



Clínica  
Universidad  
de Navarra

# **SEMINARIO DE CASOS: DIAGNÓSTICOS AL LÍMITE DE LO INVISIBLE**



## **VALORACIÓN POR BIOPSIA RENAL DE POSIBLE DONANTE VIVA DE RIÑÓN CON HEMATURIA PERSISTENTE**

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# HISTORIA CLÍNICA

- Mujer de 46 años.
- Estudio para donación de trasplante renal (marido).

- **Antecedentes personales:**

- Dos episodios de cólicos nefríticos en riñón izquierdo que fue tratado con litotricia (se encuentra asintomática desde entonces).
- Infección del tracto urinario.
- Hepatitis por VHB en 2000 (core +, carga viral -).

- **Antecedentes familiares:**

- Padre fallecido de Ca de pulmón a los 72 años, **sin alteraciones renales.**
- Madre de 71 años con bocio, **no alteraciones renales.**
- 3 hermanos sanos.

# Analítica

- Ligera hipercalciuria normocalcémica
- Perfil tiroideo: normal. Proteinograma normal.
- GFR normal(114ml/min), creatinina 0,6 mg/dL e ionograma normal.
- ANAs y ANCA's negativos. C3 normal con C4 bajo.
- Antígenos tumorales normales (CA-125, CA19,9, CEA)

## Anormales y Sedimento :

- Ligera **Hematuria dismórfica persistente**
- Ligera proteinuria de 414 mg/24 h, sin albuminuria (alb/cr<30 mg)

- Ecografía renal: normal, con **2 quistes** en riñón derecho.
- Examen ginecológico normal

## **Table 3 | Causes of persistent microscopic hematuria**

*(A) Glomerular bleeding (common causes, not associated with proteinuria or casts)*

1. TBMN
2. AS (early stage) or carrier state
3. IgAN

*(B) Extraglomerular bleeding*

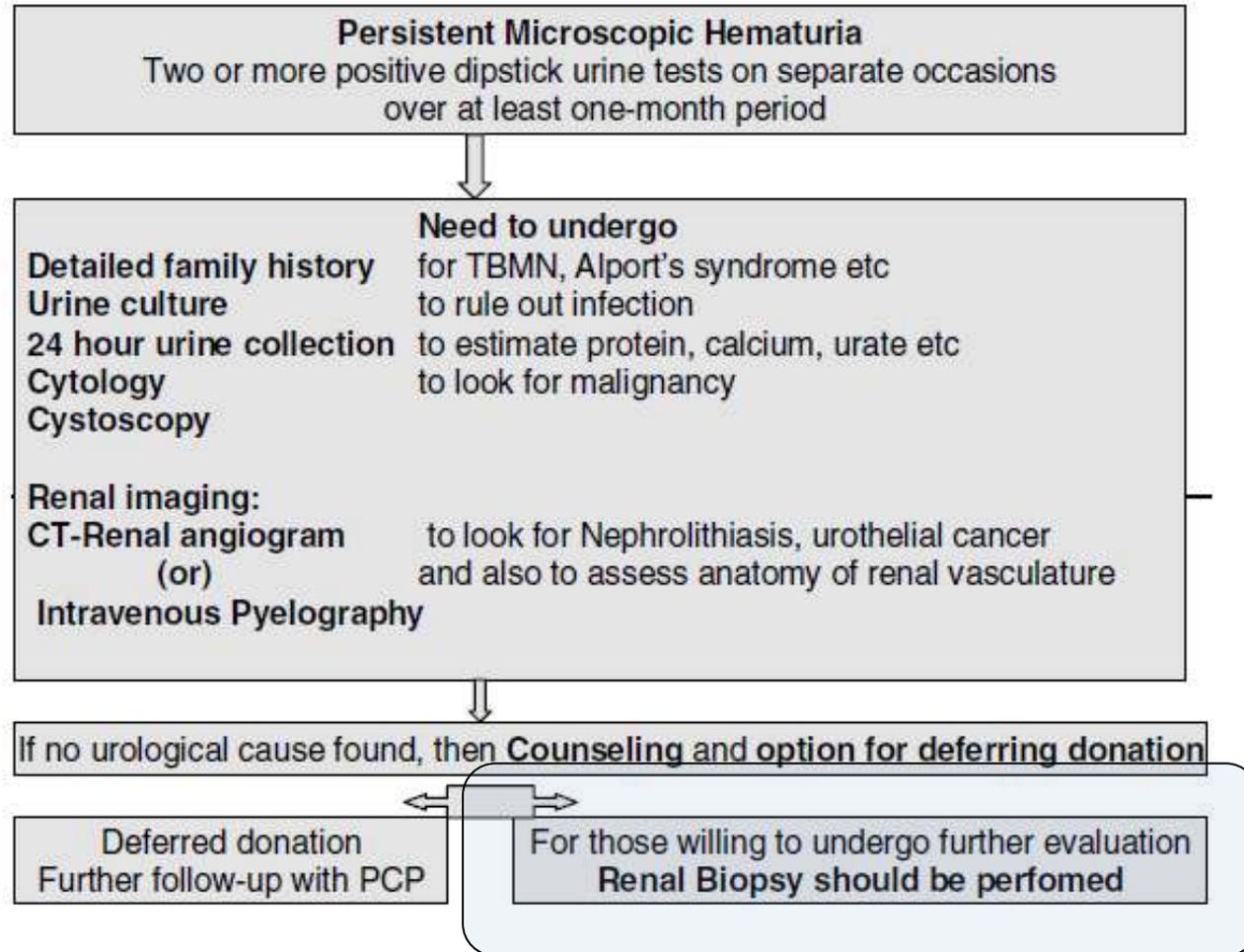
1. Stone disease
2. Hemoglobinopathy (SS/SA hemoglobin)
2. Polycystic kidney disease
3. Benign prostatic hyperplasia (elderly donors)
4. Malignancy (bladder, kidney, prostate)
5. Arteriovenous malformations and fistulas
6. Schistosomiasis (in endemic areas)
7. Hypercalciuria, hyperuricosuria, etc.

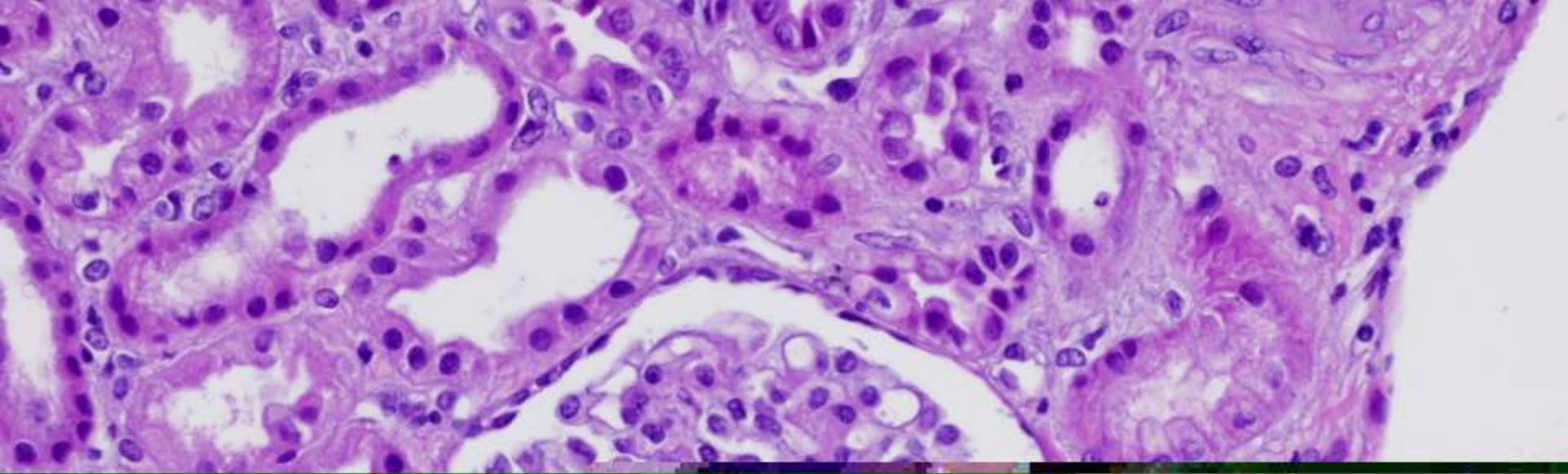
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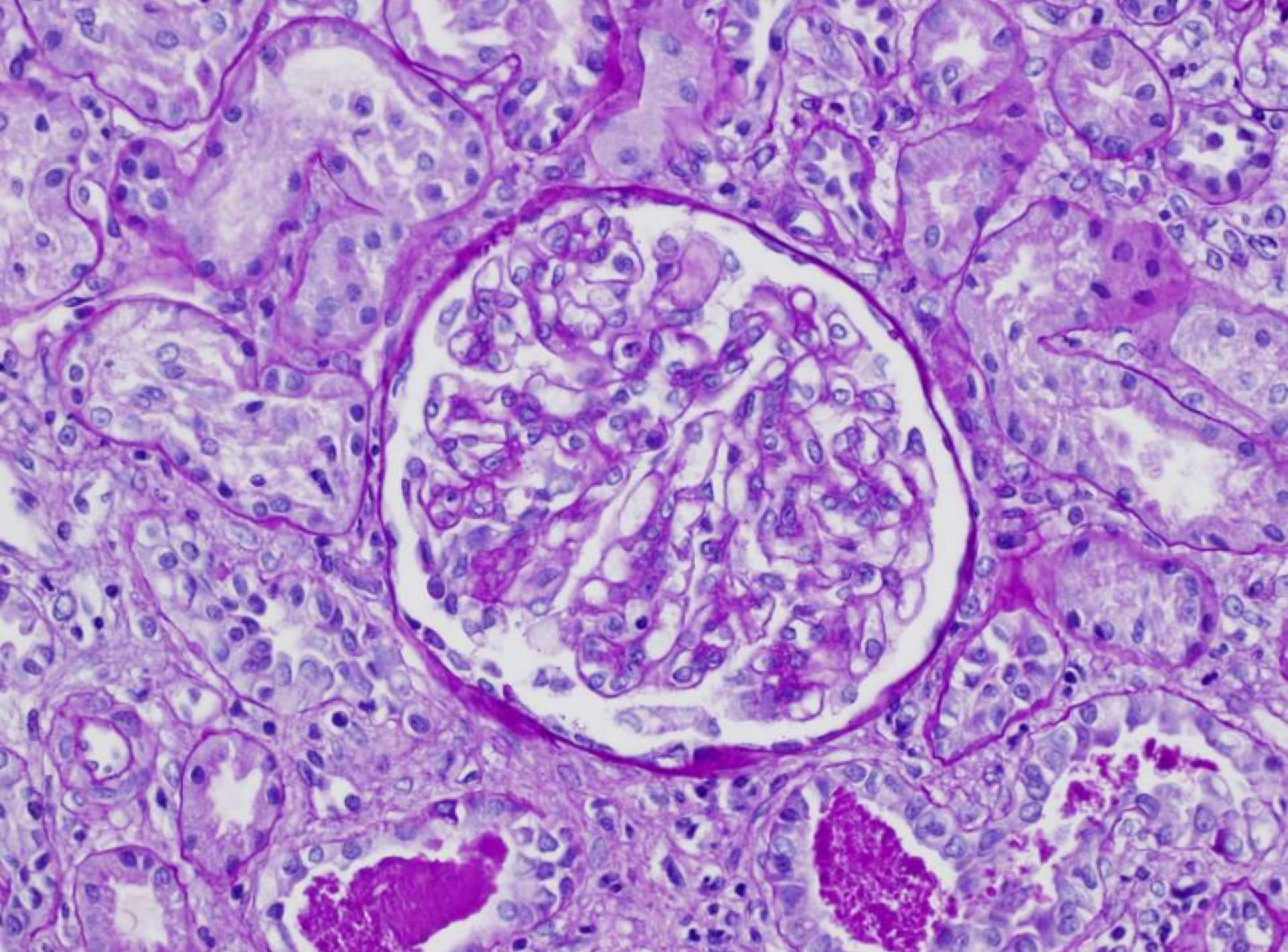
AS, Alport's syndrome; IgAN, IgA nephropathy; TBMN, thin basement membrane nephropathy.

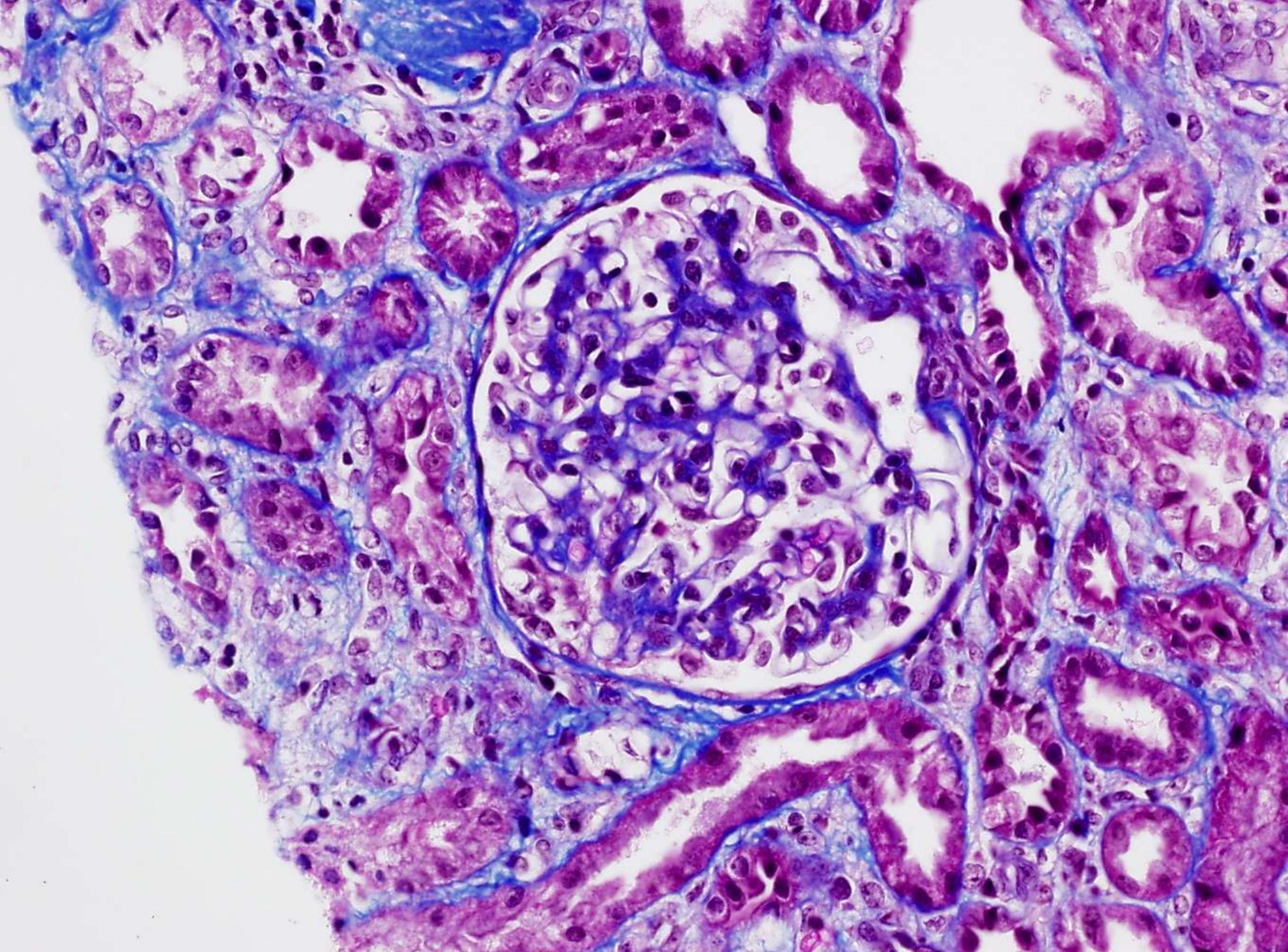
# ¿CONTRAINDICA LA HEMATURIA LA DONACIÓN DEL RIÑÓN?

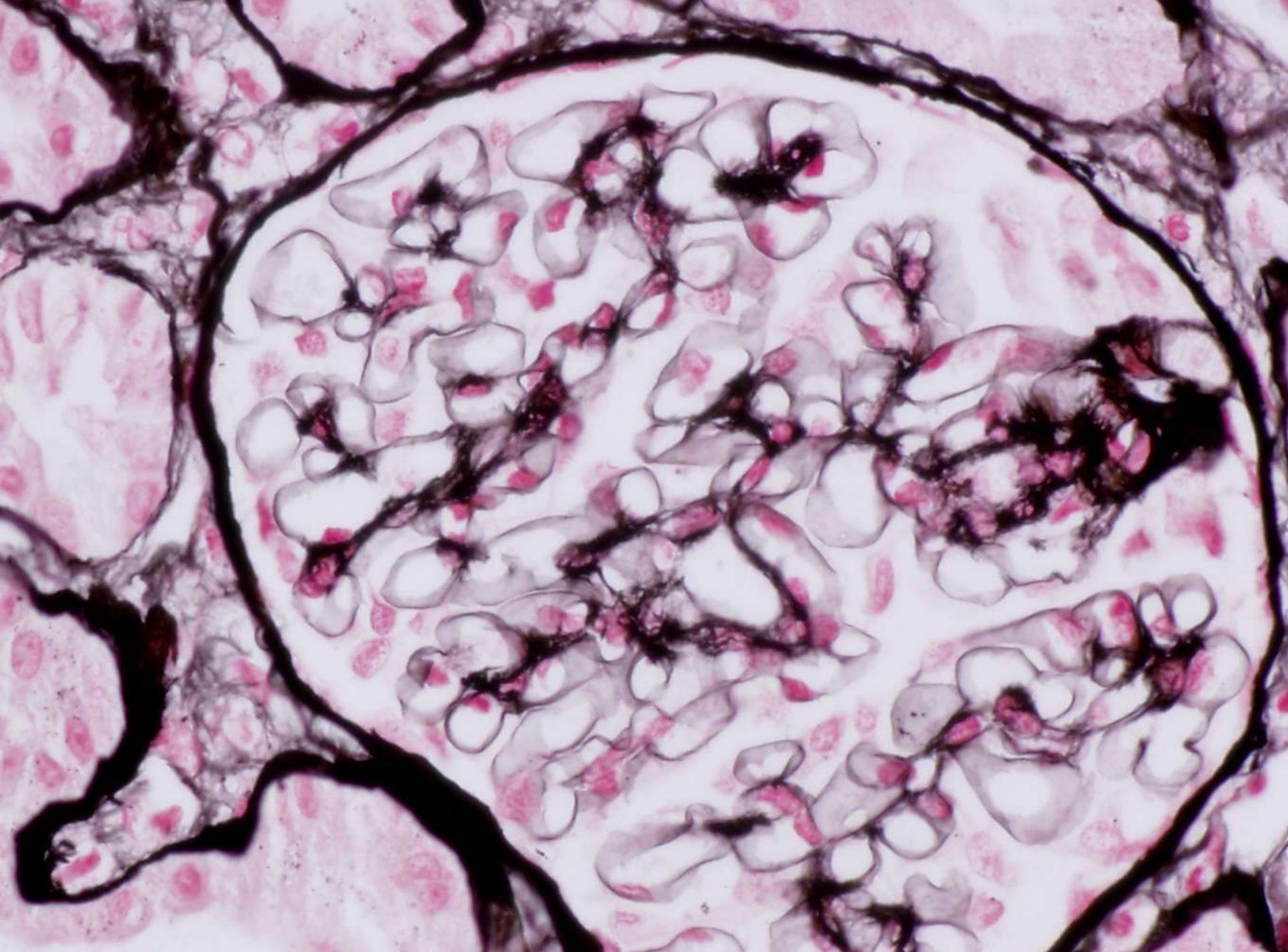
Table 2 | Algorithm to investigate microscopic hematuria in donors

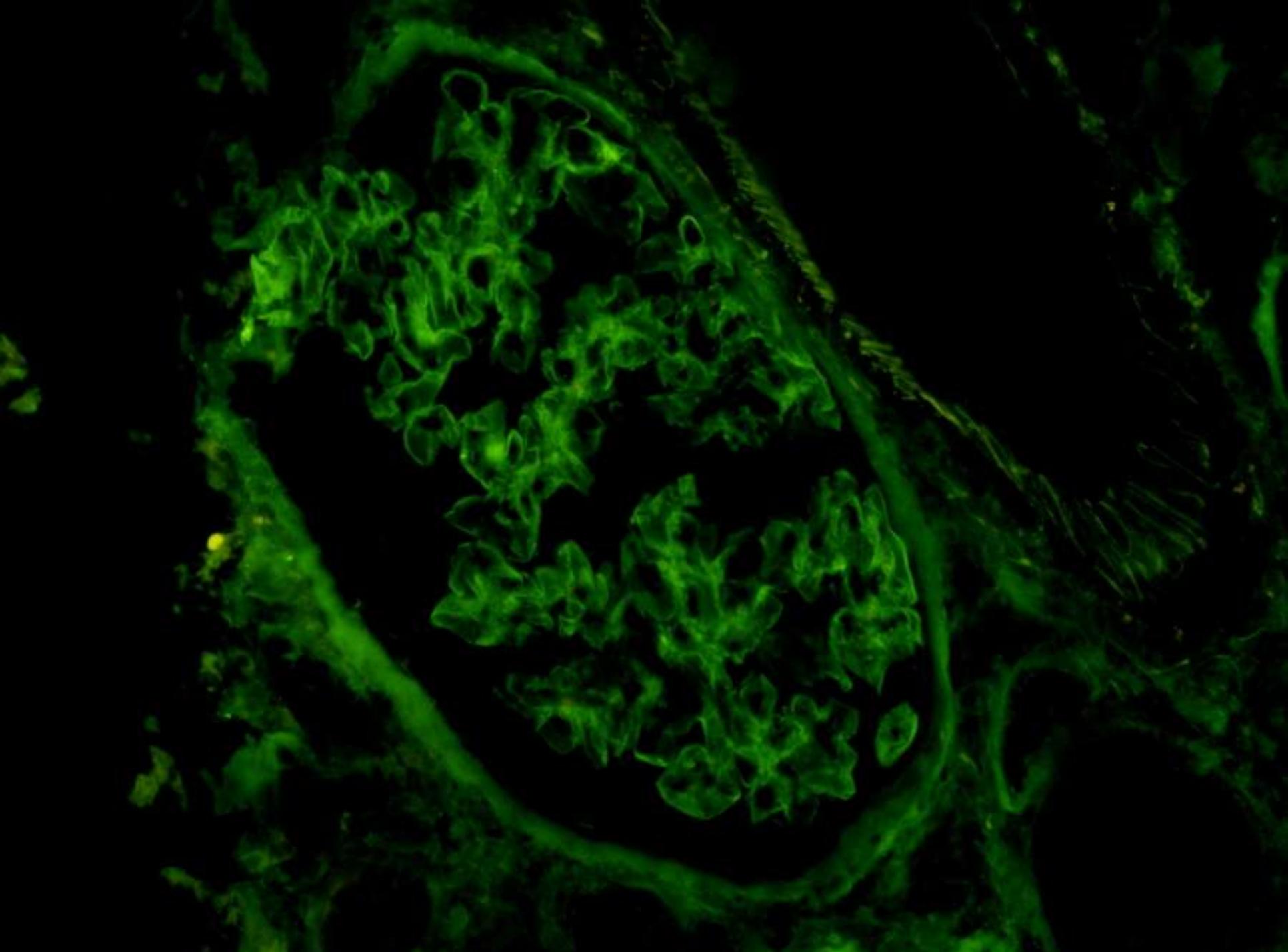


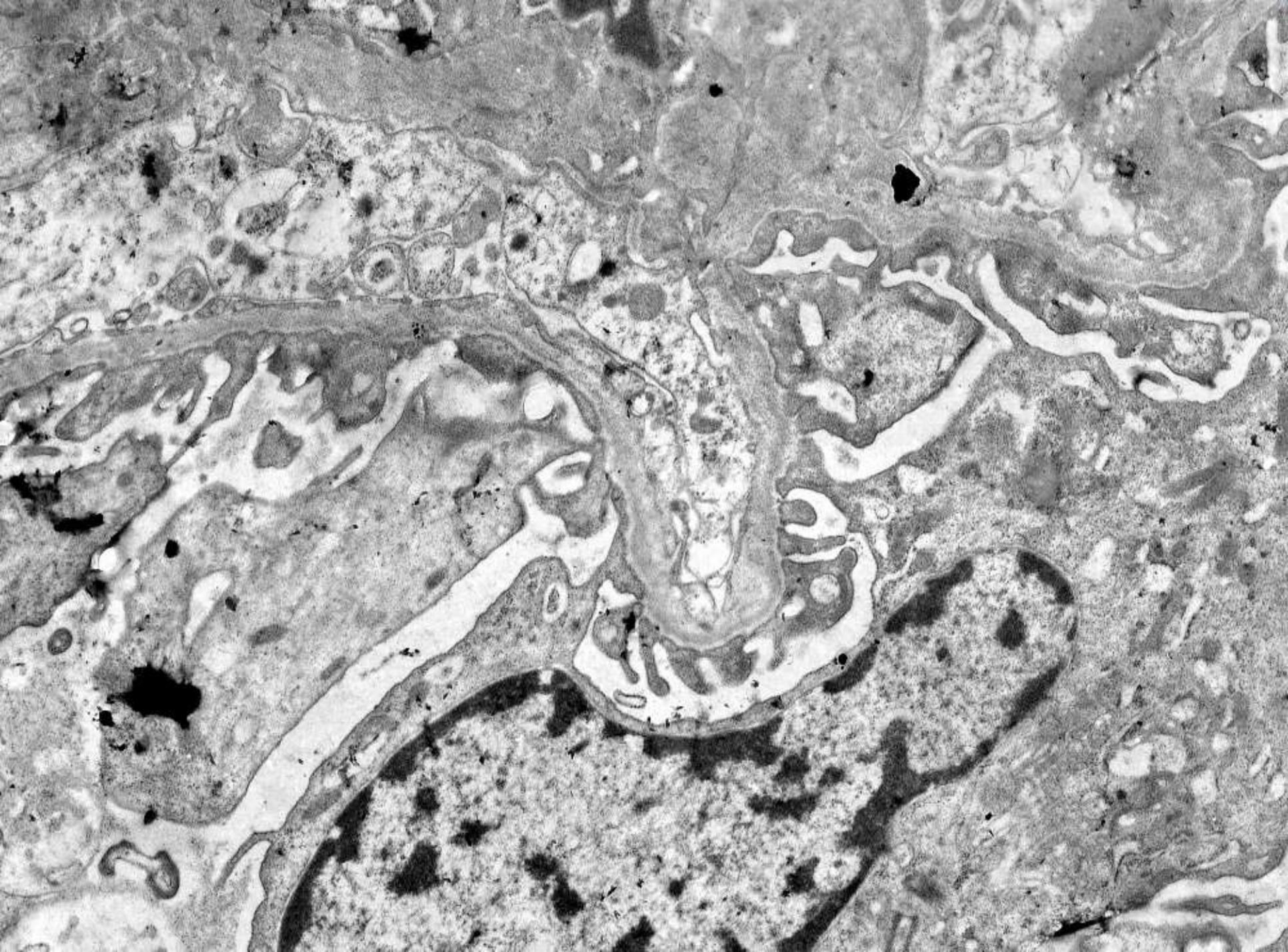


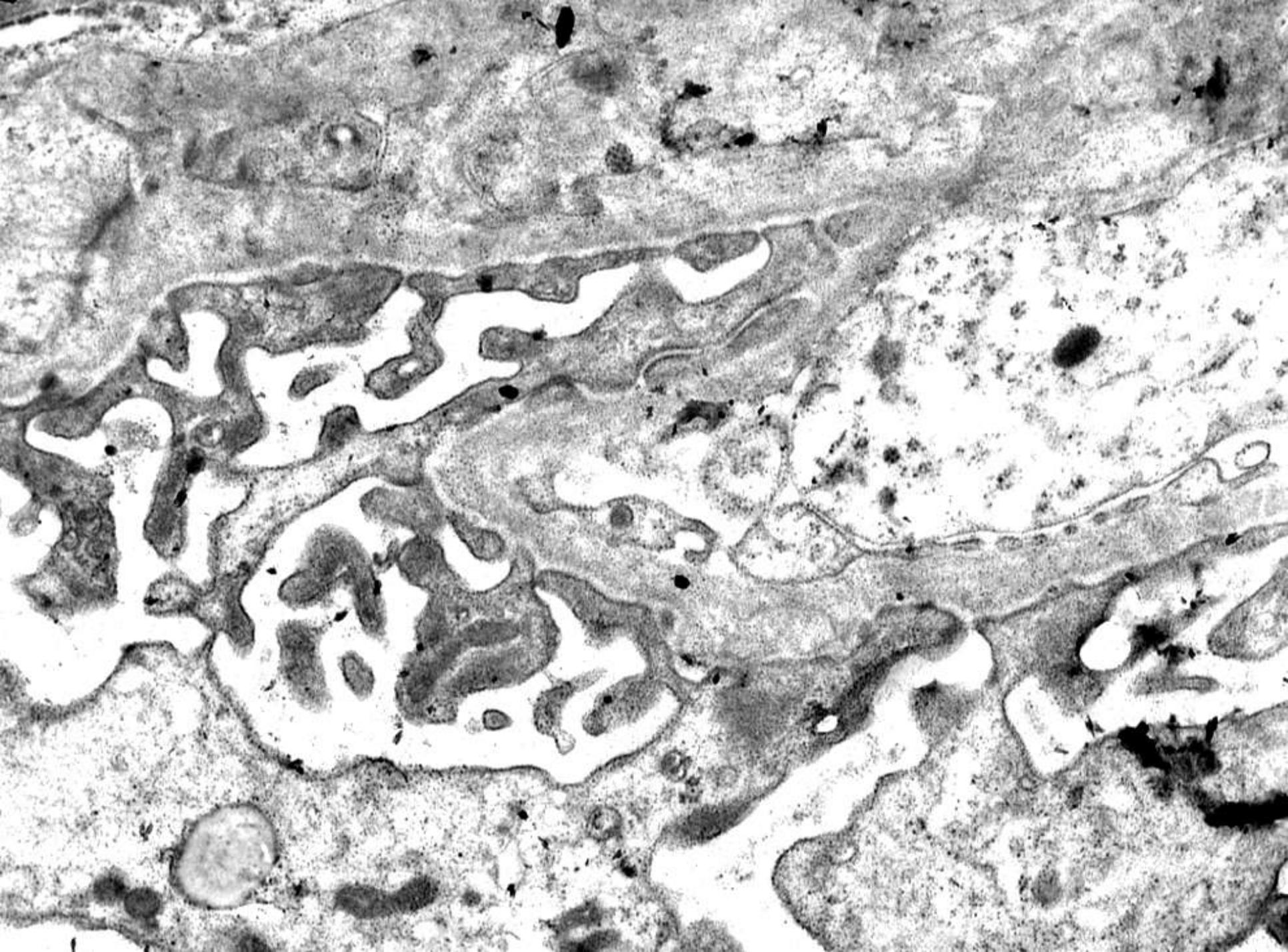


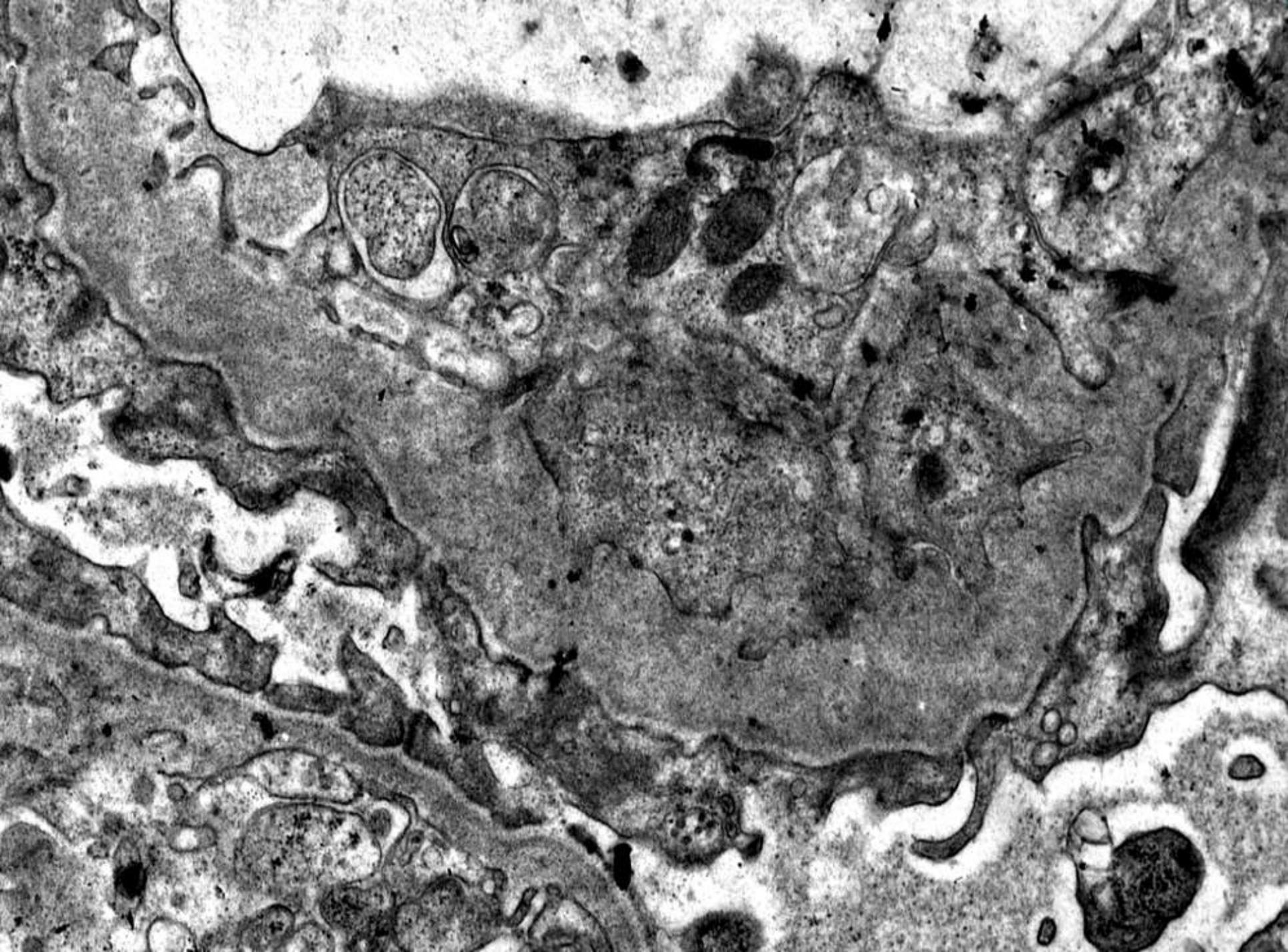












¿Está indicada la donación del riñón  
para trasplante?

## Persistent Glomerular Hematuria in Living Kidney Donors Confers a Risk of Progressive Kidney Disease in Donors After Heminephrectomy

In parallel with the global increase in the prevalence of end-stage kidney disease (ESKD) (1), the number of living donor kidney transplantations for ESKD has steadily increased (2). Because living kidney donation is justified on the premise of acceptable safety (3), the risks involved in heminephrectomy should be clearly determined. Although long-term (up after donation, which may progress to ESKD (7–11). Nevertheless, factors imposing a postdonation risk on donors, including kidney disease, have been poorly investigated

Here, to clarify whether hematuria is a risk factor for progressive kidney disease after living kidney donation, we investigated the prevalence of hematuria in donors before and after donation. We then checked the association between hematuria and signs of progressive kidney disease, such as the development of persistent proteinuria or decline in renal function in living kidney donors.

-242 donantes  
vivos  
-8,3% presentan  
**hematuria**  
**persistente** antes  
del trasplante

**Table 2:** Association between baseline characteristics and persistent proteinuria after donation in 242 living kidney donors

Variable	Proteinuria after donation		p-Value
	No or occasional (n = 222)	Persistent (n = 20)	
Male, %	40.0	45.0	0.67
Age, years	57 (49 to 64)	61 (54 to 69)	0.15
<b>Variables estudiadas con resultados significativos</b>		<b>Valor p</b>	
Aumento de la presión arterial		<0,01	11
-Sistólica			32
-Diastólica			00
			02
			001
			02
			07
			96
			66
			001
			16
-Tratamiento con antihipertensivos		0,07	48
			001
			01
-Proteinuria ocasional con moldes		<0.01	00
			42
-HEMATURIA DISMÓRFICA PERSISTENTE		<0,01	37
			49
			38
			46
Hemoglobin, g/dL	13.5 (12.8 to 14.3)	13.6 (12.2 to 14.6)	0.73
Uric acid, mg/dL	4.8 (4.1 to 5.9)	4.9 (4.0 to 6.0)	0.74
Total cholesterol, mg/dL	202 (178 to 232)	209 (171 to 221)	0.78
Triglyceride, mg/dL	111 (76 to 177)	118 (85 to 155)	0.66
Time since donation, month	27 (15 to 48)	35 (18 to 47)	0.37

Variables were expressed as median (interquartile range) or percentage.

CKD = chronic kidney disease; eGFR = estimated glomerular filtration rate.

# La hematuria microscopica persistente después de la donación se asocia a menor GFR

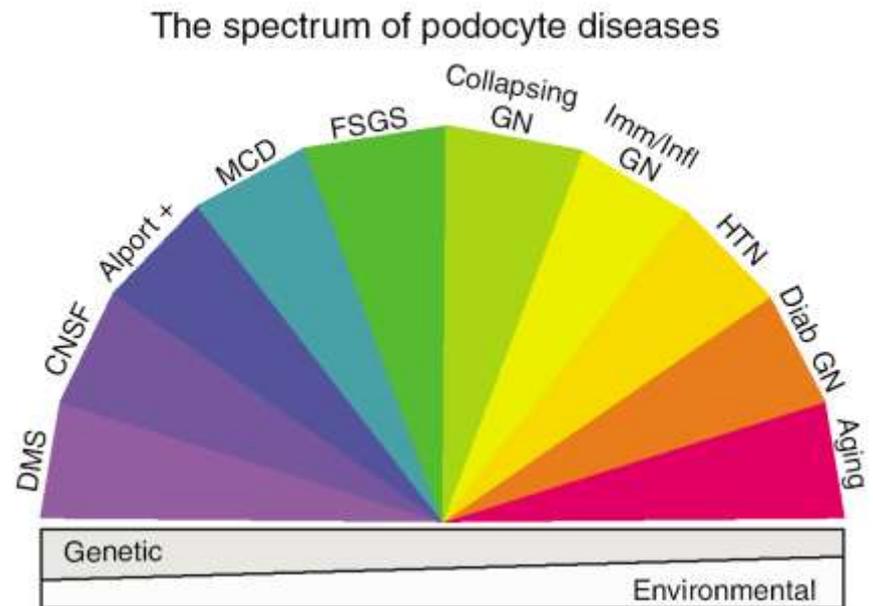
**Table 4:** Association between yearly percent changes in glomerular filtration rate from 1 year after donation and clinical characteristics in 163 living kidney donors

Variable	Univariate analysis		Multivariate analysis <sup>1</sup>	
	β (95%CI)	p-Value	β (95%CI)	p-Value
Predonation characteristic				
Body-mass index, kg/m <sup>2</sup> , 1 unit	0.26 (0.11 to 0.42)	0.001	0.20 (0.02 to 0.38)	0.03
eGFR, mL/min/1.73m <sup>2</sup> , 1 unit	0.03 (−0.01 to 0.06)	0.12	0.04 (0.01 to 0.07)	0.04
Proteinuria by dipstick (vs. negative test)				
Occasional	−0.01 (−2.49 to 2.48)	1.00	−0.27 (−2.79 to 2.24)	0.83
Postdonation characteristic				
Hematuria (vs. negative test)				
Occasional	0.23 (−1.56 to 2.03)	0.80	0.29 (−1.50 to 2.08)	0.75
Persistent without d-RBC	−0.10 (−3.53 to 3.33)	0.96	−0.74 (−4.21 to 2.73)	0.67
Persistent with d-RBC	−2.07 (−3.70 to −0.44)	0.01	−1.69 (−3.36 to −0.01)	0.048
Time since donation, month, 1 unit	−0.02 (−0.05 to 0.01)	0.12	−0.03 (−0.06 to 0.01)	0.08

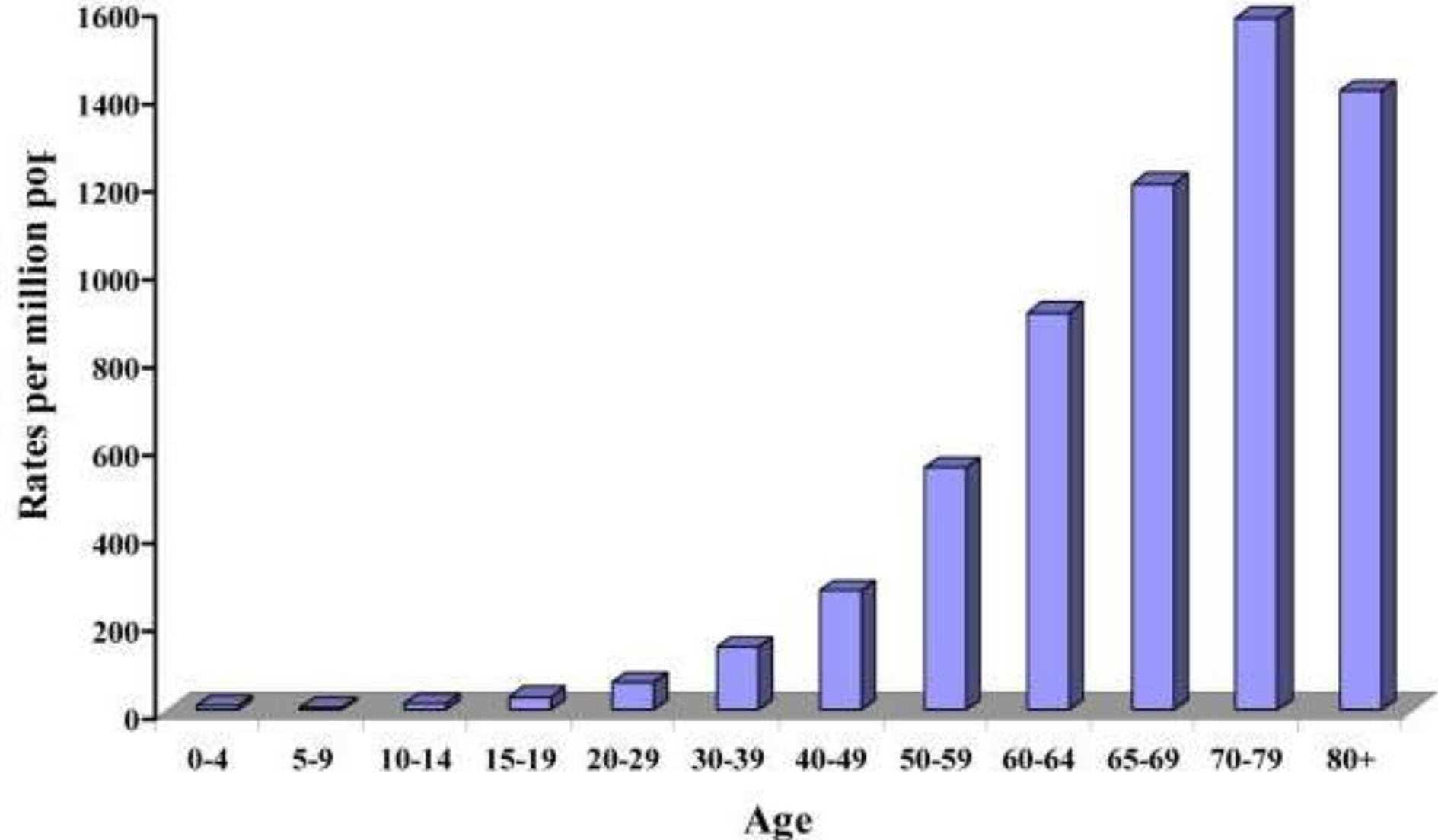
<sup>1</sup>Adjusted for age, sex, proteinuria, eGFR and body-mass index before donation, and hematuria after donation and variables with p-value less than 0.2 in univariate analysis, namely serum albumin, uric acid, total cholesterol at donation and donation for sibling ( $R^2 = 0.15$ ). Excluded variables from covariates which had a p-value more than 0.2 in univariate analysis: predonation variables of hematuria, smoking, family history of IgA nephropathy or Alport syndrome, serum hemoglobin and triglyceride; and postdonation variables of proteinuria. CI = confidence interval; d-RBC = dysmorphic red blood cells; eGFR = estimated glomerular filtration rate.

# Significado de la fusión parcial de podocitos

- Puede estar relacionada con episodios cortos de proteinuria o con la edad



# Incident Rates of Treated ESKD



Wiggins J. Podocytes and glomerular function with aging. *Semin Nephrol.* 2009;29:587-93.

# CONCLUSIONES

- Los pacientes con microhematuria glomerular persistente y alteraciones renales morfológicas deben excluirse como donantes vivos.

phic. Although histological findings could not be evaluated in this study, we expect that light microscopy in many of the donors would have revealed minor glomerular abnormality, mild mesangial proliferative change, or partially global sclerosis in glomeruli, with either mesangial IgA and C3 deposition by immunofluorescence (IgAN), as well as thin glomerular basement membrane (GBM) and segmental attenuation of the lamina densa (TBMN), or thickening

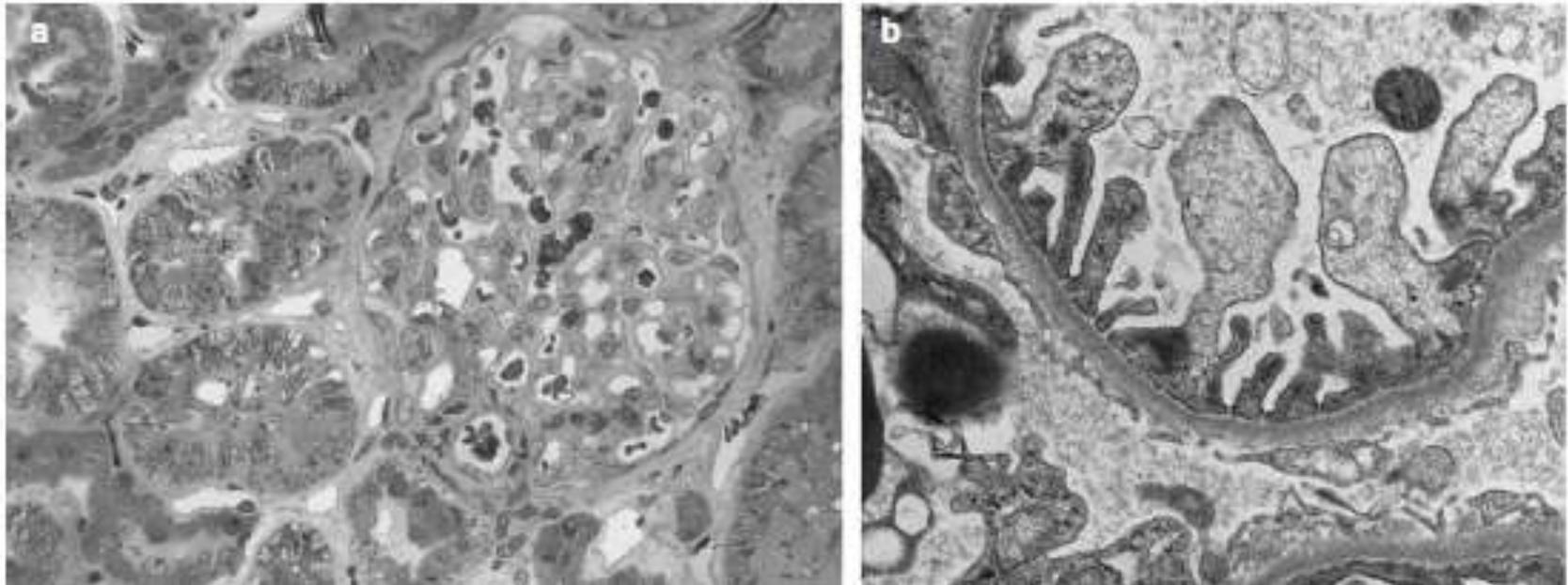
and lamellation of GBM (AS) on electron microscopy, as previously reported (19,20,29,30).

Although the paucity of evidence for the eligibility of candidates with isolated hematuria and their risk after donation has meant that the acceptability of glomerular hematuria in kidney donors remains controversial (19,20,29,31,32), our results clearly show that donors with persistent hematuria and d-RBC are at significant risk of developing persistent proteinuria or lower GFR, which also confer worse renal outcomes in patients with TBMN, IgAN or AS (31,33–36). The eligibility of potential donors with isolated but persistent dysmorphic hematuria should therefore be closely considered.

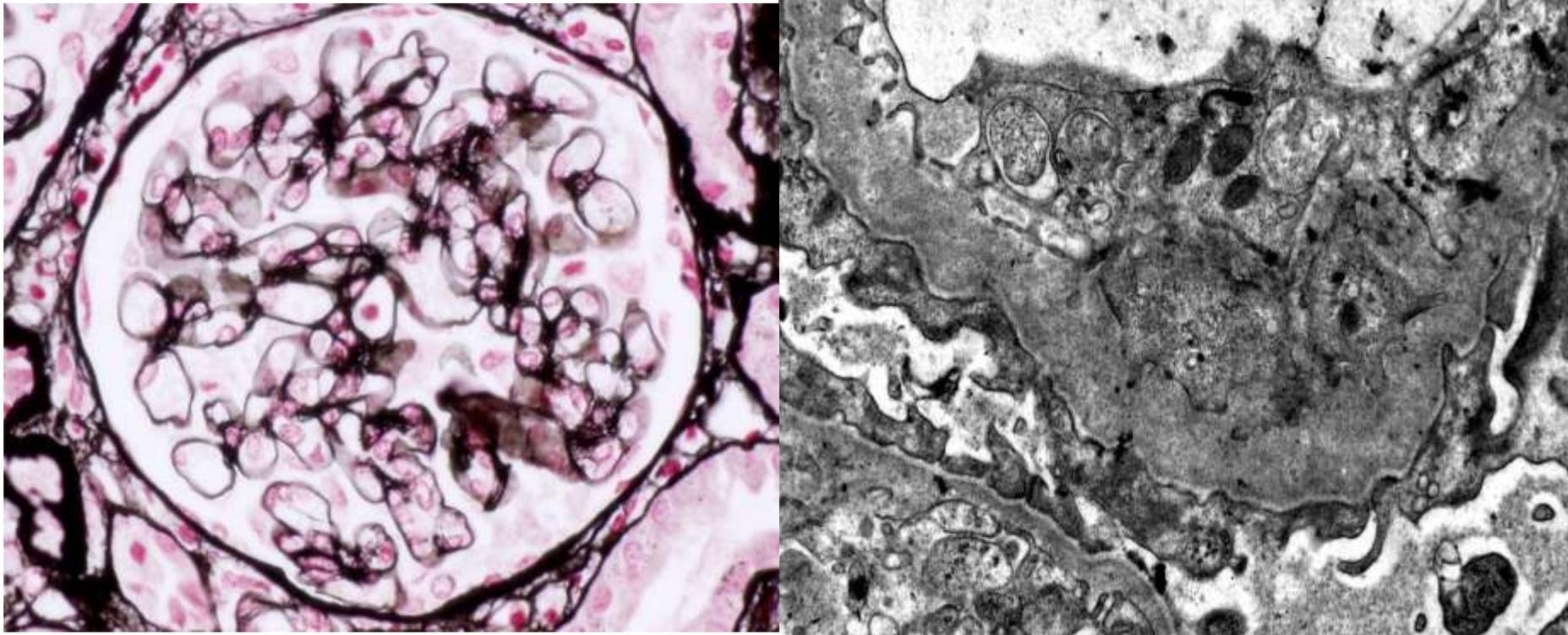
## Accepting prospective kidney donors with asymptomatic urinary abnormalities: Are we shooting in the dark?

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- Mujer 46 años, posible donante viva
- Hematuria dismórfica persistente
- Sin antecedentes familiares



FUSIÓN PARCIAL DE PODOCITOS  
Membrana basal glomerular normal

**SE DESCARTA A LA PACIENTE COMO DONANTE**

**MUCHAS GRACIAS POR SU ATENCIÓN**