# Identifying opportunities to improve evaluation for hematuria in a large health system

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#### **KEY FINDINGS & CONCLUSIONS**

- In a large US regional health system, nephrology and urology referral rates were low following hematuria diagnosis, highlighting substantial opportunities to improve the evaluation of hematuria
- Follow-up microscopy testing occurred in only 41.6% of patients despite guidelines recommending repeat microscopic evaluation<sup>5,6</sup>
- Referral to nephrology was minimal, even in patients with more severe concomitant proteinuria
- Overall, patients with appointments or referrals were more likely to be male, of White race, have hypertension, and have worse urinalysis proteinuria or hematuria categories
- Based on the findings presented here, subsequent analyses are being planned to help determine strategies to improve first-line management and follow-up of hematuria in primary care



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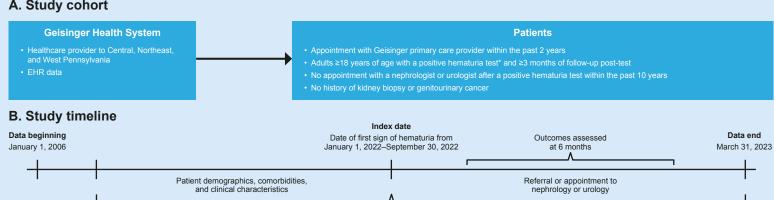
#### INTRODUCTION

- Hematuria is commonly encountered in primary care and presents in 1–18% of patients, depending on age, gender, frequency of testing, and the presence of risk factors1
- · As hematuria can be an initial sign of conditions such as glomerulonephritis and urologic cancer, adherence to medical society guidelines is key for timely diagnosis and patient management<sup>1-6</sup>
- There is currently limited evidence describing referral patterns and management of patients with hematuria to nephrology or urology by primary care physicians
- This study aimed to examine opportunities to improve hematuria management in the primary care setting for the early diagnosis of serious hematuria-associated conditions

#### **METHODS**

- This was a retrospective cohort study of adults with EHR data in the US Geisinger Health System, who tested positive for hematuria (defined as a positive dipstick test with blood 1+ or greater) between January 1, 2022, and September 30, 2022 (Figure 1)
- Descriptive statistics were used to compare characteristics of patients who were referred vs not referred and to summarize patients who underwent follow-up testing or presented with concomitant proteinuria
- For the univariate analysis, continuous variables were analyzed using a t-test, and categorical variables were analyzed using the Chi-square test

#### Figure 1. Study design A. Study cohort



C. Study objectives

**Primary objectives** To evaluate the proportion of patients with hematuria who had a referral to, or appointment with a nephrologist or urologist within 6 months of the index date

Medical history period

(January 1, 2021-September 30, 2021)

\*Defined as dipstick with blood 1+ or greater; †Defined as dipstick with protein 1+ or greater ACR, albumin/creatinine ratio; EHR, electronic health record; PCR, protein/creatinine ratio.

Secondary objectives

Follow-up period

(January 1, 2022-March 31, 2023)

· Confirmation of hematuria by repeat urinalysis or urine microscopy within 3 months of the index date

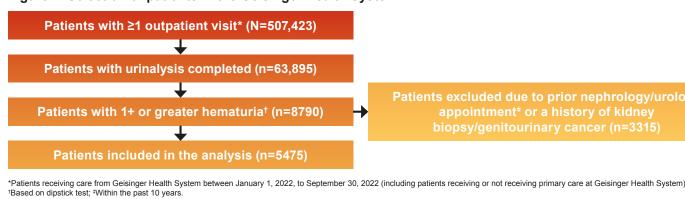
 Patients with concomitant proteinuria<sup>†</sup> at presentation · Patients with concomitant proteinuria who underwent quantitative ACR or PCR testing within

#### **RESULTS**

#### Patient selection

 Of 507,423 patients with ≥1 outpatient visit to a Geisinger primary care provider, 5475 patients were included in the analysis (Figure 2)

Figure 2. Selection of patients in the Geisinger Health System



# **Demographics and clinical characteristics**

- The majority of patients were female (78.0%), White (92.1%), and not Hispanic or Latino (93.1%), with a mean age of 60.5 years (**Table 1**)
- Overall, the most frequent comorbidities included hypertension (33.4%), diabetes (15.3%), and CKD (9.9%) (Table 2)

# Characteristics associated with appointment or referral to urology or nephrology

- In total, 774 of 5475 patients (14.1%) had an appointment or referral to urology (12.9%) or nephrology (1.8%) within 6 months of hematuria assessment
- In a univariate analysis, appointment or referral was higher for patients with greater hematuria (1+: 11.7% vs 2+: 18.0%; *P*<0.001) and patients with concomitant proteinuria (negative: 11.9%, trace: 15.6%, 1+: 15.8%, 2+: 14.9%, 3+: 17.5%; *P*=0.001) (**Table 2**)
- Other factors associated with higher appointment or referral rates included a tendency to receive primary care at Geisinger Health System (P<0.001), male sex (P<0.001), White race (P=0.019), and hypertension (*P*=0.001) (**Tables 1 and 2**)

# Patients with concomitant hematuria and proteinuria

- A total of 4952 patients had concurrent urinalysis protein and blood data available, of whom 2375 (48.0%) had 1+ or greater protein (Figure 3)
- Of these 2375 patients, 348 (14.7%) had quantitative ACR and/or PCR testing performed within 3 months of the index date
- This included 252 patients (72.4%) who had ACR testing and 197 patients (56.6%) who had PCR testing ACR and PCR levels by urinalysis proteinuria category are shown in Table 3
- Appointment or referral with nephrology or urology within 6 months of the index date was observed in 373 out of 2375 patients (15.7%) with concomitant proteinuria
- Of those with more severe proteinuria, 1.9% with 2+ and 5.1% with 3+ urinalysis proteinuria categories had an appointment or were referred to nephrology

# Confirmation of hematuria by repeat urinalysis and urine microscopy

- Of the 4187 patients with urine microscopy data available at the index date, 2905 (69.4%) had ≥3 RBCs/HPF (**Figure 4**)
- Among 1743 patients (41.6%) who had ≥1 repeat microscopy conducted, 1010 (57.9%) had ≥3 RBCs/HPF
- Overall, 1582 of 5475 patients (28.9%) had ≥1 repeat urinalysis test (Figure 5) Of the 1582 patients, 1086 (68.6%) had 1+ or greater hematuria

Figure 3. Index urinalysis proteinuria results (n=4952)

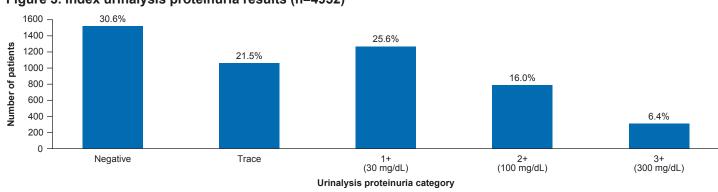


Figure 4. Index urine microscopy results (n=4187)

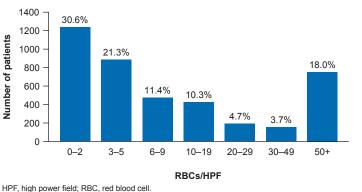


Figure 5. Repeat urinalysis hematuria results (n=1582)

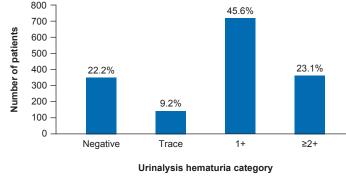


Table 1. Demographics of patients with hematuria

Category	Overall N=5475	No appointment or referral within 6 months n=4701	Appointment or referral within 6 months n=774	<i>P</i> -value*	
Receives Geisinger primary ca	re, n (%)				
Yes	2058 (37.6)	1643 (35.0)	415 (53.6)	<0.001	
Age, years					
Mean (SD)	60.5 (18.4)	60.5 (21.1)	60.9 (18.4)	0.600	
Sex, n (%)					
Male	1204 (22.0)	937 (19.9)	267 (34.5)	<0.001	
Race, n (%)					
White	5041 (92.1)	4312 (91.7)	729 (94.2)		
Black or African American	211 (3.9)	187 (4.0)	24 (3.1)	0.200	
Asian	75 (1.4)	67 (1.4)	8 (1.0)		
Other <sup>†</sup>	58 (1.1)	51 (1.1)	7 (0.9)		
Unknown	90 (1.6)	84 (1.8)	6 (0.8)		
Ethnicity, n (%)					
Hispanic or Latino	239 (4.4)	212 (4.5)	27 (3.5)		
Not Hispanic or Latino	5096 (93.1)	4362 (92.8)	734 (94.8)	0.100	
Unknown	140 (2.6)	127 (2.7)	13 (1.7)		
Non-Hispanic White	4799 (87.7)	4101 (87.2)	698 (90.2)	0.020	

'Native Hawaiian or other Pacific Islander,' 'two or more,' and 'other

Table 2. Comorbidities and clinical characteristics of patients with hematuria

Category	Overall N=5475	No appointment or referral within 6 months n=4701	Appointment or referral within 6 months n=774	<i>P-</i> value*	
Comorbidities,† n (%)					
Hypertension	1829 (33.4)	1529 (32.5)	300 (38.8)	0.001	
Diabetes	840 (15.3)	701 (14.9)	139 (18.0)	0.030	
Genitourinary infection	201 (3.7)	184 (3.9)	17 (2.2)	0.020	
CKD	542 (9.9)	467 (9.9)	75 (9.7)	0.800	
Acute kidney injury	95 (1.7)	83 (1.8)	12 (1.6)	0.100	
Glomerular disease	8 (0.1)	8 (0.2)	0	0.300	
Urinalysis hematuria category,	n (%)				
1+	3342 (61.0)	2952 (62.8)	390 (50.4)	<0.001	
2+	2133 (39.0)	1749 (37.2)	384 (49.6)		
Urinalysis proteinuria category	v, n (%)				
Negative	1514 (27.7)	1334 (28.4)	180 (23.3)		
Trace	1063 (19.4)	897 (19.1)	166 (21.5)	0.001	
1+ (30 mg/dL)	1267 (23.1)	1067 (22.7)	200 (25.8)		
2+ (100 mg/dL)	793 (14.5)	675 (14.4)	118 (15.3)		
3+ (300 mg/dL)	315 (5.8)	260 (5.5)	55 (7.1)		
Missing	523 (9.6)	468 (10.0)	55 (7.1)		

\*Continuous variables were analyzed using a t-test, and categorical variables were analyzed using the Chi-square test. †Comorbidities were identified using separate ICD-10 diagnostic codes during the 12 months prior to index date

Table 3 Quantification of proteinuria in nationts with homaturis

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Urinalysis proteinuria category	ACR levels, mg/g			PCR levels, mg/g					
	n	Mean (±SD)	Median (IQR)	n	Mean (±SD)	Median (IQR)			
1+ (30 mg/dL)	91	188 (355)	94 (33–215)	47	492 (403)	333 (145–750)			
2+ (100 mg/dL)	61	595 (598)	383 (195–793)	53	1403 (1116)	1193 (475–2053)			
3+ (300 mg/dL)	39	3131 (2791)	2262 (981–4203)	42	5509 (4405)	4558 (2726–6581)			

ACR, albumin/creatinine ratio; IQR, interquartile range; PCR, protein/creatinine ratio; SD, standard deviatio

# LIMITATIONS

- Data recorded in the database may be subject to human or technical error or data omission
  - Laboratory values, appointments, and/or referrals may be incomplete for patients who received care
  - outside of Geisinger Health System Ascertainment of clinical characteristics relied on ICD-10 diagnosis codes
- Patients who received more care due to illness are more likely to be represented in the database
- The Geisinger Health System database only includes individuals who have interacted with the medical system and may not be representative of populations beyond those identified in the database, such as patients with undiagnosed hematuria who did not receive initial urinalysis

#### **Abbreviations**

ACR, albumin/creatinine ratio; CKD, chronic kidney disease; EHR, electronic health record; HPF, high power field; ICD-10, International Classification of Diseases, 10th Revision; IQR, interquartile range; KDIGO, Kidney Disease Improving Global Outcomes; PCR, protein/creatinine ratio; RBC, red blood cell; SD, standard deviation;

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# **Disclosures**

AC, TJ, and YH are employees of Geisinger Health System, Danville, Pennsylvania, USA. BN and CA are employees of Novartis Pharmaceuticals Corporation, East Hanover, New Jersey, USA.

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