

# Lupus Nephritis

## HISTOPATHOLOGY

Histopathologic findings in lupus nephritis are heterogenous and affect all compartments of the kidney. The site of deposition of immune complexes determines the histological pattern. To date, the ISN/RPS 2018 Classification is used to interpret the kidney biopsy based on glomerular features.

### CLASS I Minimal mesangial



- Normal by light microscopy [LM]
- Mesangial deposits on immunofluorescence [IF] or electron microscopy [EM]

### CLASS II Mesangial



- Pure mesangial hypercellularity with  $\geq 4$  nuclei surrounded by mesangium per mesangial area, away from the hilum
- Mesangial deposits in IF & EM

### CLASS III Focal proliferative



- Segmental or global endocapillary hypercellularity by LM in  $\geq 50\%$  of glomeruli

### CLASS IV Diffuse proliferative



- Segmental or global endocapillary hypercellularity by LM in  $\geq 50\%$  of glomeruli

### CLASS V Membranous



- Segmental to global subepithelial electron dense deposits
- If endocapillary hypercellularity is also present, an additional diagnosis of Class III or IV nephritis is made

### CLASS VI Sclerotic



- $> 90\%$  glomeruli globally sclerotic
- No evidence of endocapillary proliferation in the remaining glomeruli

### IMMUNOFLUORESCENCE

- IgG is essential for diagnosis and usually accompanied by IgA, IgM, C3 & C1q
- Kappa & Lambda stain equally
- Deposits can be seen in any compartment of the kidney
- Full house  
Ⓞ IgG, IgM, IgA, C3 & C1q

### ELECTRON MICROSCOPY

- More detailed information about location of immune deposits (Extraglomerular, subendothelial, subepithelial or mesangial deposits)
- Tubuloreticular inclusions

### TOP DIFFERENTIALS

- ✓ Type II mixed cryoglobulinemia
- ✓ MPGN with immune deposits
- ✓ Membranous nephropathy
- ✓ Post-infectious glomerulonephritis
- ✓ ANCA-associated glomerulonephritis

### 2018 MODIFIED NIH ACTIVITY AND CHRONICITY INDEX<sup>1</sup>

This standardized scoring system is used to evaluate for active and chronic lesions of lupus nephritis to help monitor for response to treatment, disease progression and prognosis.

#### ACTIVITY INDEX

Feature	Score
Endocapillary hypercellularity	0-3
Neutrophils/ karyorrhexis	0-3
Hyaline deposits/ wire loops	0-3
Fibrinoid necrosis	(0-3)x2
Cellular or fibrocellular crescents	(0-3)x2
Interstitial inflammation	0-3
<b>TOTAL</b>	<b>0-24</b>

#### CHRONICITY INDEX

Feature	Score
Global glomerulosclerosis	0-3
Fibrous crescents	0-3
Tubular atrophy	0-3
Interstitial fibrosis	0-3
<b>TOTAL</b>	<b>0-12</b>

SCORING 1 < 25% 2 25-50% 3 > 50%

### LOOK OUT FOR

- ☀ HIV Associated Immune Complex Kidney Disease
- 🦋 Lupus Podocytopathy
- ☹ Thrombotic Microangiopathy & other Vascular Lesions
- ⊖ Drug Toxicity [including hydroxychloroquine induced Pseudo-Fabry's disease]
- 🦠 Superimposed ANCA associated vasculitis

1. Sedoru M et al. Revision of the International Society of Nephrology-Renal Pathology Society classification for lupus nephritis: clarification of definitions, and modified National Institutes of Health activity and chronicity indices. Kidney Int. 2019 Apr;195(4):769-784. doi: 10.1016/j.kint.2017.12.022. Epub 2018 Feb 14. PMID: 29429925

# Lupus Nephritis

**JR Anderson**



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