

# **Nephrotic Syndrome**

## **Background**

### 1. Definition

- Characterized by massive proteinuria and lipiduria with varying degrees of edema, hypoalbuminemia, and hyperlipidemia

## **Pathophysiology**

### 1. Pathology of disease

- Increased permeability of glomerular membrane → leakage of large proteins in urine → hypoalbuminemia → edema (due to low plasma oncotic pressure)

### 2. Primary glomerular dz (90%)

- Focal segmental glomerulosclerosis
  - 35% of primary cases
  - Most common in black pts
- Membranous glomerular dz
  - 33% of primary primary cases
  - Most common in white pts
- Minimal change glomerular dz
  - 15% of primary cases
- Membranoproliferative glomerular dz (IgA nephropathy)
  - 14% pf primary cases

### 3. Secondary glomerular dz (10%)

- Diabetic nephropathy
  - MC secondary cause
- Amyloidosis
- SLE
- Myeloma
- Drugs
  - Gold, NSAIDs, penicillamine
- Infections
  - HIV, hepatitis, mycoplasma

### 4. Incidence

- 3 per 100,000

### 5. Risk factors

- Dependent on underlying dz

### 6. Morbidity, mortality

- Variable

### 7. Complications

- Renal failure
- Hyperlipidemia
- Hypercoagulable state (DVT)

## **Diagnosis**

### 1. History

- Focus on identifying a cause
- Fatigue, shortness of breath

## 2. Physical exam

- Edema
  - Dependent or generalized
- JVP increased
- Look for signs of DV
  - Calf tenderness, swelling

## 3. Labs

- Complete blood count
- Urinalysis
  - Proteinuria
  - Hematuria
  - Exclude UTI
- Protein
  - Decreased
    - Total protein <6g/dL [60 g/L]
    - Albumin <3g/dL [30 g/L]
  - Quantification of proteinuria
    - Albumin to creatinine ratio
    - 24 hr urine for proteinuria
- Comprehensive metabolic panel
  - Decreased serum albumin
  - Plasma glucose
  - Estimated GFR
  - Elevated BUN/Cr
- Coagulation profile
- Complement titers
  - Hepatitis panel
  - HIV
- Look for systemic dz
  - ESR
  - C-reactive protein
  - Immunoglobulins
  - ANA

## 4. Dx imaging

- Renal/abd ultrasound
  - Kidney size
  - Ascites

## 5. Other studies

- Kidney biopsy
  - Look for underlying dz
- Venous doppler
  - If DVT suspected

## Differential Diagnosis

1. Congestive heart failure
2. Liver failure / cirrhosis
3. Protein losing enteropathies
4. Myxedema
5. Renal failure

## **Treatment**

1. Best managed by or in consultation with a nephrologist
2. Underlying dz
  - Primary glomerular dz are best managed by or in consultation with a nephrologist
    - Pt may need steroids and/or immunosuppressants
  - Edema
    - Loop diuretics
    - Thiazide and spironolactone may need to be added
  - Proteinuria
    - ACEi + ARB
  - Hyperlipidemia
    - Statins
  - Hypercoagulability
    - Anticoagulation controversial in absence of DVT
3. Long-term care
  - Diet
    - Normal protein intake recommended
  - Renal failure
    - Best managed by or in consultation with nephrologist
  - Bone loss
    - Bone density testing and treatment of osteoporosis if needed

## **Follow-Up**

1. Nephrology consult
2. In severe cases w/ hypotension, azotemia, nausea & vomiting, admit for stabilization

## **Prognosis**

1. Variable, depends on underlying etiology

## **Prevention**

1. If systemic dz process, controlling the dz may slow down nephrotic syndrome progression

## **Patient Education**

1. <http://www.patient.co.uk/showdoc/27000748/>

## **References**

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2. Hull RP, Goldsmith DJ. Nephrotic syndrome in adults. BMJ. 2008 May 24;336(7654):1185-9.
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5. Wheeler DC, Bernard DB. Lipid abnormalities in the nephrotic syndrome: cause, consequences and treatment. Am J Kid Dis 1994;23:331-46.

**Evidence-Based Inquiry**

1. What is the differential diagnosis of chronic leg edema in primary care?

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