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Pathological and radiological irregularities in patients with chronic kidney disease - An observational study



Dushyant Pawar¹, Vinayak Bhoi², Shital Pawar³, Dilip Patil⁴

¹Associate Professor, ⁴Professor and Head, Department of Medicine, ³Tutor, Department of Community Medicine, JMFs ACPM Medical College, Dhule, ²Senior Resident, Department of Medicine, Bharati Vidyapeeth Medical College, Pune, Maharashtra, India

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ABSTRACT

Background: Chronic kidney disease (CKD) encompasses a spectrum of different pathophysiologic processes associated with abnormal renal function and progressive decline in glomerular filtration rate (GFR). Aims and Objectives: The objectives of the study are 1) to assess basic tests and kidney function tests in patients with CKD, (2) to assess coagulation profile, sugar levels, and platelets levels in patients of CKD, (3) to do the urine analysis in patients with CKD, and (4) to assess GFR among the study population. Materials and Methods: After taking the informed consent, detailed history was taken and clinical examination of patient was done. Pathological tests such as hemogram, renal function tests, liver function tests, coagulation profile, blood sugar levels, urine routine, and microscopy were performed. Ultrasonography of abdomen and pelvis was done. Correlation was done between severity of sepsis and mortality in patients. Results: Anemia was moderate (7-8.9 g/dl) in 38%, mild (9-10.9 g/dl) in 31%, while severe (<6.9) in 29 patients, that is, 29%. Thrombocytopenia was present in 58% of patients. Mean serum albumin in patients was 2.67 ± 0.551 mg/dl; serum globulin was 3.89 ± 0.345 mg/dl. Mean blood sugar of study subjects was 127.41 ± 61.387 mg/dl fasting, 199.67 ± 114.827 mg/dl post-prandial, and mean glycated hemoglobin was 5.63 ± 1.668 . On ultrasound acute pyelonephritis, 25 (25%) was the most common finding. Conclusion: Correlation of clinical findings with pathological and radiological findings is very important in patients with CKD. Severity of sepsis directly correlates with mortality of patients.

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Key words: Chronic kidney disease; Glomerular filtration rate; Pathological; Radiological, sepsis

INTRODUCTION

Chronic kidney disease (CKD) is the 12^{th} highest cause of death and 17^{th} highest cause of disability worldwide.¹ Understanding the entire depth of kidney disease becomes an integral part of preventing other organ damages as kidney damage is root cause for the most of them. Up to 90% of kidney function may be lost before symptoms are present, so annual checking of those at risk is essential.² Kidney damage for ≥ 3 months as defined by structural or functional abnormalities of the kidney, with or without decreased glomerular filtration rate (GFR), that can lead to decreased GFR, manifest by either: Pathological abnormalities or GFR <60 ml/min/1.73 m² for \geq 3 months, with or without kidney damage or markers of kidney damage, including abnormalities in imaging tests.³

Aims and objectives

The objectives of the study are as follows:

- 1. To assess basic tests and kidney function tests in patients with CKD
- 2. To assess coagulation profile, sugar levels, and platelets levels in patients of CKD

Address for Correspondence:

Dr. Dushyant Pawar, Associate Professor, Department of Medicine, JMFs ACPM Medical College, Dhule, Maharashtra, India. **Mobile:** +91-9823408616. **E-mail:** drdushyantpawar@gmail.com

- 3. To do the urine analysis in patients with CKD
- 4. To assess GFR among the study population.

MATERIALS AND METHODS

Study design

This was a observational study.

Sample size

The sample size was 100.

Sampling technique

This was simple random sampling.

Inclusion criteria

The following criteria were included in the study:

- a. Patients more than 18 year of age
- b. Male and female patients
- c. Patients who give valid informed written consent for the study
- d. Patients fulfilling the Kidney Disease Improving Global Outcome (2012) criteria for CKD.

Exclusion criteria

Patients <18 year of age, HIV, hepatitis B surface antigen, and hepatitis C virus positive patients were excluded from the study.

Procedure

After taking informed consent, detailed history was taken regarding the disease. A detailed clinical examination was done of all the study population. Routine investigations were done as and when required.

Following blood investigations were done:

- Hemogram
- Acute phase reactant: Erythrocyte sedimentation rate/C-reactive protein (CRP)
- Renal function tests: Creatinine/blood urea nitrogen (BUN)
- Serum electrolytes: $(Na^+/K^+/Ca^{++}/Po_4)$
- Liver function tests: Bilirubin: (Total/direct/indirect), serum glutamic oxaloacetic transaminase (SGOT), serum glutamic pyruvic transaminase (SGPT), and proteins: (Total/albumin/globulin/A: G)
- Coagulation profile: Bleeding time (BT), clotting time (CT), activated partial prothrombin time (aPTT), prothrombin time (PT), and international normalized ratio
- Blood sugar levels: Fasting (F) and post-prandial
- Glycated hemoglobin (HbA1C)
- Urine examination: Routine and microscopy.

RESULTS

In our study, mean total leukocyte count was 20031.05 ± 4388.859 ranging from 15600 to 37800. Mean platelet count of study population was 14.28 ± 3.761 . The mean creatinine level was 2.09 ± 1.848 , maximum levels were 16 mg/dl were as the minimum levels were 0.9 mg/dl. Mean BUN 113.57 ± 29.158 ranging within 34–229 (Table 1).

In present study, mean BT, CT, PT HbA1C was 3.86±1.064,5.49±1.291, 18.37±6.170, 5.63±1.668 respectively (Table 2).

Amongst USG findings, acute pyelonephritis was seen in 25%, small kidney was seen in 15%, renal stone with hydronephrosis and cystitis was seen in 11% (Table 3).

Sepsis was seen in 66 patients, septic shock was seen in 34% patients (Table 4).

The eGFR values of the current study compared with other studies are shown in Table 5.

According to WHO Anemia classification, in present study, 38 patients were having moderate anemia, 31 were having mild anemia, 29 were having severe anemia. Only Hemoglobin of 2 patients was within normal range (Chart 1). Majority of patients had platelet count between 1 lakh to 1.5 lakh. Platelet count more than 2 lakhs was found only in 2 patients (Chart 2).

In urine examination findings were:

- Appearance of urine cloudy in 56%, while turbid in 37% and 7% patients had clear urine
- 2. Color of urine In this study, urinary examination finding revealed white color urine in 52% patients, pale yellow color in 39% patients, and red urine was in 9% patients
- Protein/albumin Urinary proteins were 2+ in 45% patients while 42% had 3+ proteins in urine. All study subjects had proteinuria. This universal finding was ranging from trace to 4+
- 4. Urine sugar Glycosuria was present in 48% CKD patients having urosepsis
- 5. Red blood cells (RBCs) in urine RBC's in urine was observed microscopically in 51% patients
- 6. Pus cells In this study of CKD with urosepsis, urine examination of 38 (38%) patients had more than 140 pus cells, that is, abundant pus cells and 100% patients had pus cells in urine
- Casts 91% patients were had urinary casts which were white blood cell (WBC) cast, Granular cast, and RBC cast. Out of that WBC casts were predominant

in 53% patients, f/b granular casts 29%, and RBC casts in 9%

8. Out of total 100 patients urine culture, maximum growth of 52% of *Escherichia coli* was seen, 11% had growth of proteus, while *Klebsiella* and *Pseudomonas* growth was observed in 10% and 8% patients, respectively.

DISCUSSION

The present study is an observational study of total 100 patients conducted in the department of general medicine in a tertiary care institute. In the present study, patients had 9.01 ± 1.997 g/dl mean hemoglobin (Hb). Anemia was a universal finding in all patients, with maximum patients having moderate anemia, 38 patients, that is, 38%, followed by mild anemia, in 31 patients, that is, 31%, while severs anemia was present in 29 patients, that is, 29%. Only 2 (2%) patients had Hb in normal range.

The results of our study were similar with Bhattacharjee et al.,⁴ Sathyan et al.,⁵ and Afshar et al.⁶ However, James et al.,⁷ did not found Hb level <11.9 in any of the patients in their study.

In our study, mean total leukocyte count was 20031.05 ± 4388.859 ranging from 15600 to 37800. Similar findings were seen in Tambo et al.,⁸ found leukocytosis in their study and Mansur et al.,⁹ who found leukocytes $(1000/\mu)$ 15±6.

Mean platelet count of study population was 14.28 ± 3.761 (×10000). Thrombocytopenia was present in 58% of patients with the most patients had platelet in range of 1–1.49 lakhs, that is, 45 patients, 45–13% had platelets <1 lakh.

Buonaiuto et al.,¹⁰ noted thrombocytopenia in 14.3% cases of their study and Mansur et al.,⁹ found platelets $(1000/\mu l)$ 248±162–315±156 0.0004.

In our study, mean creatinine level was 2.09 ± 1.848 maximum 16 and minimum 0.9 mg/dl. Mean BUN 113.57 ±29.158 ranging within 34–229. This results were consistent with Umesha et al.,¹¹ and Tambo et al.⁸

Mean serum albumin in patients was 2.67 ± 0.551 mg/dl; serum globulin was 3.89 ± 0.345 mg/dl while total protein was 6.02 ± 0.651 mg/dl. A/G ratio was 1.528 ± 0.6632 . Other similar studies of Sr. albumin (g/dl) are mentioned below:

In this study of CKD with urosepsis, urine examination of 38 (38%) patients had more than 140 pus cells, that is, abundant pus cells and 100% patients had pus cells in urine. Urinary proteins were 2+ in 45% patients while 42% had 3+ proteins in urine. All study subjects had proteinuria ranging from trace to 4+. About 48% CKD patients had sugar in urine. RBC's in urine was present in 51% patients. About 91% patients had urinary casts which were WBC cast, granular cast, and RBC cast. Out of that WBC casts were predominant in 53% patients, followed by granular casts 29% and RBC casts in 9%.

| Table 1: Investigations of study subjects | | | | | |
|---|-----|---------|---------|----------|----------|
| Parameters | n | Minimum | Maximum | Mean | SD (±) |
| qSOFA Score | 100 | 1 | 3 | 1.60 | 0.550 |
| HB | 100 | 5 | 16 | 9.01 | 1.997 |
| TLC | 100 | 15600 | 37800 | 20031.05 | 4388.859 |
| Platelet (×10000) | 100 | 7 | 24 | 14.28 | 3.761 |
| PCT (ng/ml) | 100 | 0.9 | 16 | 7.59 | 3.9 |
| CREAT | 100 | 1 | 9 | 2.09 | 1.848 |
| BUN | 100 | 34 | 229 | 113.57 | 29.158 |
| Na⁺ | 100 | 125 | 146 | 134.18 | 3.424 |
| K⁺ | 100 | 3 | 7 | 5.53 | 0.758 |
| Ca++ | 100 | 8 | 10 | 8.51 | 0.541 |
| PO4- | 100 | 3 | 7 | 6.03 | 0.481 |
| Bilirubin total | 100 | 1 | 3 | 1.12 | 0.409 |
| Direct | 100 | 0 | 2 | 0.11 | 0.373 |
| Indirect | 100 | 0 | 3 | 0.98 | 0.284 |
| SGOT | 100 | 20 | 152 | 40.64 | 33.561 |
| SGPT | 100 | 20 | 224 | 42.16 | 37.913 |
| ALP | 100 | 41 | 262 | 158.06 | 19.787 |
| Serum globulin | 100 | 2 | 4 | 3.89 | 0.345 |
| Serum albumin | 100 | 1 | 3 | 2.67 | 0.551 |
| Total protein | 100 | 4 | 7 | 6.02 | 0.651 |
| A/G ratio | 100 | 1.0 | 5.4 | 1.528 | 0.6632 |

qSOFA: Quick sequential organ failure assessment, HB: Hemoglobin, TLC: Total leucocyte count, PCT: Procalcitonin, BUN: Blood urea nitrogen, SGOT: Serum glutamic oxaloacetic transaminase, SGPT: Serum glutamic pyruvic transaminase, ALP: Alkaline phosphatase

| Table 2: Coagulation and sugar profile ofpatients | | | | | |
|---|-----|---------|---------|--------|---------|
| Parameters | n | Minimum | Maximum | Mean | SD |
| BT | 100 | 3 | 8 | 3.86 | 1.064 |
| СТ | 100 | 2 | 10 | 5.49 | 1.291 |
| PT | 100 | 13 | 43 | 18.37 | 6.170 |
| aPTT | 100 | 30 | 38 | 32.58 | 1.939 |
| FBS | 100 | 68 | 385 | 148.41 | 79.176 |
| PPBS | 100 | 100 | 458 | 224.44 | 123.788 |
| HbA1C | 100 | 3 | 10 | 5.63 | 1.668 |

BT: Bleeding time, CT: Clotting time, aPTT: Activated partial prothrombin time, PT: Prothrombin time, FBS: Fasting blood sugar, PPBS: Post-prandial blood sugar, HbA1C: Glycated hemoglobin

| Table 3: USG abdomen findings in patients | | | | |
|---|-----------|---------|--|--|
| Finding | Frequency | Percent | | |
| APN | 25 | 25.0 | | |
| Small kidney | 15 | 15 | | |
| Renal stone with HN | 11 | 11 | | |
| Cystitis | 11 | 11.0 | | |
| Ureteric calculi, HN HU | 9 | 9.0 | | |
| Stent <i>in situ</i> | 9 | 9.0 | | |
| Renal abscess | 7 | 7.0 | | |
| Renal papillary necrosis | 6 | 6 | | |
| EPN | 3 | 3.0 | | |
| Prostatic abscess | 2 | 2.0 | | |
| Bladder tumor | 2 | 2.0 | | |
| Total | 100 | 100.0 | | |

USG: Ultrasound, HU: Hounsfield units, EPN: Emphysematous pyelonephritis, APN: Acute pyelonephritis

| Table 4: Correlation of severity of sepsis and | |
|--|--|
| mortality in patients | |

| Severity of | Mor | Mortality | |
|--------------------|------|-----------|-------|
| sepsis | N | Y | |
| Severity of sepsis | | | |
| Sepsis | | | |
| N | 59 | 7 | 66 |
| % | 89.4 | 10.6 | 100.0 |
| Septic shock | | | |
| N | 14 | 20 | 34 |
| % | 41.2 | 58.8 | 100.0 |
| Total | | | |
| Ν | 73 | 27 | 100 |
| % | 73.0 | 27.0 | 100.0 |
| P=0.001 (S) | | | |

P=0.001(S)

| Table 5: Comparision | of eGFR with other studies |
|----------------------|----------------------------|
| | |

| Author and | eGFR (ml/min/1.73 m²) | | | | |
|-------------------------------------|-----------------------|-----------|-----------|-----------|--|
| year of study | ≥60 | 45–59 | 30–44 | <30 | |
| Wang et al. (2011) ¹² | 4.10±0.02 | 4.04±0.02 | 3.98: | ±0.05 | |
| James et al. (2008) ⁷ | 3.7 (0.4) | 3.6 (0.5) | 3.6 (0.5) | 3.5 (0.5) | |
| Tambo et al. (2013) ⁸ | 2.8±0.7 | | | | |
| Present study | 2.67±0.551 | | | | |

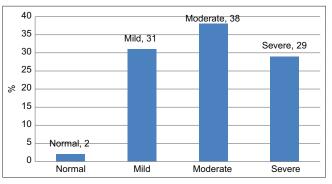


Chart 1: The World Health Organization anemia classification in the study subjects

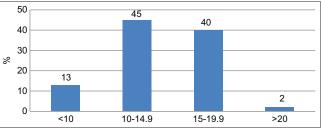


Chart 2: Platelet range among study subjects

Buonaiuto et al.,¹⁰ found similar results of pyuria and microscopic hematuria in their study, Tambo et al.,⁸ also observed pyuria in their study.

In our study, serum procalcitonin (PCT) level ranging from 0.9 to 16 ng/ml with mean of 7.59±3.9. Similar findings were noted by van Nieuwkoop et al.,¹³ Meisner (2014),¹⁴ Uzzan et al.,¹⁵ and Sugimoto et al.,¹⁶ who found PCT as a good marker for diagnosis urosepsis.

Luzzani et al.,¹⁷ and Simon et al.,¹⁸ found PCT more reliable than the acute-phase protein CRP and Mansur et al.,⁹ also had similar finding of raised PCT (ng/dl) of (n) 5.1 ± 7.3 in 47% in their study. Thus, according to the various studies PCT is a good marker even for urosepsis.

On ultrasound of CKD patients having urosepsis, the most common finding was acute pyelonephritis (APN) 25 (25%) followed by small kidney which was present in 15%, cystitis and renal stone 11 (11%) each, stent is situ and ureteric calculus with HN HU was finding present in 9 (9%) patients and renal abscess present in 7 (7%). Renal papillary necrosis was present in 6 (6%) patients of 3 (3%) patient had emphysematous pyelonephritis whereas 2 (2%) patient each had prostatic abscess and bladder tumor in CKD patients having urosepsis in the study.

These findings were consistent with Tambo et al.,⁸ Sørensen et al.,¹⁹ Ruiz-Mesa et al.,²⁰ and Buonaiuto et al.¹⁰

In the present study, sepsis led to mortality in 10.6% of patients of sepsis; however, mortality was found in 58.8% patients of uroseptic shock thus uroseptic shock causes more mortality than urosepsis alone. Lee et al.,²¹ also found similar results, sepsis-related mortality was (4.9) in health-care associated (HCA) whereas hospital associated (HOA) caused (14.6%) death. However, total mortality at discharge in there study was HCA (9.8%) mortality and HOA (41.5%) mortality.

Padkin et al.,²² reported 47.3% mortality, but their study only includes severe sepsis that occurred during the first 24 h after intensive care unit admission.

Furthermore, Ferrer et al.,²³ in a recent article on outcome of a nationwide educational intervention based on the Surviving Sepsis Campaign guidelines, reported a basal hospital mortality rate of 44%. Degoricija et al.,²⁴ also found mortality rate for sepsis as 17%, severe sepsis 33.7%, septic shock 72.1%, and multiple organ dysfunction syndrome 74.4%.

Limitations of the study

We could study only 100 patients. By increasing the sample size, prevalence of CKD could have been stidied.

CONCLUSION

All patients had albuminuria and pyuria. Abundant pus cells were present in 38% patients. The maximum number of patients belonged to CKD Stage 3 (eGFR- 30–59). On ultrasonography, APN (25%) was the most common cause of urosepsis. Thorough pathological, microbiological, and radiological investigations are important to assess the damage and to start early treatment.

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Authors Contribution:

DP - Concept and design of the study and prepared first draft of manuscript; **VB** - Interpreted the results, reviewed the literature, and manuscript preparation; **SP and DP** - Concept, coordination, statistical analysis and interpretation, preparation of manuscript, and revision of the manuscript.

Work attributed to:

Department of Medicine, JMFs ACPM Medical College, Dhule - 424 001, Maharashtra, India.

Orcid ID:

Dr. Dushyant Pawar - © http://orcid.org/0000-0002-7485-2391 Vinayak Bhoi - © http://orcid.org/0000-0001-8201-0760 Dr. Shital Pawar - © http://orcid.org/0000-0002-0521-7781

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