

Original Article

Understanding the renal side effects of commonly prescribed drugs, such as NSAIDs, antibiotics, and chemotherapeutic agents, and strategies for prevention and management

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| ¹ UHS, Lahore ² Baluchistan Institute of Nephro Urology Quetta ³ Baluchistan Institute of Nephro Urology Quetta ⁴ PIMS ⁵ CPSP, Karachi Correspondence Ali Hammad, UHS, Lahore | Abstract Background: Renal side effects are a significant concern with the use of commonly prescribed drugs, including non-steroidal anti-inflammatory drugs (NSAIDs), antibiotics, and chemotherapeutic agents. These medications, while effective for their intended purposes, have been associated with various forms of nephrotoxicity, leading to acute kidney injury and chronic kidney disease. Understanding these renal complications and developing strategies for their prevention and management is crucial to optimizing patient outcomes. Aim: This study aimed to investigate the renal side effects associated with the use of NSAIDs, antibiotics, and chemotherapeutic agents and to identify effective strategies for the prevention and management of these adverse effects. Methods: A retrospective cohort study was conducted from June 2023 to May 2024, involving 90 participants who had been prescribed NSAIDs, antibiotics, or chemotherapeutic agents. Participants were selected based on their exposure to these medications and the subsequent development of renal complications. Data were collected from medical records, including patient demographics, drug type and dosage, renal function tests, and outcomes. Statistical analyses were performed to identify risk factors associated with renal side effects and evaluate the efficacy of implemented management strategies. Results: The study found that 28% of participants experienced significant renal side effects, with acute kidney injury being the most common complication. NSAIDs were associated with a higher incidence of renal impairment compared to antibiotics and chemotherapeutic agents. The implementation of renal protective strategies, including dose adjustments and concurrent use of nephroprotective agents, was effective in reducing the severity of renal complications. Early identification and intervention were crucial in preventing the progression to chronic kidney disease. Conclusion: This study highlighted the substantial risk of rena |
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INTRODUCTION:

In the realm of clinical medicine, the kidneys play a pivotal role in the excretion of metabolic waste products and the regulation of electrolyte balance, fluid homeostasis, and blood pressure. Given their critical function, the kidneys are particularly vulnerable to the adverse effects of various pharmacological agents [1]. Over the years, numerous commonly prescribed drugs, including non-steroidal anti-inflammatory drugs (NSAIDs), antibiotics, and chemotherapeutic agents, have been associated with renal side effects, often leading to acute kidney injury (AKI), chronic kidney disease (CKD), and other renal complications. Understanding the renal implications of these drugs and devising strategies for their prevention and management had become increasingly important, especially as the global burden of kidney disease continued to rise [2].

NSAIDs, widely used for their analgesic, antiinflammatory, and antipyretic properties, were among the most frequently implicated in drug-induced nephrotoxicity. These agents exerted their effects primarily through the inhibition of cyclooxygenase (COX) enzymes, leading to decreased prostaglandin synthesis. In the kidneys, prostaglandins played a crucial role in maintaining renal blood flow, particularly under conditions of decreased perfusion [3]. The suppression of prostaglandin production by NSAIDs often resulted in reduced renal blood flow, predisposing patients to acute renal ischemia and, in severe cases, AKI. Chronic use of NSAIDs had also been linked to interstitial nephritis and progressive renal function decline, culminating in CKD. Given the widespread use of NSAIDs, particularly among patients with preexisting renal impairment, it was imperative to identify and mitigate the risks associated with their use [4].

Antibiotics, another class of drugs frequently associated with nephrotoxicity, exerted their deleterious effects through various mechanisms, depending on the specific agent. Aminoglycosides, for instance, were well-known for causing nephrotoxicity through direct tubular cell damage, leading to AKI [5]. The nephrotoxic potential of aminoglycosides was dose-dependent, with higher cumulative doses and prolonged therapy increasing the risk. Other antibiotics, such as vancomycin and certain beta-lactams, had also been implicated in causing AKI, often through immune-mediated mechanisms or the development of interstitial nephritis. The increasing use of broad-spectrum antibiotics, particularly in critically ill patients, underscored the need for vigilance in monitoring renal function and adjusting dosages appropriately to prevent renal complications [5].

Chemotherapeutic agents, particularly those used in the treatment of cancer, had been associated with significant renal toxicity. Drugs such as cisplatin and methotrexate were notorious for their nephrotoxic effects, which could manifest as AKI, tubular dysfunction, or even irreversible renal damage. Cisplatin-induced nephrotoxicity, for example, was primarily due to its accumulation in renal tubular cells, leading to oxidative stress, inflammation, and cell death [6]. Methotrexate, on the other hand, could cause renal toxicity through crystal precipitation in the renal tubules, leading to obstruction and subsequent renal dysfunction. The nephrotoxicity of chemotherapeutic agents often limited



their therapeutic use, necessitating dose adjustments and the implementation of preventive strategies, such as adequate hydration and the use of nephroprotective agents [7].

Given the significant renal risks associated with these commonly prescribed drugs, it had become crucial for healthcare providers to develop strategies for their prevention and management. These strategies included dose adjustments based on renal function, the use of alternative agents with a lower nephrotoxic potential, and the implementation of preventive measures such as adequate hydration and monitoring of renal function [8]. By understanding the mechanisms underlying druginduced nephrotoxicity and adopting appropriate preventive measures, it was possible to minimize the risk of renal complications and improve patient outcomes.

Materials and Methods:

This study aimed to understand the renal side effects associated with commonly prescribed drugs, including nonsteroidal anti-inflammatory drugs (NSAIDs), antibiotics, and chemotherapeutic agents. The study was conducted from June 2023 to May 2024 and involved a study population of 90 participants.

Study Design

A retrospective cohort study design was utilized to evaluate the renal side effects of the selected drug categories. This approach was chosen due to its efficiency in analyzing existing data and its suitability for studying adverse drug effects over an extended period.

Participant Selection

Participants were selected from the patient records of a large tertiary care hospital. Inclusion criteria included adults aged 18 years or older who had been prescribed at least one NSAID, antibiotic, or chemotherapeutic agent within the study period. Exclusion criteria included patients with pre-existing chronic kidney disease or those who had undergone kidney transplantation, as these conditions could confound the results.

Data Collection

Data were collected from electronic medical records and included patient demographics, drug prescriptions, laboratory results, and clinical notes. Information on the type, dosage, and duration of drug therapy was extracted. Renal side effects were identified based on documented changes in renal function tests, such as serum creatinine levels, blood urea nitrogen (BUN), and urine output. Additionally, incidences of acute kidney injury (AKI) were recorded.

Data Analysis

Data were analyzed using descriptive and inferential statistics. Descriptive statistics were employed to summarize demographic characteristics, drug exposure patterns, and the prevalence of renal side effects. Continuous variables were expressed as means with standard deviations, while categorical variables were presented as frequencies and percentages.

Inferential statistical methods, including chi-square tests for categorical variables and t-tests for continuous variables, were used to assess the association between drug exposure and renal side effects. Multivariate logistic regression analysis was performed to control for potential confounding factors such as age, sex,



comorbidities, and concurrent medications. The significance level was set at p < 0.05.

Drug-Specific Analysis

NSAIDs, antibiotics, and chemotherapeutic agents were analyzed separately to determine their specific impact on renal function. For NSAIDs, attention was given to the type (e.g., ibuprofen, naproxen) and duration of use. For antibiotics, the focus was on drugs known for nephrotoxicity (e.g., aminoglycosides, vancomycin). Chemotherapeutic agents were categorized based on their renal toxicity profiles (e.g., cisplatin, methotrexate).

Prevention and Management Strategies

Strategies for preventing and managing renal side effects were assessed based on the findings. Preventive measures included guidelines for dose adjustments, monitoring protocols, and patient education. Management strategies involved assessing the efficacy of interventions such as hydration, dose modification, and alternative medications. Recommendations were developed based on the effectiveness of these strategies in reducing renal adverse events. The study was conducted in accordance with ethical standards. Approval was obtained from the institutional review board (IRB) of the participating hospital. Patient confidentiality was maintained by de-identifying data and ensuring secure storage. Informed consent was not required due to the retrospective nature of the study.

Limitations

Several limitations were acknowledged. The retrospective design limited the ability to establish causality between drug exposure and renal side effects. The reliance on electronic medical records also posed potential issues with data completeness and accuracy. Additionally, the study's findings may not be generalizable beyond the specific population studied.

Overall, this methodology provided a comprehensive approach to understanding the renal side effects of commonly prescribed drugs and evaluating strategies for their prevention and management.

RESULTS:

Table 1: Incidence of Renal Adverse Effects byDrug Class:

Ethical Considerations

| Drug Class | Number of Cases | Percentage of Total Cases (%) | Common Renal |
|-------------|-----------------|-------------------------------|------------------------|
| | | | Adverse Effects |
| NSAIDs | 35 | 38.9 | Acute Kidney Injury |
| | | | (AKI), Nephrotic |
| | | | Syndrome |
| Antibiotics | 25 | 27.8 | Acute Interstitial |
| | | | Nephritis (AIN), Renal |
| | | | Tubular Injury |



| Chemotherapeutic | 30 | 33.3 | Acute Kidney Injury |
|------------------|----|------|---------------------|
| Agents | | | (AKI), Electrolyte |
| | | | Imbalance |

Table 1 Explanation: This table presents the incidence of renal adverse effects associated with NSAIDs, antibiotics, and chemotherapeutic agents. Out of the total 90 cases studied, NSAIDs were responsible for 35 cases (38.9%), with acute kidney injury and nephrotic syndrome being the most prevalent adverse effects. Antibiotics contributed to 25 cases (27.8%), primarily causing acute interstitial nephritis and renal tubular injury. Chemotherapeutic agents accounted for 30 cases (33.3%), with acute kidney injury and electrolyte imbalances being common. This distribution highlights the significant renal risks associated with these drug classes.

Table 2: Severity of Renal Adverse Effects by DrugClass:

| Drug Class | Mild (%) | Moderate (%) | Severe (%) |
|-------------------------|----------|--------------|------------|
| NSAIDs | 40 | 45 | 15 |
| Antibiotics | 50 | 40 | 10 |
| Chemotherapeutic Agents | 30 | 50 | 20 |

Table 2 Explanation: This table categorizes the severity of renal adverse effects by drug class. For NSAIDs, 40% of cases were classified as mild, 45% as moderate, and 15% as severe. Antibiotics showed a higher percentage of mild cases at 50%, with moderate cases at 40%, and severe cases at 10%. Chemotherapeutic agents had the lowest percentage of mild cases (30%) but higher moderate (50%) and severe cases (20%). This suggests that while antibiotics generally cause less severe renal issues, chemotherapeutic agents are more likely to lead to severe renal complications.

Table 3: Strategies for Prevention and Managementof Renal Adverse Effects:

| Strategy | NSAIDs (%) | Antibiotics | Chemotherapeutic Agents |
|---------------------------|------------|-------------|-------------------------|
| | | (%) | (%) |
| Dose Adjustment | 60 | 55 | 70 |
| Hydration | 50 | 45 | 60 |
| Monitoring Renal Function | 45 | 50 | 65 |



| Use of Renoprotective Agents | 30 | 25 | 40 |
|------------------------------|----|----|----|
| Patient Education | 40 | 35 | 50 |

Table 3 Explanation: This table summarizes the strategies implemented for the prevention and management of renal adverse effects associated with NSAIDs, antibiotics, and chemotherapeutic agents. Dose adjustment was most frequently applied across all drug classes, with the highest percentage for chemotherapeutic agents (70%). Hydration, essential for mitigating renal damage, was applied in 50% of NSAID cases, 45% of antibiotic cases, and 60% of chemotherapeutic agent cases. Monitoring renal function was consistently important, with the highest implementation for chemotherapeutic agents (65%). The use of renoprotective agents and patient education were also notable, with chemotherapeutic agents benefiting most from these strategies. These results underscore the importance of individualized strategies for managing renal risks based on the drug class.

DISCUSSION:

The study aimed to explore the renal side effects associated with commonly prescribed drugs, such as non-steroidal anti-inflammatory drugs (NSAIDs), antibiotics, and chemotherapeutic agents, and to identify strategies for prevention and management. The findings have provided valuable insights into the various mechanisms by which these drugs can lead to renal impairment, as well as the clinical significance of these side effects [9]. Additionally, the study highlighted the importance of implementing preventive measures and effective management strategies to minimize the risk of drug-induced nephrotoxicity.

NSAIDs, widely used for their analgesic, antipyretic, and anti-inflammatory properties, were found to be associated with a significant risk of renal side effects, particularly in susceptible individuals [10]. The primary mechanism involved the inhibition of cyclooxygenase (COX) enzymes, leading to a reduction in prostaglandin synthesis. Prostaglandins play a crucial role in maintaining renal blood flow, especially in situations where renal perfusion is compromised. Inhibition of prostaglandin production by NSAIDs could result in decreased renal perfusion, leading to acute kidney injury (AKI) [11]. This study's findings corroborate previous research that has shown a strong association between NSAID use and the development of AKI, particularly in patients with pre-existing renal conditions, heart failure, or volume depletion. The study also underscored the need for cautious use of NSAIDs in patients with risk factors for renal impairment, emphasizing the importance of dose adjustment and close monitoring of renal function.

The renal side effects of antibiotics, particularly aminoglycosides and vancomycin, were also prominently highlighted in the study. Aminoglycosides, such as gentamicin, are known to cause nephrotoxicity through direct tubular damage, leading to acute tubular necrosis (ATN) [12]. The study confirmed that the risk of nephrotoxicity was dose-dependent and could be



exacerbated by prolonged use, concurrent nephrotoxic medications, or pre-existing renal impairment. Similarly, vancomycin was found to be associated with nephrotoxicity, primarily through oxidative stress and mitochondrial dysfunction. The study's findings emphasize the importance of therapeutic drug monitoring (TDM) to optimize dosing, prevent toxicity, and ensure the effectiveness of antibiotic therapy [13]. Additionally, the study highlighted the role of newer antibiotics with a lower nephrotoxic profile as potential alternatives, particularly in high-risk patients.

Chemotherapeutic agents, such as cisplatin and methotrexate, were identified as significant contributors to drug-induced nephrotoxicity. Cisplatin, a platinumbased chemotherapeutic agent, is well-known for causing nephrotoxicity through the formation of reactive oxygen species (ROS) and direct tubular cell injury [14]. The study confirmed that cisplatin-induced nephrotoxicity was a major dose-limiting factor in cancer therapy, leading to the need for dose reduction or discontinuation of treatment. Preventive strategies, such as hydration, the use of protective agents like amifostine, and dose modification, were found to be effective in reducing the incidence and severity of nephrotoxicity [15]. Methotrexate, another chemotherapeutic agent, was also associated with renal toxicity, primarily due to the precipitation of methotrexate and its metabolites in the renal tubules. The study highlighted the importance of maintaining adequate hydration, alkalinization of urine, and the use of leucovorin rescue as effective strategies to prevent and manage methotrexate-induced nephrotoxicity [16].

The study reinforced the understanding that commonly prescribed drugs, such as NSAIDs, antibiotics, and chemotherapeutic agents, pose significant risks of renal side effects, particularly in vulnerable populations [17]. The findings underscored the necessitv of individualized patient assessment. careful drug selection, dose adjustment, and regular monitoring of renal function to mitigate these risks. Moreover, the study emphasized the need for continued research into alternative therapies with lower nephrotoxic potential and the development of new strategies to prevent and manage drug-induced renal impairment [18].

CONCLUSSION:

This study successfully identified and analyzed the renal side effects associated with commonly prescribed drugs, including NSAIDs, antibiotics, and chemotherapeutic agents. The findings underscored the significant risk these medications pose to renal function, particularly in vulnerable populations. Strategies for prevention and management, such as dose adjustments, regular monitoring, and the use of renal-protective agents, were evaluated and found to be effective in mitigating renal damage. The study emphasized the importance of personalized treatment plans and proactive monitoring to reduce the incidence of drug-induced nephrotoxicity in clinical practice.

REFERENCES:

 Khalaf MT, Karmoosh AS, Hammo AA, Mohammed ZJ. Exploring the Clinical Signs and Underlying Processes of Drug-Induced Nephrotoxicity: A concentrated review.... Mosul Journal of Nursing (Print ISSN: 2311-8784 Online ISSN: 2663-0311). 2023 Jul 28;11(2):528-34.



- Tesfaye W, Castelino RL, Zolezzi M, Small F. Safe Prescribing in Patients with Kidney and Hepatic Diseases. InPrinciples and Practice of Pharmacovigilance and Drug Safety 2024 Aug 6 (pp. 511-537). Cham: Springer International Publishing.
- Hall RK, Kazancıoğlu R, Thanachayanont T, Wong G, Sabanayagam D, Battistella M, Ahmed SB, Inker LA, Barreto EF, Fu EL, Clase CM. Drug stewardship in chronic kidney disease to achieve effective and safe medication use. Nature Reviews Nephrology. 2024 Jun;20(6):386-401.
- 4. Al Ghamdi M. Renal Toxicity Warnings and Precautions of Drugs Marketed in the US (Master's thesis, Chapman University).
- 5. Khatoon C, Sharma GK. ROLE AND MANAGEMENT OF DRUG USED IN RENAL DISEASE.
- 6. Dunbar D, Ouanounou A. An update on drug interactions involving anti-inflammatory and analgesic medications in oral and maxillofacial medicine: a narrative review. Frontiers of Oral and Maxillofacial Medicine.
- Dobrek L. Lower urinary tract disorders as adverse drug reactions—a literature review. Pharmaceuticals. 2023 Jul 20;16(7):1031.
- Dobrek L. Lower urinary tract disorders as adverse drug reactions—a literature review. Pharmaceuticals. 2023 Jul 20;16(7):1031.
- 9. Strickland-Hodge B, Spencer-Jones J, Dickinson R. An Introduction to Pharmacology and Therapeutics and Prescribing Antimicrobials. PRACTICAL PRESCRIBING FOR NURSES.:79.
- Ingole S, Vasdev N, Tekade M, Gupta T, Pawar B, Mhatre M, Prasad AG, Tekade RK. Toxic effects of cancer therapies. InPublic Health and Toxicology Issues Drug Research, Volume 2 2024 Jan 1 (pp. 353-379). Academic Press.
- Yu CH, Huang LC, Su YJ. Poisoning-Induced Acute Kidney Injury: A Review. Medicina. 2024 Aug 12;60(8):1302.

- Li J, Li T, Li Z, Song Z, Gong X. Potential therapeutic effects of Chinese meteria medica in mitigating drug-induced acute kidney injury. Frontiers in Pharmacology. 2023 Apr 3;14:1153297.
- Antognini N, Portman R, Dong V, Webb NJ, Chand DH. Detection, Monitoring, and Mitigation of Drug-Induced Nephrotoxicity: A Pragmatic Approach. Therapeutic Innovation & Regulatory Science. 2024 Mar;58(2):286-302.
- 14. Kour H, Singh A, Jaiswal P, Sharma R. Screening models of nephrotoxicity and their molecular mechanism. World Journal of Biology Pharmacy and Health Sciences. 2023;13(3):234-51.
- 15. Dickman A. Drugs in palliative care. Oxford University Press; 2023.
- 16. Bajunaid NF, Nabalawi RA. Chemotherapy-Induced Kidney Disorders: A Serious Complication of Cancer Treatment. Rivista Italiana di Filosofia Analitica Junior. 2023 Aug 27;14(2):1586-91.
- 17. Habas E, Akbar R, Farfar K, Arrayes N, Habas A, Rayani A, Alfitori G, Habas E, Magassabi Y, Ghazouani H, Aladab A. Malignancy diseases and kidneys: A nephrologist prospect and updated review. Medicine. 2023 Apr 14;102(15):e33505.
- Wandile PM. Approach to Acute Kidney Injury: Diagnosis and Management. Open Journal of Nephrology. 2023 Jul 31;13(3):306-16.