green boxes labelled A–D represent the Strength of the Recommendation (SoR) as shown in the relative column of table 1. Dotted lines connect same drug or drug class across distinct clinical domains. CCB, calcium channel blocker; CYC, cyclophosphamide; ERAs, endothelin receptor antagonists; MMF, mycophenolate mofetil; MTX, methotrexate; PDE5i, phosphodiesterase five inhibitors; PPI, proton pump inhibitors; RITUX, rituximab; TCZ, tocilizumab.

IS (LoE)	RP	Digital ulcer
<b>CCB (Nifedipine)</b>	<b>1A</b>	
<b>PDE5i (sildenafil/ tadalafil)</b>	<b>1A</b>	<b>1A (healing/ prevention)</b>
<b>ERAs (Bosentan)</b>		<b>1A (prevention)</b>
<b>iv.Prostacyclin (Iloprost)</b>	<b>1A</b>	<b>1A (healing)</b>

## Raynaud's phenomenon

Calcium channel blockers and other vasodilators may be considered in management of SSc-RP 1B

Expert opinion suggests that PDE5i are effective as 2nd line agent for refractory SSc-RP 1B

For rescue therapy in severe SSc-RP IV prostanoids may be considered 1C

## Digital ulcers

Severe digital vasculopathy with new tissue necrosis or critical ischaemia requires urgent clinical assessment 1C

Sildenafil (or tadalafil) as 1st line agent in DU healing and secondary prevention and bosentan as second line treatment 1C

Expert opinion supports use of IV prostanoids in promoting DU healing 1C

Neurological co  
management w  
Peripheral Sens



MMF is recomm  
cyclophospham  
Consider adding  
disease 2C  
Nintedanib rec



PAH diagnosis b  
by a designated  
Anticoagulation



ACEi should be  
Glucocorticoid t  
Renal biopsy sh  
Referral for renal transplantation in cases without significant renal



recovery  
MMF is  
Antihist  
Options  
intense



Muscul  
immun



Severe  
requires  
Sildenafil  
prevent

Expert opinion supports use of IV prostanoids in promoting DU healing 1C

Fatigue and Health related Quality of Life

improve quality of

when investigation

be added if

lux 1C

1B

2B

th

Sexual dysfunction needs engagement of gynaecology,  
urology, and sexual health clinical services 1C

organ

C

-disciplinary

may be

s 2nd line

may be

ns

s major

impact on people with SSc

# Case 9

- 45F with SSc-ILD on MMF 2 g/day (6 mo)
- Progressive cough & DOE; afebrile; skin thickness much improved
- FVC ↓ 68% → 62%
- HRCT: increase reticulation, traction bronchiectasis and honey combing pattern

**Best add-on therapy?**

- A. Rituximab
- B. Tocilizumab
- C. Nintedanib
- D. Cyclophosphamide
- E. Sildenafil

# Thai recommendation of SSc-ILD 2564

## 1. การพิจารณาเริ่มรักษาด้วยยากดภูมิคุ้มกัน (immunosuppressive drugs)

ปัจจัยพิจารณาเริ่มการรักษาด้วยยา immunosuppressive drugs มีดังต่อไปนี้

1.1 ประเมินความรุนแรงของโรค โดยพิจารณาจากค่า FVC และปริมาณความผิดปกติจาก HRCT โดยใช้เกณฑ์ข้อใดข้อหนึ่งต่อไปนี้<sup>(3, 5, 13)</sup>

- FVC < 70% predicted
- ปริมาณความผิดปกติจาก HRCT > 20%

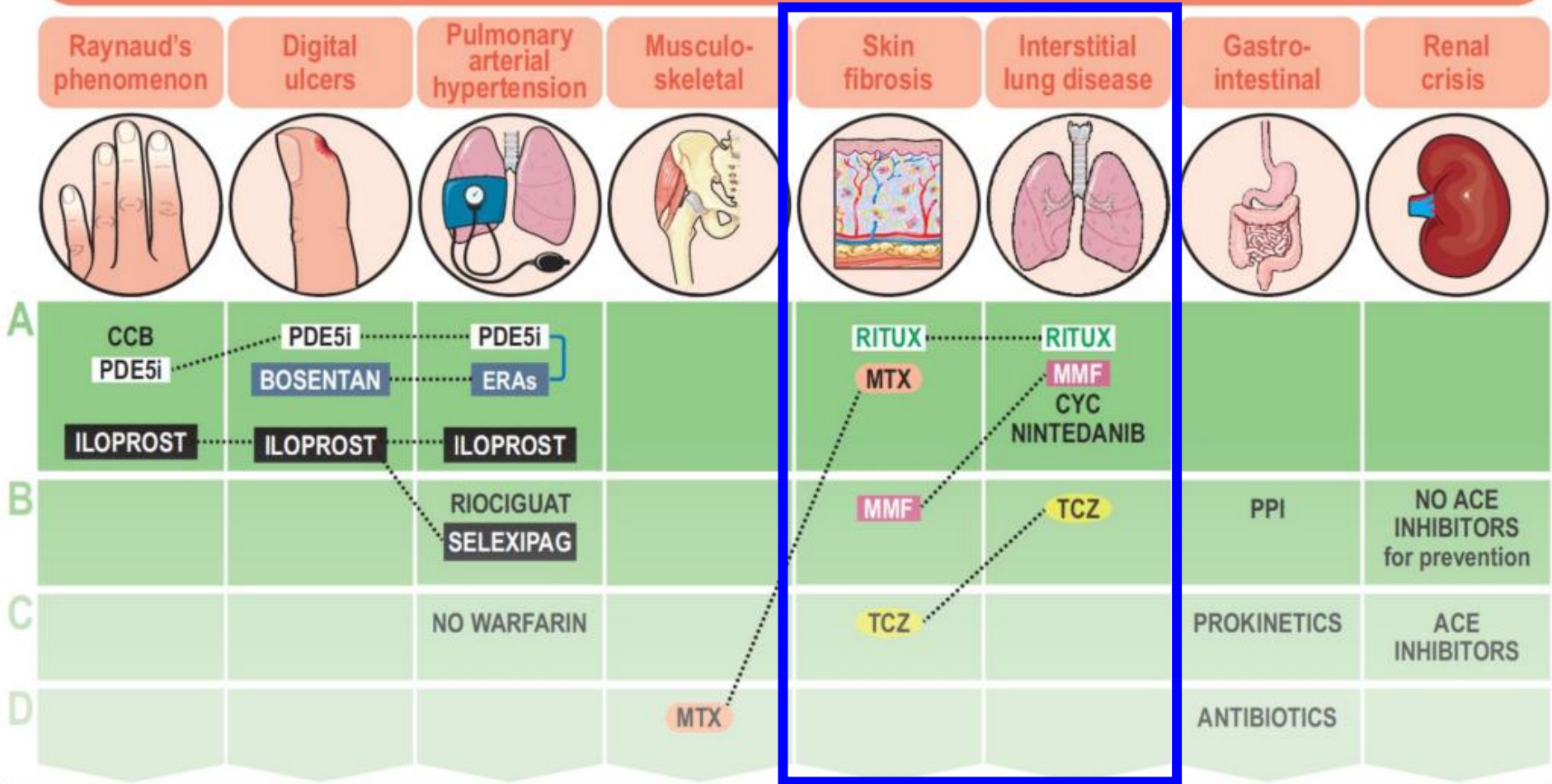
1.2 กรณีไม่เข้าเกณฑ์ข้อ 1.1 หรือไม่สามารถตรวจ pulmonary function tests ได้ ให้พิจารณาเริ่มการรักษาจากลักษณะต่อไปนี้

- ลักษณะทางคลินิกแย่งโดยไม่มีสาเหตุมาจากโรคหรือภาวะอื่น
- ผู้ป่วยที่มีปัจจัยเสี่ยงในการเกิดการลุกลามของโรค
- HRCT มีลักษณะที่เข้าได้กับ organizing pneumonia ร่วมด้วย (ตารางที่ 3) โดยไม่พบสาเหตุอื่นที่ทำให้มี HRCT เป็นแบบนี้ เช่น infection เป็นต้น
- มีข้อบ่งชี้ในการรักษาตามความผิดปกติในระบบอวัยวะอื่น

## Risk factors for disease progression in SSc-ILD

	SSc <sup>#</sup>
<b>Demographics</b>	<ul style="list-style-type: none"><li>• Older age</li><li>• Male sex</li><li>• African American ethnicity</li></ul>
<b>Circulating markers</b>	<ul style="list-style-type: none"><li>• Elevated ESR, CRP</li><li>• ATA-I <b>Anti-ScI-70</b></li></ul>
<b>Pulmonary function/markers</b>	<ul style="list-style-type: none"><li>• Baseline PFTs (FVC, <math>D_{LCO}</math>)</li></ul>
<b>Imaging/histology</b>	<ul style="list-style-type: none"><li>• Higher extent of ILD on HRCT</li></ul>
<b>Extrapulmonary involvement</b>	<ul style="list-style-type: none"><li>• Recent onset of SSc with rapid skin progression, more extensive skin fibrosis (mRSS)</li></ul>

# Systemic sclerosis



IS (LoE)	Skin	ILD
RTX	1A	1A
MMF	1B	1A
CYC	1B	1A
*TCZ	1B	1B
MTX	1B	
**Nintedanib		1A

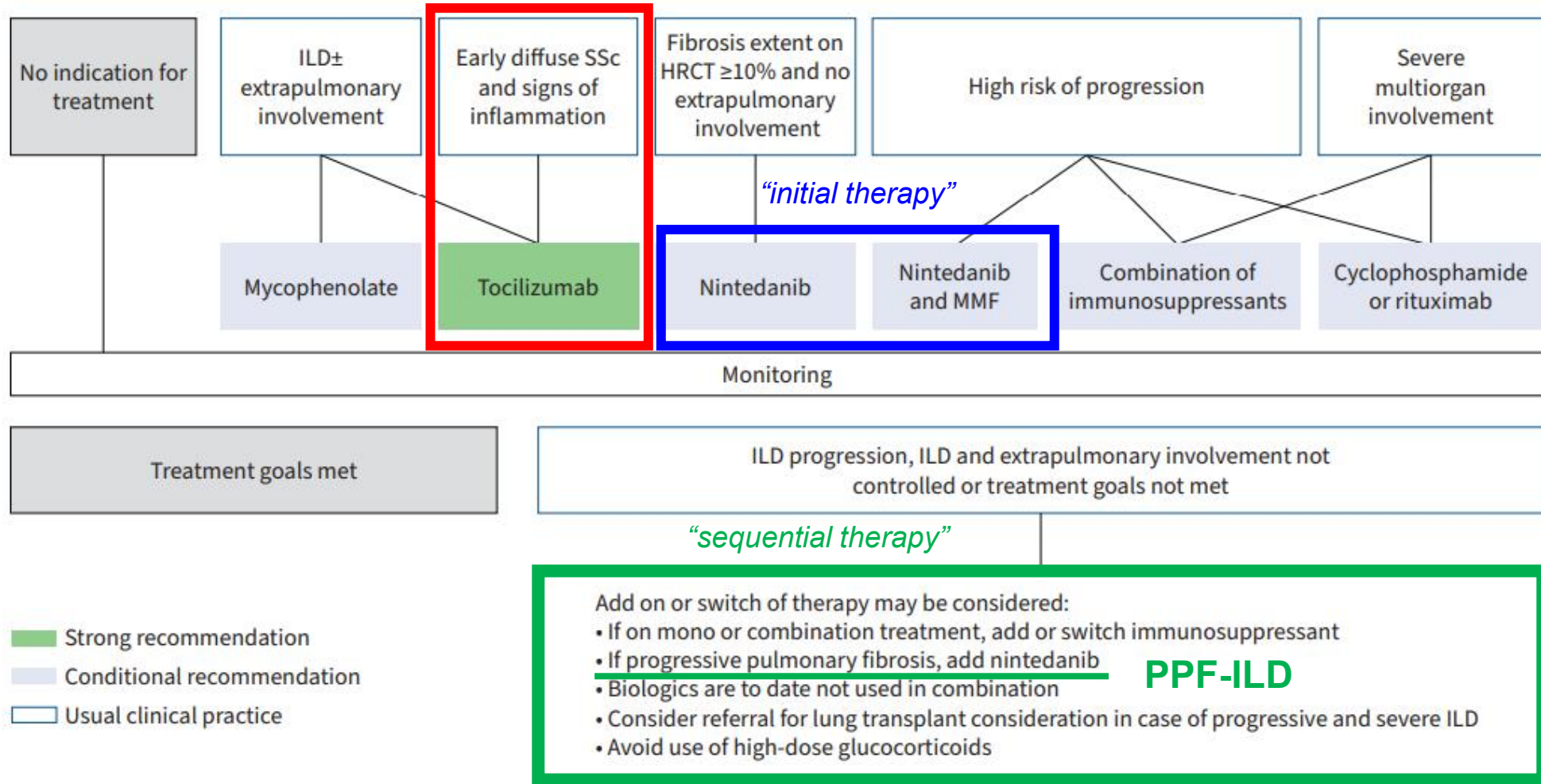
\*TCZ = early inflammatory (mild or subclinical) ILD

\*\*Nintedanib = Progressive Pulmonary Fibrosis (PF-ILD)

**Figure 1** Schematic representation of the eight clinical domains covered by the 2023 recommendations. Note that severe prognosis is not represented. The different shades of green boxes labelled A–D represent the Strength of the Recommendation (SoR) as shown in the relative column of table 1. Dotted lines connect same drug or drug class across distinct clinical domains. CCB, calcium channel blocker; CYC, cyclophosphamide; ERAs, endothelin receptor antagonists; MMF, mycophenolate mofetil; MTX, methotrexate; PDE5i, phosphodiesterase five inhibitors; PPI, proton pump inhibitors; RITUX, rituximab; TCZ, tocilizumab.

# 2025: ERS/EULAR guidelines for SSc-ILD

## Treatment of SSc-ILD



**FIGURE 4a** Treatment algorithms for patients with systemic sclerosis (SSc)-associated interstitial lung disease (ILD). HRCT: high resolution computed tomography; MMF: mycophenolate mofetil.

# แนวเวชปฏิบัติการวินิจฉัยและรักษาภาวะปอดเป็นพังผืดชนิดดุกตามในผู้ใหญ่ พ.ศ. 2567

เกณฑ์	เกณฑ์บังคับ	
การวินิจฉัย	1. ต้องได้รับการรักษาตามมาตรฐานของโรคหรือภาวะที่เป็นสาเหตุของ non-IPF fibrotic ILD แล้ว 2. ไม่มีโรคหรือภาวะอื่นที่เป็นสาเหตุของการแย่ง เช่น ปอดติดเชื้อ น้ำท่วมปอด ลมรั่วในช่องเยื่อหุ้มปอด ลิ้มเลือดอุดตันในหลอดเลือดแดงปอด และความดันหลอดเลือดแดงปอดสูง เป็นต้น	
	เกณฑ์ร่วม (ใช้เกณฑ์อย่างน้อย 2 ใน 3 ข้อ)	เกณฑ์เดี่ยว (ใช้เกณฑ์ทางสรีรวิทยาข้อเดียว)
1. ลักษณะทางคลินิก	อาการและอาการแสดงแย่ง*	-
2. ลักษณะทางสรีรวิทยา	ข้อใดข้อหนึ่งต่อไปนี้ • Absolute FVC decline $\geq$ 5% • Absolute DLCO decline $\geq$ 10%	Relative FVC decline $\geq$ 10%
3. ลักษณะทางรังสีวิทยา	ปริมาณหรือลักษณะที่บ่งถึงพยาธิสภาพพังผืด มากขึ้นหรือแย่งจากภาพเอกซเรย์คอมพิวเตอร์ปอดชนิดความละเอียดสูง**	-
ระยะเวลาการประเมิน	12 เดือน	24 เดือน

\* อาการและอาการแสดงแย่ง หมายถึง เหนื่อยมากขึ้น แนะนำให้ประเมินจากแบบประเมินอาการเหนื่อย เช่น modified Medical Research Council (mMRC) dyspnea scale เป็นต้น ไอมากขึ้น และระดับออกซิเจนปลายนิ้ว (pulse oximetry) ต่ำลง โดยประเมินทั้งขณะพักและหลังออกกำลังกายหรือมีกิจกรรม (exertional desaturation) เป็นต้น

\*\* ปริมาณหรือลักษณะที่บ่งถึงพยาธิสภาพพังผืดมากขึ้นหรือแย่งจากภาพเอกซเรย์คอมพิวเตอร์ปอดชนิดความละเอียดสูง ได้แก่ traction bronchiectasis หรือ bronchiolectasis ที่มีปริมาณมากขึ้น, ground-glass opacity ร่วมกับ traction bronchiectasis ที่เกิดขึ้นใหม่, fine reticulation ที่เกิดขึ้นใหม่, reticular abnormalities ที่มีการหนาตัวหรือมีปริมาณมากขึ้น, honeycombing ที่มีปริมาณมากขึ้นหรือเกิดขึ้นใหม่ และปริมาตรปอดที่ลดลง เป็นต้น

# Case 10.

- A 40-year-old woman with fever, dyspnea (4 weeks), SpO<sub>2</sub> 90%.
- Exam: Gottron's ulceration, no weakness
- CXR: bilateral ground-glass + consolidation.
- Sputum negative.
- CK 400, Anti-MDA5 positive.

**What is the best treatment in addition to high-dose steroids?**

- A. Rituximab**
- B. Mycophenolate**
- C. Cyclophosphamide**
- D. Cyclophosphamide + tacrolimus**
- E. IVIG**

# Autoantibody Phenotypes in IIM

Anti-p155/140 (TIF-1 $\gamma$ )



- Psoriasis-like lesions
- Palmar hyperkeratosis
- Hypopigmented and telangiectatic patches ("red on white")

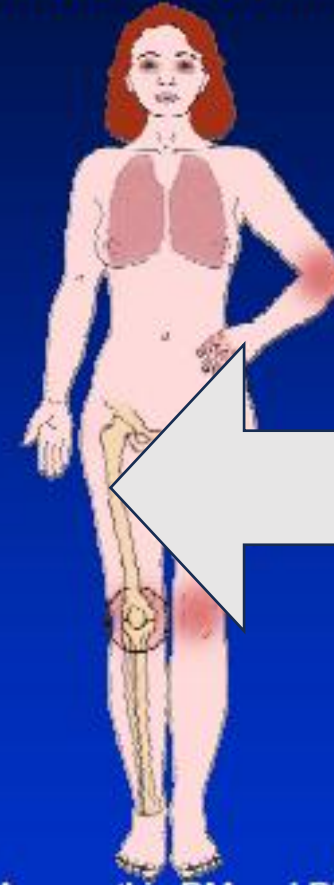
DM/JDM, CTM, CAM;  
Mod to severe weakness, ulcers,  
Erythroderma, V- and shawl sign,  
Edema, generalized LD

Anti-MJ (NXP-2)



DM/JDM;  
Calcinosis,  
Contractures,  
No trunk rash

Anti-CADM-140 (MDA-5)



- Skin ulceration
- Scalp ulceration with alopecia
- Mucosal ulcer
- Palmar papules (inverse Gottron' papules)

Amyopathic DM and DM;  
Rapidly progressive ILD,  
Classic DM rashes,  
Arthritis

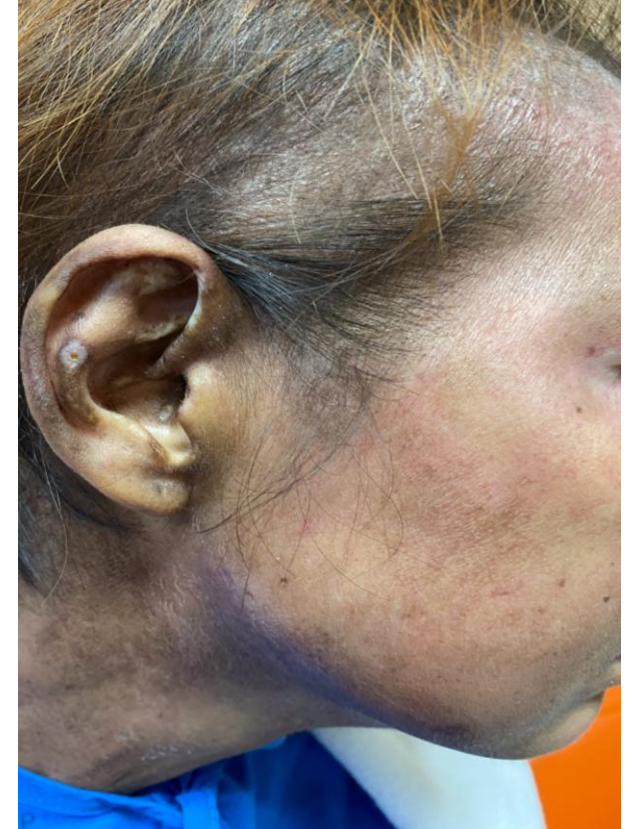
# Skin Manifestations of Anti-MDA-5 CADM



**Palmar papules  
(inverse Gottron' sign)**



**Cutaneous ulcerations**



# Autoantibody Phenotypes in IIM

## Anti-aminoacyl-tRNA Synthetases



Interstitial lung disease, arthritis,  
Fever, mechanic's hands  
PM > DM  
75% 5-year survival

## Anti-Signal Recognition Particle



Acute, severe muscle weakness in PM,  
Myalgias, cardiac involvement  
25% 5-year survival

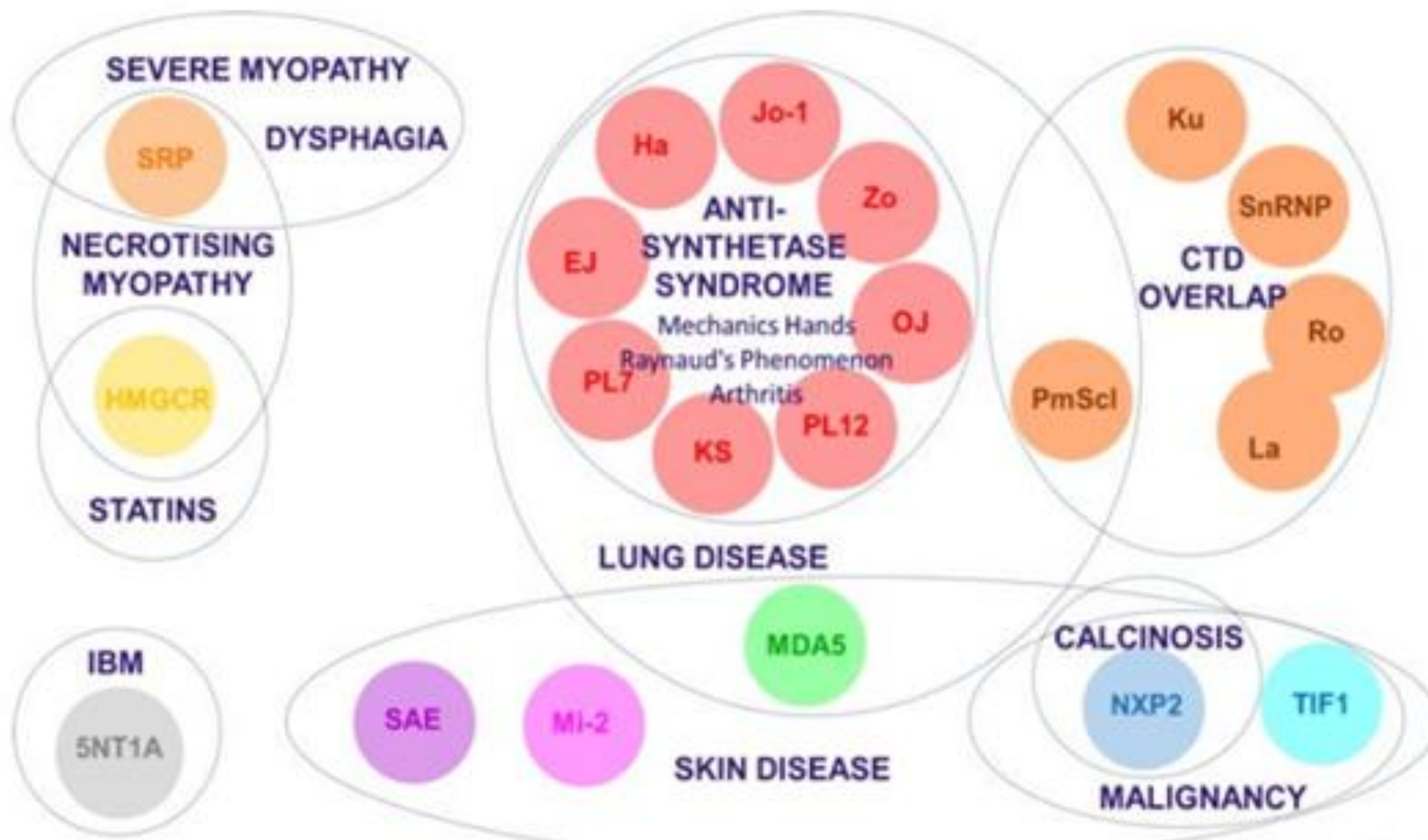
## Anti-Mi-2: Chromodomain Helicase DNA Binding Protein 4



Classic dermatomyositis, V-sign &  
shawl rashes, cuticular overgrowth  
100% 5-year survival

37  
Love et al. 1991 Medicine; Miller 1993 JAMA

# Myositis antibodies & Clinical associations



# 2023 ACR/CHEST Guideline

## Initial treatment options for the treatment of SARD-ILD

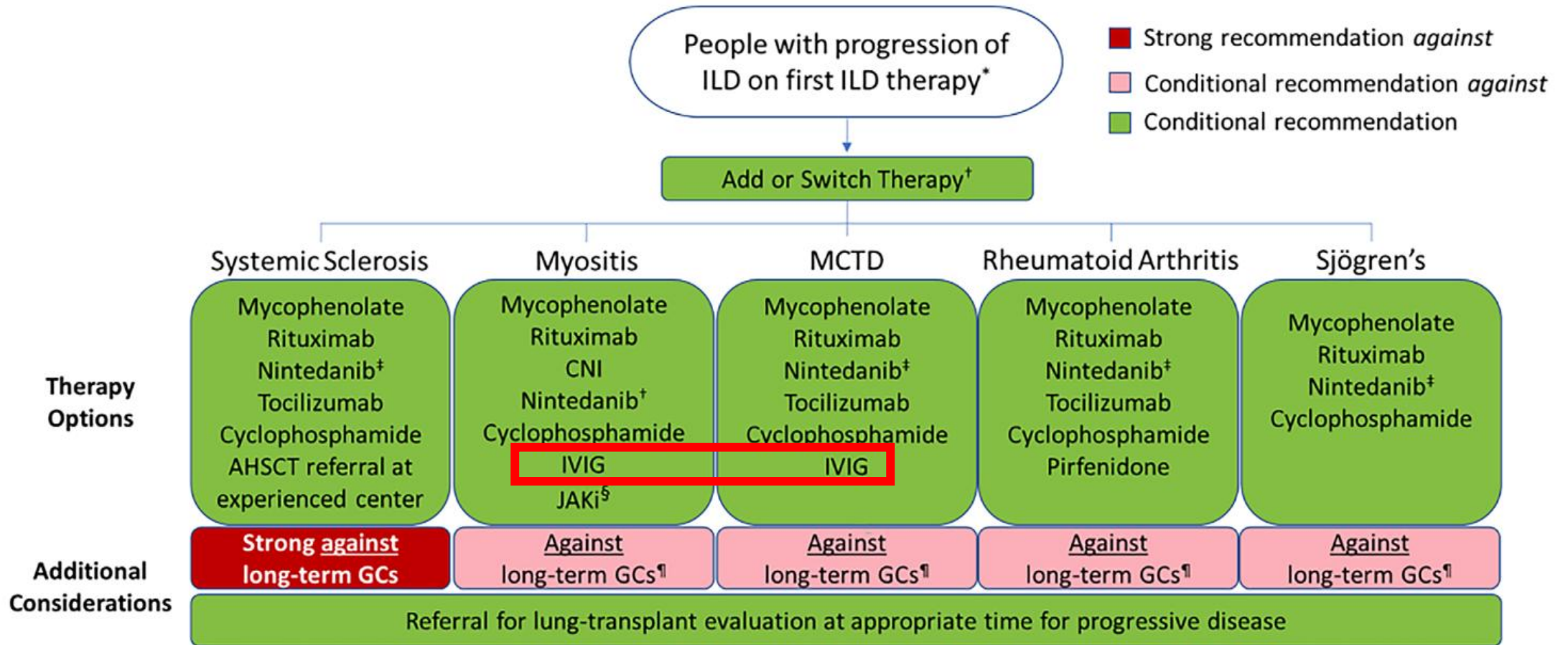
	Systemic Sclerosis	Myositis	MCTD	Rheumatoid Arthritis	Sjögren's
First-line ILD therapy	Preferred Mycophenolate <sup>†</sup> Tocilizumab Rituximab	Mycophenolate <sup>†</sup> Azathioprine Rituximab CNI	Mycophenolate <sup>†</sup> Azathioprine Rituximab	Mycophenolate <sup>†</sup> Azathioprine Rituximab	Mycophenolate <sup>†</sup> Azathioprine Rituximab
	Additional options Cyclophosphamide Nintedanib Azathioprine	JAKi Cyclophosphamide	Tocilizumab Cyclophosphamide	Cyclophosphamide	Cyclophosphamide
+ Glucocorticoids	Strong recommendation against GCs	Short-term GCs*	Short-term GCs*	Short-term GCs*	Short-term GCs*

■ Strong recommendation *against*    ■ Conditional recommendation

Calcineurin inhibitors (CNI) = Cyclosporin A, tacrolimus (most evidences from IIM-ASS)  
Janus kinase inhibitor (JAKi) = Tofacitinib (indirect evidence from anti-MDA-5 not RP-ILD)

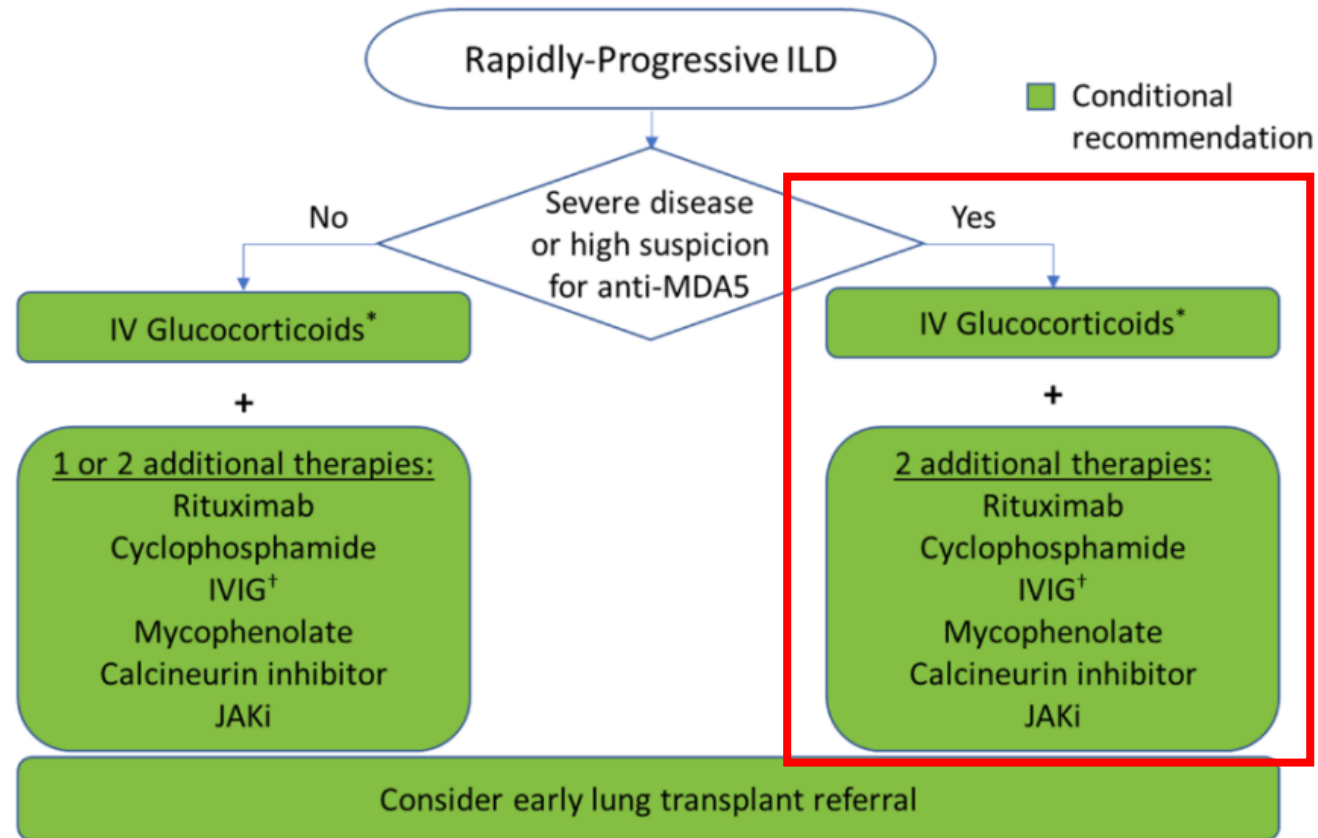
# 2023 ACR/CHEST Guideline

## Management of SARD-ILD with progression of ILD despite first ILD therapy.



1. IVIG is effective for **myositis and dysphagia IIM** (myositis predominant–MCTD).
2. IVIG may be useful when rapid onset of action is desired, eg, presence of **severe respiratory muscle weakness**.
3. IVIG may be used in the acute setting if infection is a concern, **BUT limited ILD efficacy data**.

# Guideline for the Treatment of RP-ILD



\* In rare patients with systemic sclerosis with rapidly progressive ILD, there was no consensus on whether or not to use glucocorticoids – if used, patients should be monitored closely for evidence of renal crisis.

† Rituximab and cyclophosphamide recommended over IVIG, but IVIG may be preferred if there is high concern for infection.

ILD = interstitial lung disease; IV = intravenous; IVIG = intravenous immune globulin; JAKi = janus kinase inhibitor

# Case 11.

- A 60-year-old man presents with 1 week of painful proximal weakness.
- Hx: gout, dyslipidemia.
- Meds: colchicine, statin.
- Recently took clarithromycin due to CAP.
- Exam: proximal weakness, ↓DTR
- CK 1500, ANA negative.

**What is the cause?**

- A. Inclusion body myositis
- B. Statin-induced myopathy
- C. Colchicine-induced myopathy
- D. Cancer-associated myopathy
- E. Infectious myopathy

# Drug induced myopathy

	Steroid	Anti-malarials	Colchicine	Statin
Onset	Subacute-chronic	Chronic	Acute-subacute	
Risk	<ul style="list-style-type: none"> <li>▪ Prednisolone &gt; 20 mg/day at least 2 WK</li> <li>▪ Cushingoid</li> </ul>	<ul style="list-style-type: none"> <li>▪ Duration &gt; 5 Y</li> <li>▪ Cardiomyopathy</li> <li>▪ Arrhythmia</li> <li>▪ Retinopathy</li> </ul>	<ul style="list-style-type: none"> <li>▪ CKD</li> <li>▪ Drug interaction with CYP3A4/P-glycoprotein inhibitors (<i>cyclosporine, clarithromycin, ketoconazole, itraconazole, ritonavir, diltiazem, verapamil, erythromycin, fluconazole</i>)</li> <li>▪ Concomitant drug (<i>statin, fibrate, or digoxin</i>)</li> </ul>	<ul style="list-style-type: none"> <li>▪ CKD</li> <li>▪ Drug interaction</li> <li>▪ Hypothyroid</li> <li>▪ Anti-HMGCR (ANA)</li> </ul>
Myopathy	<ul style="list-style-type: none"> <li>▪ Myopathic</li> </ul>	Myopathic and neuropathic pattern		Myopathic
CK	Normal	Abnormal		
Biopsy	<ul style="list-style-type: none"> <li>▪ Muscle fiber type 2 atrophy</li> <li>▪ No inflammation</li> </ul>	<ul style="list-style-type: none"> <li>▪ Muscle fiber hypertrophy</li> <li>▪ Vacuoles</li> <li>▪ Rarely inflammation</li> <li>▪ Curvilinear bodies (anti-malarials)</li> </ul>		<ul style="list-style-type: none"> <li>▪ Necrotizing myositis</li> <li>▪ Myocyte necrosis</li> <li>▪ Rarely inflammation (macrophage)</li> </ul>
Rx	<ul style="list-style-type: none"> <li>▪ Steroid discontinuation or tapering with steroid-sparing agents</li> </ul>	Anti-malarial/colchicine discontinuation		<ul style="list-style-type: none"> <li>▪ Statin discontinuation</li> <li>▪ Short course of steroid (Anti-HMGCR)</li> <li>▪ ±Re-challenging</li> </ul>

# Case 12.

- A 65-year-old man presents with acute painless vision loss (3 days).
- He had temporal headache, fever (1 month) and shoulder morning stiffness.
- Exam: temporal tenderness, pale swollen optic disc.
- He was on warfarin due to AF.
- ESR 90 mm/hr. CT brain normal.

**What is the most appropriated management in this patient?**

- A. Temporal artery biopsy**
- B. Temporal artery ultrasound**
- C. Oral high dose prednisolone**
- D. Pulse methylprednisolone**
- E. Tocilizumab**

<b>Features</b>	<b>Giant cell arteritis</b>	<b>Takayasu's arteritis</b>
<b>F:M; Age</b>	2:1; >50 years (Caucasian)	8:1; <40 years (Asian)
<b>Fever</b>	Common	Uncommon
<b>PMR</b>	Common	Rare
<b>Headache</b>	Common (temporal arteritis)	Rare
<b>AION/stroke (carotid and vertebrobasilar hypoperfusion)</b>	Common	Uncommon
<b>Claudication</b>	Jaw	Extremities
<b>Aorta and major branches</b>	Uncommon (LV-GCA)	Common
<b>Bruit, unequal BP, renovascular hypertension</b>	Rare	Common

# 2021 ACR/Vasculitis Foundation Guideline for Diagnostic Testing in GCA

- We conditionally recommend **temporal artery biopsy** over temporal artery ultrasound/MRI for establishing a diagnosis of GCA.
- We conditionally recommend an initial **unilateral** temporal artery biopsy over bilateral biopsies.
- We conditionally recommend a long segment temporal artery biopsy **specimen (>1 cm)**.
- We conditionally recommend obtaining a temporal artery biopsy specimen within **2 weeks of starting oral glucocorticoids**.
- For patients with newly diagnosed GCA, we conditionally recommend obtaining noninvasive vascular imaging to **evaluate large vessel involvement**.

# 2023 EULAR Recommendations for the Use of Imaging in Large Vessel Vasculitis in Clinical Practice

Suspected GCA (mural inflammation)	Ultrasound	CT	High-resolution MRI	FDG-PET
Cranial arteries	+ (temporal arteries)	-	+	+
Extracranial arteries	+ (axillary arteries)	+	+	+
Structural damage	+	+	+	+
Suspected relapse when ESR/CRP unreliable	+	-	+	+

**Conventional angiography is not recommended for the diagnosis of GCA or TAK**

**In patients with suspected GCA,**

- An early imaging test is recommended to support the clinical diagnosis of GCA, assuming high expertise and prompt availability of the imaging technique.

- **Imaging should not delay initiation of treatment (ideally within 72 hours of commencing GC therapy).**

## 2022 AMERICAN COLLEGE OF RHEUMATOLOGY / EULAR

CLASSIFICATION CRITERIA FOR **GIANT CELL ARTERITIS****CONSIDERATIONS WHEN APPLYING THESE CRITERIA**

- These classification criteria should be applied to classify the patient as having giant cell arteritis when a diagnosis of medium-vessel or large-vessel vasculitis has been made
- Alternate diagnoses mimicking vasculitis should be excluded prior to applying the criteria

**ABSOLUTE REQUIREMENT**

Age  $\geq$  50 years at time of diagnosis

**ADDITIONAL CLINICAL CRITERIA**

Morning stiffness in shoulders/neck	+2
Sudden visual loss	+3
Jaw or tongue claudication	+2
New temporal headache	+2
Scalp tenderness	+2
Abnormal examination of the temporal artery <sup>1</sup>	+2

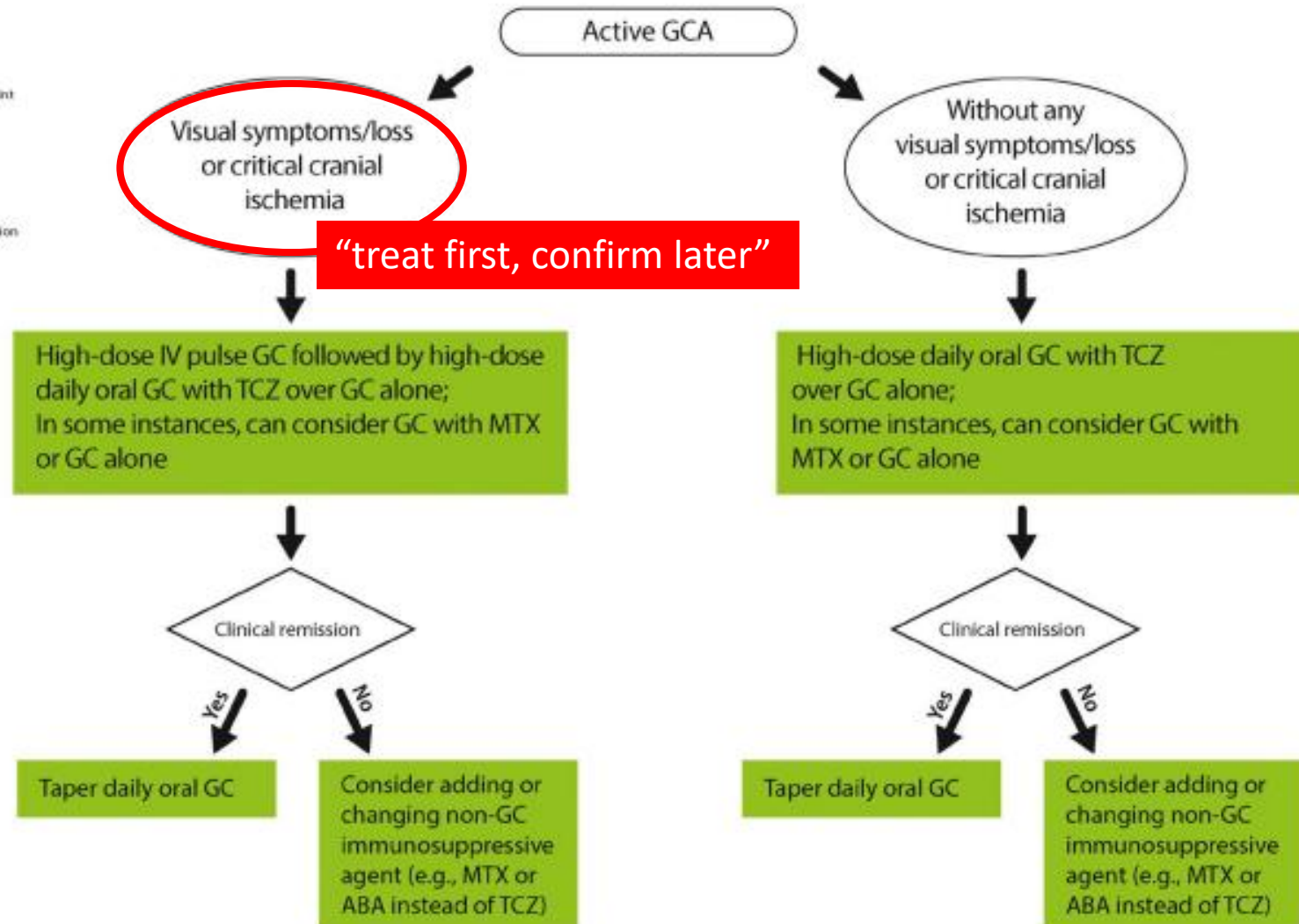
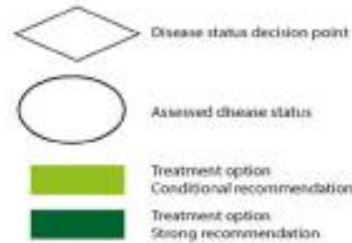
**LABORATORY, IMAGING, AND BIOPSY CRITERIA**

Maximum ESR $\geq$ 50 mm/hour or maximum CRP $\geq$ 10 mg/liter <sup>2</sup>	+3
Positive temporal artery biopsy or halo sign on temporal artery ultrasound <sup>3</sup>	+5
Bilateral axillary involvement <sup>4</sup>	+2
FDG-PET activity throughout aorta <sup>5</sup>	+2

**Sum the scores for 10 items, if present. A score of  $\geq$  6 points is needed for the classification of **GIANT CELL ARTERITIS**.**

# 2021 ACR/Vasculitis Foundation Guideline for GCA

## Overview of treatment of giant cell arteritis (GCA)



### IV pulse GCs

- IV methylprednisolone 500–1,000 mg/day (adults) or equivalent for 3–5 days

### High-dose oral GCs

- Prednisone 1 mg/kg/day up to 80 mg or equivalent

### Adjunctive treatment

- Statin in high CVD risk
- LD-ASA in critical or flow-limiting involvement of the vertebral or carotid arteries