

# Acute kidney injury in Coronavirus disease-19 related pneumonia in the intensive care unit: a retrospective multicenter study, Saudi Arabia

Safwat A.M. Eldaboosy,<sup>1,2</sup> Amgad Awad,<sup>2,3</sup> Abdullah Farouk,<sup>2,4</sup> Waheed Mahdy,<sup>2,5</sup> Eman Abdelsalam,<sup>6,7</sup> Sameh O. Nour,<sup>1</sup> Ahmed Kabil,<sup>1</sup> Ahmad Taha,<sup>1</sup> Sameh Makled,<sup>1</sup> Ahmed Lotfi,<sup>1</sup> Usama Nabway,<sup>1</sup> Hatem Kanany<sup>8</sup>

<sup>1</sup>Department of Chest Diseases, Faculty of Medicine, Al-Azhar University, Cairo, Egypt

<sup>2</sup>Almoosa Specialist Hospital, Al Ahsa, Saudi Arabia

<sup>3</sup>Department of Internal Medicine, Faculty of Medicine, Al-Azhar University, Cairo, Egypt

<sup>4</sup>Department of Critical Care, Alexandria Faculty of Medicine, Alexandria, Egypt

<sup>5</sup>Department of Chest Diseases, Banha Faculty of Medicine, Banha, Egypt

<sup>6</sup>Department of Internal Medicine, Al-Azhar Faculty of Medicine for Girls, Cairo, Egypt

<sup>7</sup>King Khalid hospital, Hail, Saudi Arabia

ABSTRACT

<sup>8</sup>Department of Anesthesia and Critical Care, Faculty of Medicine, Al-Azhar University, Cairo, Egypt

**Background:** Acute kidney injury (AKI) poses a significant morbidity and mortality risk to critically ill COVID-19 patients. The aim of this study was to investigate the incidence, predictors, and outcomes of AKI in patients admitted to the intensive care unit (ICU) with critically ill COVID-19 pneumonia.

**Methods:** A multicenter retrospective study in Saudi Arabia of adult patients aged at least 18 years diagnosed with COVID-19 pneumonia and admitted to the intensive care unit between May 2020 and May 2021 was conducted. The occurrence of AKI and associated risk factors, the need for continous renal replacement therapy (CRRT), and the outcome were reported.

**Results:** The study included 340 patients admitted to the ICU with COVID-19. Their mean age was  $66.7\pm13.4$  years, ranging from 49 to 84 years, and most of them were men (63.8%). The most common concomitant diseases were hypertension (71.5%), diabetes (62.4%), IHD (37.6%), CKD (20%), heart failure (19.4%), and 81.2% suffered from ARDS. AKI occurred in 60.3% of patients, 38% were stage 1, 16.6% were stage 2, and 45.4% were stage 3. Approximately, 39% of patients required CRRT, out of which 76.2% were stage 3, which was significantly higher than the other stages (p<0.001). AKI patients suffered significantly from asthma and had lower levels of C-reactive protein (CRP), ferritin, lactate dehydrogenase (LDH), and blood urea nitrogen (BUN) and higher creatinine levels than patients without AKI (p<0.05 all). The overall mortality rate was 39.4%, and the mortality rate was significantly higher in patients with AKI than in patients without AKI (48.3% versus 25.9%; p<0.001).

**Conclusion**: AKI is common in adults admitted to the ICU with COVID-19 and is associated with an increased risk of death. Early detection of AKI and appropriate treatment can positively impact COVID-19 outcome. CRRT is the preferred dialysis method in critically ill ICU patients with AKI.

Key words: Acute kidney injury; COVID-19 pneumonia; intensive care unit; renal replacement therapy; Saudi Arabia.

**Correspondence:**Dr. Safwat Eldaboosy, Department of Pulmonary Medicine, Almoosa Specialist Hospital, box code 5098, Al Ahsa, Saudi Arabia, Tel. +966.590050316. E-mail: safwatchest@gmail.com

**Contributions:** All authors contributed to the conception and design of the work, reviewing the literature, data acquisition, manuscript writing, and revision. SE, AA, AF wrote the first draft of the manuscript and critically reviewed and revised the manuscript. All authors read, reviewed, and approved the final text and agreed to be accountable for all aspects of the work.

Conflict of interest: The authors declare that they have no competing interests, and all authors confirm accuracy.

**Ethics approval and consent to participate:** Approval was obtained from Almoosa Specialist Hospital Institutional Review Board (IRB protocol number: ARC -21.03.3).

Availability of data and material: All data obtained or analyzed as part of this study are available from the corresponding author on reasonable request.

Funding: This work was not funded by any organization, institution, or individual.

Acknowledgments: We would like to thank the research centers in our hospitals.



## Introduction

Coronavirus disease-19 (COVID-19) is associated with high morbidity and mortality worldwide and has overwhelmed many healthcare systems and economies. It is caused by a strain of severe acute respiratory syndrome coronavirus 2 (SARS-CoV-2) [1]. According to the World Health Organization (WHO), more than 590,000,000 people worldwide have been diagnosed with COVID-19 as of August 18, 2022, resulting in more than 6.4 million deaths [2]. About 30% of people infected with COVID-19 remain asymptomatic, 80% of symptomatic cases have mild and moderate disease (40% each) and 15% have severe disease requiring oxygen support. Approximately 5% develop critical COVID-19 illness characterized by pneumonia, acute respiratory failure, septic shock, and/or multiple organ dysfunction [2].

The manifestations of renal involvement in patients with COVID-19 may include proteinuria, hematuria, or acute kidney injury (AKI) and are associated with a high risk of adverse prognosis [3]. The main pathogenesis of COVID-19 associated renal disease is unknown and probably multifactorial and could be due to direct viral cytotoxic damage, the renin-angiotensin-aldosterone system (RAAS) imbalance, associated hyperinflammatory state due to re-released cytokines, microvascular injury and prothrombotic state, hypovolemia, potentially nephrotoxic agents, and nosocomial sepsis [4].

There are data on the temporal evolution of the incidence of COVID-19-associated AKI. Preliminary data suggest that AKI rates declined during the duration of the pandemic, although the reason for this evolution is unclear [5,6]. In a meta-analysis of approximately 13,000 mostly hospitalized patients, the incidence of AKI was 17 percent (range 0.5 to 80%), and about 5% of patients needed renal replacement therapy (RRT). Incidence appears to vary by geographic location and proportion of critically ill patients in each study [7].

In two large observational studies of more than 5,000 patients hospitalized with COVID-19, approximately 32 to 37 percent were found to have AKI. Out of the patients with AKI, about 50% had the mild disease (1.5- to 2-fold increase in serum creatinine) and the remainder had moderate or severe disease (more than a doubling of creatinine) [5,8]. Independent predictors of AKI included advanced age or maleness, obesity, essential hypertension, diabetes mellitus, cardiovascular disease, and low estimated glomerular filtration rate (eGFR), as well as higher interleukin-6 levels and need for mechanical ventilation or vasopressor therapy [9,10]. Continuous renal replacement therapy (CRRT) is the preferred treatment for AKI or end-stage renal disease (ESRD) in critically ill patients in the ICU, usually with hemodynamic instability, when dialysis is required (e.g., volume overload, electrolyte disturbances including hyperkalemia, acidosis, and complications of uremia) [11]. The aim of this study was to investigate the incidence, risk factors, and consequences of AKI in patients admitted to the intensive care unit with critically ill COVID-19 pneumonia.

## Methods

This study was part of a larger project investigating various clinical and epidemiological aspects of COVID-19 pneumonia in Saudi Arabia. It was a retrospective, multicenter study of 340 adult patients  $\geq$ 18 years of age admitted to the intensive care unit of a Saudi Almoosa Specialist Hospital-Al Ahsa, Obeid General Hospital-Hofuf, and King Khaled Hospital-Hail for confirmed COVID-19 pneumonia. This study was conducted from May 2021 to May 2022. Ethical approval was obtained from the Institutional

Review Board of our hospitals (Almoosa Academic Affairs: IRB log No: ARC -21.02.02).

### **Inclusion criteria**

Adult patients ( $\geq$ 18 years) with COVID-19 pneumonia and hypoxemia (positive SARS-CoV-2 real-time polymerase chain reaction (PCR) nasopharyngeal swabs or respiratory secretions) were admitted to the ICU.

#### **Exclusion criteria**

Patients with COVID-19 pneumonia who did not not fulfill the criteria of ICU admission, patients admitted to the ICU for reasons other than COVID-19 pneumonia, patients who had COVID-19 infection but were not hospitalized, and pediatric COVID-19 patients were excluded. Patients with ESRD and kidney transplant patients were also excluded.

### Data collections and outcomes

All the cases were diagnosed and treated according to the Saudi Ministry of Health protocol for confirmed cases of COVID-19 infection [12].

The following data were collected (within 24 h of hospitalization): age, sex, body mass index (BMI), comorbidities (such as hypertension, asthma, type 2 diabetes, and other concomitant diseases), patient symptoms, and general and local examination findings, including vital signs. The following examination results were obtained from the electronic medical record: identification of SARS-CoV-2 virus by nasopharyngeal swab using PCR, chest radiograph (portable), oxygen saturation with a pulse oximeter, arterial blood gases, complete blood count (CBC), complete metabolic panel (serum sodium, potassium, and magnesium), serum ferritin, D-dimer, lactate dehydrogenase (LDH), C-reactive protein (CRP), renal function test (urea and creatinine), liver function test, procalcitonin, troponin, and electrocardiogram (ECG). ICU course, including length of stay, need for ventilatory support (noninvasive or invasive ventilation), need for vasopressors, prophylactic anticoagulants, antibiotics, systemic steroids, anti-interleukin 6 (tocilizumab), empiric antibiotics (per local protocol), and need for CRRT and AKI outcome and outcomes either mortality or discharge to ward. Acute kidney injury was defined as per the Kidney Disease Improving Global Outcome (KDIGO) criteria [13].

Baseline creatinine is the last serum creatinine in the last 7-365 days before admission. In patients in whom no previous serum creatinine has been measured, the serum creatinine at admission is considered the baseline creatinine. Estimated glomerular filtration rate (eGFR) is calculated based on a Modification of Diet in Renal Disease (MDRD) equation. Renal recovery is defined as a decrease in serum creatinine of more than 50%.

### Statistical analysis

We used the SPSS program for Windows (IBM SPSS Statistics V 25.0, IBM Corp., Armonk, NY, USA). Mean  $\pm$  SD and median and interquartile range (IQR) were used for quantitative variables, whereas frequency and percentage were used for qualitative variables. Chi-square or Fischer exact tests were used to assess differences in frequencies of qualitative variables. Independent-samples *t*-test was used to evaluate the differences in the means of the quantitative variables, whereas the Mann-Whitney U test was used for nonparametric statistics. Logistic regression analysis was used with odds ratios (OR) and 95% confidence intervals (CI) to predict factors associated with COVID-19 mortality. Only significant independent variables from the univariate analysis were included in the logistic analysis. Statistical methods were reviewed, using a significance level of p<0.05 (double-tailed).



# Results

The study included 340 patients admitted to the ICU with COVID-19. Their mean age was 66.7±13.4 years and ranged from 49 to 84 years; most of them were men (63.8%). The most common concomitant diseases were hypertension (71.5%), diabetes (62.4%), IHD (37.6%), CKD (20%), heart failure (19.4%), and 81.2% suffered from ARDS. Associated clinical and laboratory parameters showed elevated mean HR (97.5±19.3) and RR (27.7±4.2), elevated median CRP (81 [36-136]), ferritin (964 [792-1642]), LDH (652 [452-895]), D-dimer (2.9 [2.5-3.4]), creatinine (89 [73.3-142]), and lower median PaO<sub>2</sub>/FIO<sub>2</sub> ratio (62 [53-112]). The majority (91.8%) received steroid therapy and 43.5% received vasopressors. The mortality rate was 134 patients (39.4%) (Table 1). AKI occurred in 205 patients (60.3%), out of whom 78 (38%) were stage 1, 34 (16.6%) were stage 2, and 93 (45.4%) were stage 3. Eighty of 205 patients (39%) required CRRT, including 15 of 80 patients (18.8%) in stage 1, 4 patients (5%) in stage 2, and 61 patients (76.2%) in stage 3, which was significantly higher than the other stages (p<0.001) (Table 2). More than three-quarters (76.8%) of the patients received mechanical ventilation (MV), 14.6% received oxygen therapy with a high-flow nasal cannula (HFNC), and 4.4% received oxygen therapy with a nasal cannula (NC) (Figure 1). Patients with AKI suffered significantly from asthma, required CRRT, and had lower median CRP, ferritin, LDH, and BUN levels, and higher median creatinine levels than patients without AKI (p<0.05 all) (Table 3). The mortality rate was significantly higher in patients with AKI than in patients without AKI (48.3% vs 25.9%; p<0.001). Overall, 106 out of 205 patients (51.7%) with AKI were discharged; complete improvement occurred in 97 patients (47.3%), and 9 patients (4.4%) required regular dialysis because of ESRD (Figure 2).

## Discussion

The pandemic COVID-19 causes numerous cases of illness and death and has overwhelmed health services and disrupted normal life and the economy. COVID-19 is characterized by a systemic inflammatory response and an increased risk of respiratory failure and AKI [1,2]. AKI is one of the most common complications of COVID-19 in hospitalized patients and is associated with a highly unfavorable outcome [5].

In this study, AKI occurred in 60.3% of patients admitted to the ICU for critical COVID-19 pneumonia. This is consistent with many studies from the United States, in which 60-80% of critical COVID-19 patients were found to have AKI [14,15]. Other studies from the USA involving COVID-19 cases admitted to the ICU, showed that more than 60% (61-78%) of them had AKI [16,17]. The incidence of AKI reached 68% in critically ill COVID-19 patients admitted to the intensive care unit. In a large cohort study in New York City [18]. Alenezi et al. conducted a systematic review and meta-analysis (out of 618 studies identified and reviewed, 31 studies met inclusion criteria), and the incidence of AKI was 50% in eight studies that included only COVID-19 patients admitted to the ICU (n=1,540) [19]. Our results on the incidence of AKI were lower than those of other studies. Schaubroeck et al. performed a multicenter cohort analysis of AKI in critically ill patients of COVID-19 in Belgium (from seven large hospitals) and found a high rate of AKI (85.1% of 1,286 cases) [20]. Lumlertgul et al. reported that AKI occurred in 76% of critical COVID-19 patients [21]. de Almeida et al. observed an increased incidence of AKI > 70%, with more than half of these patients meeting KDIGO 3 criteria within 7 days of hospitalization [22]. Our results on AKI rates in critically ill COVID-19 cases in the ICU were higher than in many

 Table 1. General, clinical, laboratory and outcome characteristics of the studied patients.

Age (years)       66.7±13.4         Sex       63.8         Male       217       63.8         Female       123       36.2         BMI (kg/m²)       26.9±7.0         HTN       243       71.5         DM2       212       62.4         Asthma       19       5.6         IHD       128       37.6         CHF       66       19.4         CKD       68       20.0         HR (beats/min)       97.5±19.3       7         Temperature (°C)       37.8±1.01       8         RK (breaths/min)       27.7±4.2       6         GCS       14.5±0.78       7         PaO2 (mmHg)       59 (50.3-65)       7         CRP       81 (36-136)       7         Ferritin (ng/mL)       964 (792-1642)       7         LDH (IU/L)       652 (452-895)       7         D-dimer       2.9 (2.5-3.4)       7         HB (gm/dl)       11.3 (8.2-15.4)       7         Creatinine (mg/dl)       89 (73.3-142)       8         BUN (mg/dl)       12.4 (9.4-18.8)       8         PaO_/FIO <sub>2</sub> ratio       62 (53-112)       <300         <300 </th <th>Variables</th> <th>n=340</th> <th></th> <th>(%)</th>	Variables	n=340		(%)
Male Female         217         63.8 36.2           BMI (kg/m²)         26.9±7.0           HTN         243         71.5           DM2         212         62.4           Asthma         19         5.6           IHD         128         37.6           CHF         66         19.4           CKD         68         20.0           HR (beats/min)         97.5±19.3         Temperature (°C)           Temperature (°C)         37.8±1.01         RR           RR (breaths/min)         27.7±4.2         GCS           GCS         14.5±0.78         PaO2 (mmHg)           S9 (50.3-65)         CRP         81 (36-136)           Ferritin (ng/mL)         964 (792-1642)         LDH (1U/L)           LDH (1U/L)         652 (452-895)         D-dimer           LDH (1U/L)         652 (452-895)         D-dimer           VBCS (cells/m³)         11.3 (82-15.4)         Creatinine (mg/dl)           RUN (mg/dl)         12.4 (9.4+18.8)         PaO_/FIO2 ratio           <300	Age (years)		66.7±13.4	
Female         123         36.2           BMI (kg/m²)         26.9±7.0           HTN         243         71.5           DM2         212         62.4           Asthma         19         5.6           IHD         128         37.6           CHF         66         19.4           CKD         68         20.0           HR (beats/min)         97.5±19.3         Temperature (°C)           RR (breaths/min)         27.7±4.2         GCS           GCS         14.5±0.78         PaO2 (mmHg)         59 (50.3-65)           CRP         81 (36-136)         Ferritin (ng/mL)         964 (792-1642)           LDH (IU/L)         652 (452-895)         D-dimer         2.9 (2.5-3.4)           HB (gm/dl)         11.4±2.2         WBCs (cells/mm³)         11.3 (8.2-15.4)           Creatinine (mg/dl)         89 (73.3-142)         BUN (mg/dl)         12.4 (9.4-18.8)           PAO_YFIO_ratio         62 (53-112)         <300	Sex			
BMI (kg/m²)         26.9±7.0           HTN         243         71.5           DM2         212         62.4           Asthma         19         5.6           IHD         128         37.6           CHF         66         19.4           CKD         68         20.0           HR (beats/min)         97.5±19.3         Temperature (°C)           Temperature (°C)         37.8±1.01         RR           RR (breaths/min)         27.7±4.2         GCS           GCS         14.5±0.78         PaO2 (mmHg)           PaO2 (mmHg)         59 (50.3-65)         CRP           RI (36-136)         Ferritin (ng/mL)         964 (792-1642)           LDH (IU/L)         652 (452-895)         D-dimer           D-dimer         2.9 (2.5-3.4)         HB (gm/dl)           HB (gm/dl)         11.4±2.2         WBCs (cells/mm³)           UN (mg/dl)         12.4 (9.4-18.8)         PaO2 (53-112)           <300				
HTN       243       71.5         DM2       212       62.4         Asthma       19       5.6         IHD       128       37.6         CHF       66       19.4         CKD       68       20.0         HR (beats/min)       97.5±19.3       Temperature (°C)         Temperature (°C)       37.8±1.01       RR (breaths/min)         RR (breaths/min)       27.7±4.2       GCS         GCS       14.5±0.78       PaO2 (mmHg)         PaO2 (mmHg)       59 (50.3-65)       CRP         RP       81 (36-136)       Ferritin (ng/mL)       964 (792-1642)         LDH (IU/L)       652 (452-895)       D-dimer       2.9 (2.5-3.4)         HB (gm/dl)       11.4±2.2       WBCs (cells/mm³)       11.3 (8.2-15.4)         Creatinine (mg/dl)       89 (73.3-142)       BUN (mg/dl)       12.4 (9.4-18.8)         PaO2/FIO2 ratio       62 (53-112)       <300		123		36.2
DM2         212         62.4           Asthma         19         5.6           IHD         128         37.6           CHF         66         19.4           CKD         68         20.0           HR (beats/min)         97.5±19.3         Temperature (°C)           Temperature (°C)         37.8±1.01         RR (breaths/min)           RR (breaths/min)         27.7±4.2         GCS           GCS         14.5±0.78         PaO2 (mmHg)           S9 (50.3-65)         CRP         81 (36-136)           Ferritin (ng/mL)         964 (792-1642)         LDH (IU/L)           LDH (IU/L)         652 (452-895)         D-dimer           LDH (IU/L)         652 (452-895)         D-dimer           VBCs (cells/mm³)         11.3 (8.2-15.4)         Creatinine (mg/dl)           Bl (gm/dl)         11.4 ± 2.2         WBCs (cells/mm³)         11.3 (8.2-15.4)           Creatinine (mg/dl)         89 (73.3-142)         BUN (mg/dl)         12.4 (9.4-18.8)           PaO_yFIO_2 ratio         62 (53-112)         <300			$26.9 \pm 7.0$	
Asthma         19         5.6           IHD         128         37.6           CHF         66         19.4           CKD         68         20.0           HR (beats/min)         97.5±19.3         Temperature (°C)           Temperature (°C)         37.8±1.01         RR (breaths/min)           QCS         14.5±0.78         PaO2 (mmHg)           PaO2 (mmHg)         59 (50.3-65)         CRP           RP         81 (36-136)         Ferritin (ng/mL)           Pervisin (ng/mL)         964 (792-1642)         LDH (IU/L)           Chier         2.9 (2.5-3.4)           HB (gm/dl)         11.4±2.2         WBCS (cells/mm³)           D-dimer         2.9 (2.5-3.4)           HB (gm/dl)         11.4±2.2           WBCS (cells/mm³)         11.3 (8.2-15.4)           Creatinine (mg/dl)         89 (73.3-142)           BUN (mg/dl)         12.4 (9.4-18.8)           PaO2/FIO2 ratio         62 (53-112)           <300				
IHD       128       37.6         CHF       66       19.4         CKD       68       20.0         HR (beats/min)       97.5 $\pm$ 19.3       Temperature (°C)         Temperature (°C)       37.8 $\pm$ 1.01       RR (breaths/min)         RR (breaths/min)       27.7 $\pm$ 4.2       GCS         GCS       14.5 $\pm$ 0.78       PaO <sub>2</sub> (mmHg)         PaO <sub>2</sub> (mmHg)       59 (50.3-65)       CRP         RP       81 (36-136)       Ferritin (ng/mL)       964 (792-1642)         LDH (IU/L)       652 (452-895)       D-dimer       2.9 (2.5-3.4)         HB (gm/dl)       11.4 $\pm$ 2.2       WBCs (cells/mm <sup>3</sup> )       11.3 (8.2-15.4)         Creatinine (mg/dl)       89 (73.3-142)       BUN (mg/dl)       12.4 (9.4-18.8)         PaO <sub>2</sub> /FIO <sub>2</sub> ratio       62 (53-112)       <300	DM2	212		62.4
CHF6619.4CKD6820.0HR (beats/min) $97.5 \pm 19.3$ Temperature (°C) $37.8 \pm 1.01$ RR (breaths/min) $27.7 \pm 4.2$ GCS $14.5 \pm 0.78$ PaO <sub>2</sub> (mmHg)59 (50.3-65)CRP81 (36-136)Ferritin (ng/mL)964 (792-1642)LDH (IU/L)652 (452-895)D-dimer2.9 (2.5-3.4)HB (gm/dl)11.4 $\pm 2.2$ WBCs (cells/mm <sup>3</sup> )11.3 (8.2-15.4)Creatinine (mg/dl)89 (73.3-142)BUN (mg/dl)12.4 (9.4-18.8)PaO <sub>2</sub> /FIO <sub>2</sub> ratio62 (53-112)<300	Asthma	19		5.6
CKD         68         20.0           HR (beats/min)         97.5±19.3           Temperature (°C)         37.8±1.01           RR (breaths/min)         27.7±4.2           GCS         14.5±0.78           PaO <sub>2</sub> (mmHg)         59 (50.3-65)           CRP         81 (36-136)           Ferritin (ng/mL)         964 (792-1642)           LDH (IU/L)         652 (452-895)           D-dimer         2.9 (2.5-3.4)           HB (gm/dl)         11.4±2.2           WBCs (cells/mm <sup>3</sup> )         11.3 (8.2-15.4)           Creatinine (mg/dl)         89 (73.3-142)           BUN (mg/dl)         12.4 (9.4-18.8)           PaO <sub>2</sub> /FIO <sub>2</sub> ratio         62 (53-112)           <300	IHD	128		37.6
HR (beats/min)       97.5±19.3         Temperature (°C) $37.8\pm1.01$ RR (breaths/min) $27.7\pm4.2$ GCS $14.5\pm0.78$ PaO <sub>2</sub> (mmHg)       59 (50.3-65)         CRP $81$ (36-136)         Ferritin (ng/mL)       964 (792-1642)         LDH (IU/L)       652 (452-895)         D-dimer       2.9 (2.5-3.4)         HB (gm/dl) $11.4\pm2.2$ WBCs (cells/mm <sup>3</sup> ) $11.3$ (8.2-15.4)         Creatinine (mg/dl)       89 (73.3-142)         BUN (mg/dl) $12.4$ (9.4-18.8)         PaO <sub>2</sub> /FIO <sub>2</sub> ratio       62 (53-112)         <300	CHF	66		19.4
Temperature (°C) $37.8\pm1.01$ RR (breaths/min) $27.7\pm4.2$ GCS $14.5\pm0.78$ PaO <sub>2</sub> (mmHg) $59$ (50.3-65)         CRP $81$ (36-136)         Ferritin (ng/mL)       964 (792-1642)         LDH (IU/L) $652$ (452-895)         D-dimer $2.9$ (2.5-3.4)         HB (gm/dl) $11.4\pm2.2$ WBCs (cells/mm <sup>3</sup> ) $11.3$ (8.2-15.4)         Creatinine (mg/dl) $89$ (73.3-142)         BUN (mg/dl) $12.4$ (9.4-18.8)         PaO_/FIO <sub>2</sub> ratio $62$ (53-112)         <300	CKD	68		20.0
RR (breaths/min) $27.7\pm4.2$ GCS $14.5\pm0.78$ PaO <sub>2</sub> (mmHg) $59$ (50.3-65)         CRP $81$ (36-136)         Ferritin (ng/mL)       964 (792-1642)         LDH (IU/L) $652$ (452-895)         D-dimer $2.9$ (2.5-3.4)         HB (gm/dl) $11.4\pm2.2$ WBCs (cells/mm <sup>3</sup> ) $11.3$ (8.2-15.4)         Creatinine (mg/dl) $89$ (73.3-142)         BUN (mg/dl) $12.4$ (9.4-18.8)         PaO <sub>2</sub> /FIO <sub>2</sub> ratio $62$ (53-112)         <300	HR (beats/min)		$97.5 \pm 19.3$	
GCS $14.5\pm0.78$ PaO <sub>2</sub> (mmHg)       59 (50.3-65)         CRP       81 (36-136)         Ferritin (ng/mL)       964 (792-1642)         LDH (IU/L)       652 (452-895)         D-dimer       2.9 (2.5-3.4)         HB (gm/dl)       11.4 $\pm 2.2$ WBCs (cells/mm <sup>3</sup> )       11.3 (8.2-15.4)         Creatinine (mg/dl)       89 (73.3-142)         BUN (mg/dl)       12.4 (9.4-18.8)         PaO <sub>2</sub> /FIO <sub>2</sub> ratio       62 (53-112)         <300	Temperature (°C)		$37.8 \pm 1.01$	
PaO <sub>2</sub> (mmHg)       59 (50.3-65)         CRP       81 (36-136)         Ferritin (ng/mL)       964 (792-1642)         LDH (IU/L)       652 (452-895)         D-dimer       2.9 (2.5-3.4)         HB (gm/dl)       11.4 $\pm$ 2.2         WBCs (cells/mm <sup>3</sup> )       11.3 (8.2-15.4)         Creatinine (mg/dl)       89 (73.3-142)         BUN (mg/dl)       12.4 (9.4-18.8)         PaO <sub>2</sub> /FIO <sub>2</sub> ratio       62 (53-112)         <300	RR (breaths/min)	17	$27.7 \pm 4.2$	
CRP         81 (36-136)           Ferritin (ng/mL)         964 (792-1642)           LDH (IU/L)         652 (452-895)           D-dimer         2.9 (2.5-3.4)           HB (gm/dl)         11.4±2.2           WBCs (cells/mm <sup>3</sup> )         11.3 (8.2-15.4)           Creatinine (mg/dl)         89 (73.3-142)           BUN (mg/dl)         12.4 (9.4-18.8)           PaO <sub>2</sub> /FIO <sub>2</sub> ratio         62 (53-112)           <300	GCS	$\overline{\mathbf{V}}$	$14.5 \pm 0.78$	
Ferritin (ng/mL)       964 (792-1642)         LDH (IU/L)       652 (452-895)         D-dimer       2.9 (2.5-3.4)         HB (gm/dl)       11.4±2.2         WBCs (cells/mm <sup>3</sup> )       11.3 (8.2-15.4)         Creatinine (mg/dl)       89 (73.3-142)         BUN (mg/dl)       12.4 (9.4-18.8)         PaO_/FIO2 ratio       62 (53-112)         <300	$PaO_2$ (mmHg)		59 (50.3-65)	
LDH (IU/L)       652 (452-895)         D-dimer       2.9 (2.5-3.4)         HB (gm/dl)       11.4±2.2         WBCs (cells/mm <sup>3</sup> )       11.3 (8.2-15.4)         Creatinine (mg/dl)       89 (73.3-142)         BUN (mg/dl)       12.4 (9.4-18.8)         PaO <sub>2</sub> /FIO <sub>2</sub> ratio       62 (53-112)         <300	CRP		81 (36-136)	
D-dimer       2.9 (2.5-3.4)         HB (gm/dl) $11.4\pm 2.2$ WBCs (cells/mm <sup>3</sup> ) $11.3$ (8.2-15.4)         Creatinine (mg/dl)       89 (73.3-142)         BUN (mg/dl) $12.4$ (9.4-18.8)         PaO_/FIO2 ratio       62 (53-112)         <300	Ferritin (ng/mL)		964 (792-1642)	
HB (gm/dl) $11.4\pm 2.2$ WBCs (cells/mm <sup>3</sup> ) $11.3$ (8.2-15.4)         Creatinine (mg/dl)       89 (73.3-142)         BUN (mg/dl)       12.4 (9.4-18.8)         PaO <sub>2</sub> /FIO <sub>2</sub> ratio       62 (53-112)         <300	LDH (IU/L)	. ,		
WBCs (cells/mm <sup>3</sup> )       11.3 (8.2-15.4)         Creatinine (mg/dl)       89 (73.3-142)         BUN (mg/dl)       12.4 (9.4-18.8)         PaO <sub>2</sub> /FIO <sub>2</sub> ratio       62 (53-112)         <300	D-dimer			
Creatinine (mg/dl)       89 (73.3-142)         BUN (mg/dl)       12.4 (9.4-18.8)         PaO <sub>2</sub> /FIO <sub>2</sub> ratio       62 (53-112) $<300$ 276       81.2 $>300$ 64       18.8         Need for vasopressors       148       43.5         ARDS       276       81.2         Mild (200-300)       Moderate (100-200)       81.2         Steroid therapy       312       91.8         Hospital stays (days)       11 (4-18)       0utcome         Discharged       206       60.6	HB (gm/dl)	× /		
BUN (mg/dl)       12.4 (9.4-18.8)         PaO <sub>2</sub> /FIO <sub>2</sub> ratio       62 (53-112)         <300	WBCs (cells/mm <sup>3</sup> )	11.3 (8.2-15.4)		
PaO <sub>2</sub> /FIO <sub>2</sub> ratio       62 (53-112)         <300	Creatinine (mg/dl)	· · · ·		
<300	BUN (mg/dl)			
>300         64         18.8           Need for vasopressors         148         43.5           ARDS         276         81.2           Mild (200-300)         Moderate (100-200)         8           Severe <100	PaO <sub>2</sub> /FIO <sub>2</sub> ratio		62 (53-112)	
Need for vasopressors14843.5ARDS27681.2Mild (200-300) Moderate (100-200) Severe <100				
ARDS Mild (200-300) Moderate (100-200) Severe <10027681.2Steroid therapy31291.8Hospital stays (days)11 (4-18)Outcome Discharged20660.6				
Mild (200-300) Moderate (100-200) Severe <10091.8Steroid therapy31291.8Hospital stays (days)11 (4-18)Outcome Discharged20660.6				
Moderate (100-200) Severe <100Steroid therapy31291.8Hospital stays (days)11 (4-18)Outcome Discharged20660.6		276		81.2
Severe <100Steroid therapy31291.8Hospital stays (days)11 (4-18)Outcome Discharged20660.6				
Hospital stays (days) 11 (4-18) Outcome Discharged 206 60.6	· · · · · · · · · · · · · · · · · · ·			
Hospital stays (days) 11 (4-18) Outcome Discharged 206 60.6	Steroid therapy	312		91.8
Outcome Discharged 206 60.6				
Discharged 206 60.6				
Death 134 39.4	Discharged			
	Death	134		39.4

Qualitative variables present as number and percent; quantitative variables present as mean  $\pm$ SD or as median (IQR).

Table 2. Renal impairment among the studied patients.

Variables	n=340	(%)
AKI	205	60.3
AKI stage		
Stage 1	78/205	38.0
Stage 2	34/205	16.6
Stage 3	93/205	45.4
CRRT	80/205	39.0
CRRT <i>vs</i> AKI stage		
Stage 1	15/80	18.8
Stage 2	4/80	5.0
Stage 3	61/80	76.2
p	<0.001*	

Qualitative variables are present as numbers and percent and analyzed by Chi-square test; \*significant.



other studies. The incidence of AKI was reported to be 36% in hospitalized patients with COVID-19, in a study from Saudi Arabia conducted by Farooqui et al. [23]. In another Saudi Arabian study examining critically ill COVID-19 patients in a multicenter study, AKI occurred in 46.8% of cases [24]. In contrast, AKI is common

in critically ill patients with COVID-19 and affects approximately 20-40% of patients admitted to the ICU [25, 26]. About 29% of patients admitted to the intensive care unit have AKI, and the figure is as high as 78% for patients requiring intubation [18]. Several studies have reported that more than 30-50% of hospitalized

Table 3. Comparing different variables among COVID-19 patients w	vith and without acute kidney injury.
--	---------------------------------------

Variables	Patients with AKI, n=205 (%)	Patients without AKI, n=135 (%)	р
Age	67.5±11.8	65.4±15.4	0.169
Sex			
Male	128 (62.4)	89 (65.9)	0.565
Female	77 (37.6)	46 (34.1)	
BMI	$27.3 \pm 6.8$	$26.4 \pm 7.1$	0.275
HTN	154 (75.1)	89 (65.9)	0.085
DM2	133 (64.9)	79 (58.5)	0.254
Asthma	19 (9.3)	2 (1.5)	0.003*
IHD	83 (40.5)	45 (33.3)	0.209
CHF	42 (20.5)	24 (17.8)	0.577
CKD	42 (20.5)	26 (19.3)	0.890
HR (beats/min)	$98.4{\pm}20.2$	96.1±17.8	0.286
CRRT	80 (39.0)	0 (0.0)	<0.001*
Need for vasopressors	84 (41.0)	64 (47.4)	0.264
ARDS	164 (80.0)	112 (83.0)	0.571
Temp (°C)	$37.9 \pm 1.0$	37.9±1.1	0.254
RR (breaths/min)	$27.9 \pm 4.4$	27.3±3.9	0.265
GCS	$14.55 \pm 0.78$	$14.5 \pm 0.78$	0.621
Need for MV	48 (23.4)	34 (25.2)	0.796
PaO <sub>2</sub> (mmHg)	62 (51-67)	55 (50-64)	0.784
PaO <sub>2</sub> /FIO <sub>2</sub> ratio	64 (56-125)	59 (52-66)	0.087
CRP	68 (34-124)	101 (43-164)	0.006*
Ferritin (ng/mL)	900 (735-1348)	1261 (854-2354)	0.040*
LDH (IU/L)	635 (364-831)	745 (521-952)	0.014*
D-dimer	2.9 (2.5-3.4)	2.9 (2.55-3.36)	0.137
HB (gm/dl)	11.5±2.2	11.3±2.4	0.359
WBCs (cells/mm³)	9.6 (8.2-14.8)	12.6 (8.2-16.4)	0.765
Creatinine (mg/dl)	95 (74-133)	88.5 (72-195)	0.004*
BUN (mg/dl)	12 (8.4-18.2)	14 (11-20.4)	<0.001*

Qualitative variables are present as numbers and percent an analyzed by Fisher exact test; quantitative variables present as mean ±SD or as median (IQR) and analyzed by independent samples t-test or Mann-Whitney U tests; \*significant.

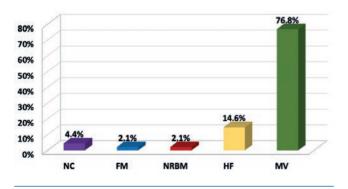
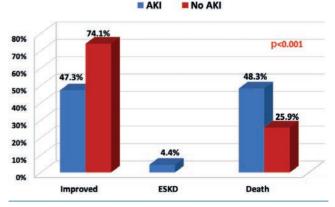


Figure 1. Type of  $O_2$  therapy among the studied patients. NC = nasal cannula; FM = face mask; NRBM = non-rebreather mask function; HF = high-flow; MV = mechanical ventilation.



AKI

Figure 2. Outcome of COVID-19 pneumonic patients with and without acute kidney injury. ESKD = End Stage Kidney Disease.



patients with COVID-19 develop AKI, with a higher proportion of patients requiring ICU admission [27,28]. Yang et al. performed a meta-analysis of 51 studies, and the incidence of AKI in ICU patients was 39% and may reach 42% in deceased patients, and 16.3% of ICU patients required CRRT [29]. Jewell et al. analyzed data from hospitalized adults with COVID-19 in two London hospitals and reported that 39% developed AKI (51% in stage 1 and 49% in stages 2 and 3) [30]. Oweis et al. conducted a study of AKI in hospitalized patients with COVID-19 from Jordan. The incidence of AKI was 25.3%, and most patients were at stage 1 [31]. CRRT is the preferred treatment for AKI in critically ill patients in the ICU because it provides hemodynamic stability and large fluid removal. Selection of this modality should be based on local experience but is expensive and time-consuming. In this study, 39% of AKI in critically ill COVID-19 pneumonia cases required CRRT, and most of them (81.2%) were in stages 2 and 3. Out of the patients with AKI, 81.2% had ARDS, 43.5% of them required vasopressors, and 76.8% of them required mechanical ventilation. The use of CRRT in our critical patients with COVID-19 was consistent with previous epidemiological studies in which the use of CRRT was required in approximately 44% of patients [16]. Many studies reported that AKI is a common complication in critical COVID-19 cases and occurred in 25-76% of cases, with 5-44% of them requiring CRRT [32,33]. About 35-50.9% of patients with AKI in the ICU associated with COVID-19 pneumonia required CRRT in some studies from the United States [9,15]. Many studies report that 30-40% of COVID-19 infected patients in the ICU require RRT [33,34]. Cummings et al. reported that out of 257 critically ill patients with COVID-19, 31% received RRT [27]. In one study it was reported that up to 20% of patients underwent renal replacement therapy (KRT) [5]. This is in contrast to a study from India and Pakistan, which reported that CRRT was required in 22.9% of cases [35]. The incidence of AKI was 43.7%, and 18.2% of patients underwent CRRT, and in this group, 90-day in-hospital mortality was 45.1%. A study by Eriksson et al. on CRRT in ICU patients with COVID-19 [36]. In a study from Saudi Arabia, CRRT was required in 18.9% of cases with AKI associated with COVID-19 pneumonia [24]. This is in contrast also to other studies reporting that CRRT was required in 4% to 23% of patients with AKI in critically ill COVID-19 patients [8,37].

In this study, the mortality rate was significantly higher in patients with AKI (48.3%) than in patients without AKI (25.9%), which is often the case in several studies. According to Jewell et al., the mortality rate was significantly higher in patients WHO with AKI (44.4%) than in patients WHO without AKI (17.3%) [30]. Alenezi et al. performed a scientific review and meta-analysis of the incidence and risk factors of AKI in COVID-19 patients with and without white lung and found that mortality was 38.7% once the studies considered only COVID-19 patients admitted to the department [19]. In a study from Asia conducted by Farooqui et al., the presence of AKI was associated with a higher 30-day mortality of 40.7%, compared with 3.7% in patients without AKI [23]. In another study from an Asian country, AKI was found to be much more common in patients who died WHO within thirty days on the ward (74.7%) than in patients who survived WHO on the ward (26.2%) [24]. Eriksson et al. performed a study of CRRT in patients with COVID-19 and found that the mortality rate in this group was 45.1% [36]. Our results were lower than those of many other studies. Fominskiy et al. found that mortality was higher in patients with AKI (52.9%) than in patients without AKI (38.9%) [37]. Several studies found that overall hospital mortality in patients with COVID-19 and AKI was 66.2% [3,39], and in several studies, it ranged from 60-80% within the AKI-RRT cluster [40, 41]. According to Oweis et al., 75% of patients with AKI died within the unit [31]. Compared with patients without AKI, mortality was higher in patients with AKI (32.5% versus 10.4%), and

39.2% of patients with AKI did not recover from urinary organ performance by the end of the follow up period or after ninety days [42]. On the other hand, other studies have not shown AKI to be a risk factor for COVID-19 deaths [30,43].

In this study, the mean age of patients with AKI was  $67.5\pm11.8$ years, 62.4% were male, the mean BMI was 27.3±6.8, and the major comorbidities were HTN 75.1%, DM2 64.9%, IHD 40.5%, CHF 20.5%, and CKD 20.5% without significant differences when compared to the group without AKI. These findings were comparable with other studies that showed variable values and percentages. In a study from Saudi Arabia, the mean age was 65 years, 74.5% of patients were male, and the most common comorbidities were DM2 (57.7%), HTN (53.6%), and dyslipidemia (22.7%) [23]. In another study from Saudi Arabia, the mean age was 66 years, 56.7% of participants were male, and the most common comorbidities were DM2 (70.2%), HTN (73.9%), heart failure (21.4%), CHF (16.8%), and a mean BMI of 29.7 [24] also, without significant differences between both groups with and without AKI. In a multicenter ICU study involving 5,866 COVID-19 patients from 55 hospitals in Spain, the main age was 63 years, most of them were men (70.4%), and the most common concomitant diseases were HTN (50.4%), obesity (35.5%), and DM2 (24.9%) [43]. In a systematic review of patients with COVID-19 and AKI, the most common comorbidities were HTN 61.4%, hyperlipidemia 57.1%, DM2 40%, and CKD 22.2 % [44].

In contrast, other studies reported that independent significant predictors of AKI included being older, black American, or male; being overweight; having diabetes; having HTN; having the cardiovascular disease; having a low eGFR or higher interleukin-6 levels; or requiring mechanical ventilation or vasopressor medications [9,10]. Most biomarkers that showed a significant correlation with AKI have been established in relation to the severity of COVID-19, including d-dimer, LDH, neutrophil and leukocyte counts, troponin-I, and CRP [44-47]. In our study, inflammatory markers were high in both groups (AKI and without AKI). Oweis *et al.* reported that comorbidities such as HTN and diabetes, as well as previous renal disease and increasing age, increase the risk of AKI in patients with COVID-19 but not significantly in terms of the degree of inflammation and the increase in CRP [31].

The differences between our results and the others can be explained by many factors: inclusion criteria, quality of the health care system, differences in referral policies, duration of follow up, the experience of the first centers affected by COVID-19 outbreaks, and differences in population characteristics or prevalence of comorbidities.

#### **Study limitations**

First: the retrospective nature of the study raises the possibility that differences in the quality of care may affect patient recovery. Second, we used KIGO rather than GMFR to define AKI. Third, some data were not collected, such as urine output, and the mortality rate was used only as in-hospital mortality without post-discharge follow up. In addition, the relation between the percentage of lung parenchyma affected by COVID-19 and the severity of renal involvement was not investigated in the current study. Further prospective studies are recommended to cover these issues.

# Conclusion

AKI is common in adults admitted to the intensive care unit with COVID-19 and is associated with an increased risk of death. Early recognition of AKI and appropriate treatment can have a positive impact on the outcome of COVID-19. CRRT is the preferred dialysis method in critically ill ICU patients with AKI.



## Abbreviations

AKI: acute kidney injury; ARDS: acute respiratory distress syndrome; BMI: body mass index; BUN: blood urea nitrogen; CBC: Complete blood count; CHF: congestive heart failure; CI: confidence intervals: CKD: chronic kidney disease: COVID-19: coronavirus disease 19: CRP: C reactive protein; CRRT: continuous replacement therapy; DM2: type 2 diabetes mellitus; ECG: electrocardiogram; FIO2: fractionated inspired oxygen; GCS: Glasgow coma scale; GMFR: glomerular filtration rate; HB: hemoglobin; HR: heart rate; HTN: hypertension; ICU: intensive care unit; IHD: ischemic heart disease; IQR: interquartile range, KDIGO: Kidney Disease Improving Global Outcome; LDH: lactate dehydrogenase; MV: mechanical ventilation; OR: odds ratios; PaO<sub>2</sub>: partial pressure of oxygen tension; RAAS: renin-angiotensin-aldosterone system; RR: respirator rate; RRT: renal replacement therapy; SARS: severe acute respiratory distress; WBCs: white blood cells; WHO: World Health Organization.

## References

- Cascella M, Rajnik M, Aleem A, Dulebohn SC, Di Napoli R. Features, evaluation, and treatment of coronavirus (COVID-19). In: StatPearls [Internet]. Treasure Island: StatPearls Publishing; 2022.
- World Health Organization. Clinical management of COVID-19: Living guideline, 15 September 2022. Available from: https://www.who.int/publications/i/item/WHO-2019-nCoV-Clinical-2022.2
- Cheng Y, Luo R, Wang K, Zhang M, Wang Z, Dong L, et al. Kidney disease is associated with in-hospital death of patients with COVID-19. Kidney Int 2020;97:829-38.
- 4. Gabarre P, Dumas G, Dupont T, Darmon M, Azoulay E, Zafrani L. Acute kidney injury in critically ill patients with COVID-19. Intensive Care Med 2020;46:1339-48.
- Bowe B, Cai M, Xie Y, Gibson AK, Maddukuri G, Al-Aly Z. Acute kidney injury in a national cohort of hospitalized US veterans with COVID-19. Clin J Am Soc Nephrol 2020;16:14-25.
- Charytan DM, Parnia S, Khatri M, Petrilli CM, Jones S, Benstein J, et al. Decreasing incidence of acute kidney injury in patients with COVID-19 critical illness in New York City. Kidney Int Rep 2021;6:916-27.
- Robbins-Juarez SY, Qian L, King KL, Stevens JS, Husain SA, Radhakrishnan J, et al. Outcomes for patients with COVID-19 and acute kidney injury: A systematic review and meta-analysis. Kidney Int Rep 2020;5:1149-60.

- 8. Hirsch JS, Ng JH, Ross DW, Sharma P, Shah HH, Barnett RL, et al. Acute kidney injury in patients hospitalized with COVID-19. Kidney Int 2020;98:209-18.
- Chan L, Chaudhary K, Saha A, Chauhan K, Vaid A, Zhao S, et al. AKI in hospitalized patients with COVID-19. J Am Soc Nephrol 2021;32:151-60.
- 10. Xia P, Wen Y, Duan Y, Su H, Cao W, Xiao M, et al. Clinicopathological features and outcomes of acute kidney injury in critically ill COVID-19 with prolonged disease course: A retrospective cohort. J Am Soc Nephrol 2020;31:2205-21.
- Rizo-Topete LM, Claure-Del Granado R, Ponce D, Lombardi R. Acute kidney injury requiring renal replacement therapy during the COVID-19 pandemic: what are our options for treating it in Latin America? Kidney Int 2021;99:524-7.
- Saudi MOH Protocol for Patients Suspected of/Confirmed with COVID-19. Supportive care and antiviral treatment of suspected or confirmed COVID-19 infection. V2.0. June 16, 2020.
- Kidney Disease: Improving Global Outcomes (KDIGO) [Internet]. Acute Kidney Injury (AKI). Accessed: Jan 21, 2022. Available from: https://kdigo.org/guidelines/acute-kidneyinjury/
- 14. Naar L, Langeveld K, El Moheb M, El Hechi MW, Alser O, Kapoen C, et al. Acute kidney injury in critically-ill patients with COVID-19: A single-center experience of 206 consecutive patients. Ann Surg 2020;272:e280-1.
- 15. Alser O, Mokhtari A, Naar L, Langeveld K, Breen KA, El Moheb M, et al. Multisystem outcomes and predictors of mortality in critically ill patients with COVID-19: Demographics and disease acuity matter more than comorbidities or treatment modalities. J Trauma Acute Care Surg 2021;90:880-0.
- 16. Chand S, Kapoor S, Orsi D, Fazzari MJ, Tanner TG, Umeh GC, et al. COVID-19-associated critical illness-report of the first 300 patients admitted to intensive care units at a New York City Medical Center. J Intensive Care Med 2020;35:963-70.
- 17. Mohamed MMB, Lukitsch I, Torres-Ortiz AE, Walker JB, Varghese V, Hernandez-Arroyo CF, et al. Acute kidney injury associated with coronavirus disease 2019 in urban New Orleans. Kidney360 2020;1:614-22.
- Argenziano MG, Bruce SL, Slater CL, Tiao JR, Baldwin MR, Barr RG, et al. Characterization and clinical course of 1000 patients with coronavirus disease 2019 in New York: retrospective case series. BMJ 2020;369:m1996.
- Alenezi FK, Almeshari MA, Mahida R, Bangash MN, Thickett DR, Patel JM. Incidence and risk factors of acute kidney injury in COVID-19 patients with and without acute respiratory distress syndrome (ARDS) during the first wave of COVID-19: a systematic review and meta-analysis. Ren Fail 2021;43:1621-33.
- Schaubroeck H, Vandenberghe W, Boer W, Boonen E, Dewulf B, Bourgeois C, et al. Acute kidney injury in critical COVID-19: a multicenter cohort analysis in seven large hospitals in Belgium. Crit Care 2022;26:225.
- 21. Lumlertgul N, Pirondini L, Cooney E, Kok W, Gregson J, Camporota L, et al. Acute kidney injury prevalence, progression and long-term outcomes in critically ill patients with COVID-19: a cohort study. Ann Intensive Care 2021;11:123.
- 22. de Almeida DC, Franco MDCP, Dos Santos DRP, Santos MC, Maltoni IS, Mascotte F, et al. Acute kidney injury: Incidence, risk factors, and outcomes in severe COVID-19 patients. PLoS One 2021;16:e0251048.
- 23. Farooqui MA, Almegren A, Binrushud SR, Alnuwaiser FA, Almegren NM, Alhamied NA, et al. Incidence and outcome of acute kidney injury in patients hospitalized with coronavirus



disease-19 at a tertiary care medical center in Saudi Arabia. Cureus 2021;13:e18927.

- 24. Al Sulaiman KA, Aljuhani O, Eljaaly K, Alharbi AA, Al Shabasy AM, Alsaeedi AS, et al. Clinical features and outcomes of critically ill patients with coronavirus disease 2019 (COVID-19): A multicenter cohort study. Int J Infect Dis 2021;105:180-7.
- Puelles VG, Lütgehetmann M, Lindenmeyer MT, Sperhake JP, Wong MN, Allweiss L, et al. Multiorgan and Renal Tropism of SARS-CoV-2. N Engl J Med 2020;383:590-2.
- 26. Ronco C, Navalesi P, Vincent JL. Coronavirus epidemic: preparing for extracorporeal organ support in intensive care. Lancet Respir Med 2020;8:240-1.
- 27. Cummings MJ, Baldwin MR, Abrams D, Jacobson SD, Meyer BJ, Balough EM, et al. Epidemiology, clinical course, and outcomes of critically ill adults with COVID-19 in New York City: a prospective cohort study. Lancet 2020;395:1763-70.
- 28. Gupta S, Coca SG, Chan L, Melamed ML, Brenner SK, Hayek SS, et al. AKI treated with renal replacement therapy in critically ill patients with COVID-19. J Am Soc Nephrol 2021;32:161-6.
- 29. Yang X, Tian S, Guo H. Acute kidney injury and renal replacement therapy in COVID-19 patients: a systematic review and meta-analysis. Int Immunopharmacol 2021;90:107159.
- Jewell PD, Bramham K, Galloway J, Post F, Norton S, Teo J, et al. COVID-19-related acute kidney injury; incidence, risk factors and outcomes in a large UK cohort. BMC Nephrol 2021;22:359.
- Oweis AO, Alshelleh SA, Hawasly L, Alsabbagh G, Alzoubi KH. Acute kidney injury among hospital-admitted COVID-19 patients: A study from Jordan. Int J Gen Med 2022;15:4475-82.
- 32. Bayrakci N, Özkan G, Şakaci M, Sedef S, Erdem İ, Tuna N, et al. The incidence of acute kidney injury and its association with mortality in patients diagnosed with COVID-19 followed up in intensive care unit. Ther Apher Dial 2022;26:889-96.
- 33. Yu Y, Xu D, Fu S, Zhang J, Yang X, Xu L, et al. Patients with COVID-19 in 19 ICUs in Wuhan, China: a cross-sectional study. Crit Care 2020; 24:219.
- 34. Raina R, Mahajan ZA, Vasistha P, Chakraborty R, Mukunda K, Tibrewal A, et al. Incidence and outcomes of acute kidney injury in COVID-19: A systematic review. Blood Purif 2022;51:199-212.
- 35. Anandh U, Noorin A, Kazmi SKS, Bannur S, Shah SSA, Farooq M, et al. Acute kidney injury in critically ill COVID-19 infected patients requiring dialysis: experience from India and Pakistan. BMC Nephrol 2022;23:308.
- 36. Eriksson KE, Campoccia-Jalde F, Rysz S, Rimes-Stigare C. Continuous renal replacement therapy in intensive care

patients with COVID-19; survival and renal recovery. J Crit Care 2021;64:125-30.

- 37. Richardson S, Hirsch JS, Narasimhan M, Crawford JM, McGinn T, Davidson KW, et al. Presenting characteristics, comorbidities, and outcomes among 5700 patients hospitalized with COVID-19 in the New York City area. JAMA 2020;323:2052-9.
- 38. Fominskiy EV, Scandroglio AM, Monti G, Calabrò MG, Landoni G, Dell'Acqua A, et al. Prevalence, characteristics, risk factors, and outcomes of invasively ventilated COVID-19 Patients with acute kidney injury and renal replacement therapy. Blood Purif 2021;50:102-9.
- 39. Lowe R, Ferrari M, Nasim-Mohi M, Jackson A, Beecham R, Veighey K, et al. Clinical characteristics and outcome of critically ill COVID-19 patients with acute kidney injury: a single centre cohort study. BMC Nephrol 2021;22:92.
- 40. Ng JH, Hirsch JS, Hazzan A, Wanchoo R, Shah HH, Malieckal DA, et al. Outcomes among patients hospitalized with COVID-19 and acute kidney injury. Am J Kidney Dis 2021;77:204-15.e1.
- 41. Paek JH, Kim Y, Park WY, Jin K, Hyun M, Lee JY, et al. Severe acute kidney injury in COVID-19 patients is associated with in-hospital mortality. PLoS One 2020;15:e0243528.
- 42. Tan BWL, Tan BWQ, Tan ALM, Schriver ER, Gutiérrez-Sacristán A, Das P, et al. Long-term kidney function recovery and mortality after COVID-19-associated acute kidney injury: An international multi-centre observational cohort study. eClinicalMedicine 2023;55:101724.
- 43. Benítez ID, de Batlle J, Torres G, González J, de Gonzalo-Calvo D, Targa ADS, et al. Prognostic implications of comorbidity patterns in critically ill COVID-19 patients: A multicenter, observational study. Lancet Reg Health Eur 2022;18: 100422.
- 44. Sabaghian T, Kharazmi AB, Ansari A, Omidi F, Kazemi SN, Hajikhani B, et al. COVID-19 and acute kidney injury: A systematic review. Front Med (Lausanne) 2022;9:705908.
- 45. Goh BL, Shanmuganathan M, Peariasamy K, Misnan NA, Chidambaram SK, Wong EFS, et al. COVID-19 death and kidney disease in a multiracial Asian country. Nephrology (Carlton) 2022;27:566-76.
- 46. Aggarwal S, Garcia-Telles N, Aggarwal G, Lavie C, Lippi G, Henry BM. Clinical features, laboratory characteristics, and outcomes of patients hospitalized with coronavirus disease 2019 (COVID-19): Early report from the United States. Diagnosis (Berl) 2020;7:91-6.
- 47. Bhargava A, Fukushima EA, Levine M, Zhao W, Tanveer F, Szpunar SM, et al. Predictors for severe COVID-19 infection. Clin Infect Dis 2020;71:1962-8.

Received for publication: 15 November 2022. Accepted for publication: 13 February 2023.

©Copyright: the Author(s), 2023

Licensee PAGEPress, Italy

Multidisciplinary Respiratory Medicine 2023; 18:895 doi:10.4081/mrm.2023.895

This work is licensed under a Creative Commons Attribution-NonCommercial 4.0 International License (CC BY-NC 4.0).

Publisher's note: All claims expressed in this article are solely those of the authors and do not necessarily represent those of their affiliated organizations, or those of the publisher, the editors and the reviewers. Any product that may be evaluated in this article or claim that may be made by its manufacturer is not guaranteed or endorsed by the publisher.