

| pISSN 2586-6052 | eISSN 2586-6060

Impact of intradialytic hypotension on mortality following the transition from continuous renal replacement therapy to intermittent hemodialysis

Seong Geun Kim, Donghwan Yun, Jinwoo Lee, Yong Chul Kim, Dong Ki Kim, Kook-Hwan Oh, Kwon Wook Joo, Yon Su Kim, Seung Seok Han

Department of Internal Medicine, Seoul National University College of Medicine, Seoul, Korea

Background: The transition of dialysis modalities from continuous renal replacement therapy (CRRT) to intermittent hemodialysis (iHD) is frequently conducted during the recovery phase of critically ill patients with acute kidney injury. Herein, we addressed the occurrence of intradialytic hypotension (IDH) after this transition, and its association with the mortality risk.

Methods: A total of 541 patients with acute kidney injury who attempted to transition from CRRT to iHD at Seoul National University Hospital, Korea from 2010 to 2020 were retrospectively collected. IDH was defined as a discontinuation of dialysis because of hemodynamic instability plus a nadir systolic blood pressure <90 mm Hg or a decrease in systolic blood pressure \geq 30 mm Hg during the first session of iHD. Odds ratios (ORs) of outcomes, such as in-hospital mortality and weaning from RRT, were measured using a logistic regression model after adjusting for multiple variables.

Results: IDH occurred in 197 patients (36%), and their mortality rate (44%) was higher than that of those without IDH (19%; OR, 2.64; 95% confidence interval [CI], 1.70–4.08). For patients exhibiting IDH, the iHD sessions delayed successful weaning from RRT (OR, 0.62; 95% CI, 0.43–0.90) compared with sessions on those without IDH. Factors such as low blood pressure, high pulse rate, low urine output, use of mechanical ventilations and vasopressors, and hypoalbuminemia were associated with IDH risk.

Conclusions: IDH occurrence following the transition from CRRT to iHD is associated with high mortality and delayed weaning from RRT.

Key Words: acute kidney injury; continuous renal replacement therapy; hypotension; mortality; renal dialysis

INTRODUCTION

Acute kidney injury (AKI) requiring renal replacement therapy (RRT) is a common and serious condition that is associated with poor outcomes [1]. The incidence of RRT-requiring AKI has increased by 10% per year from 2000 to 2009 [2], and its death rate is over 50% [3,4]. Even for survivors, RRT-requiring AKI carries the risk of several complications, such as progression to chronic kidney disease [5,6] and cardiovascular events, including myocardial infarction

Original Article

Received: August 1, 2022 Revised: September 27, 2022 Accepted: October 14, 2022

Corresponding author

Seung Seok Han Department of Internal Medicine, Seoul National University College of Medicine, 103 Daehak-ro, Jongno-gu, Seoul 03080, Korea Tel: +82-2-2072-4785 Fax: +82-2-745-2264 Email: hansway7@snu.ac.kr

Copyright © 2023 The Korean Society of Critical Care Medicine

This is an Open Access article distributed under the terms of Creative Attributions Non-Commercial License (https://creativecommons.org/ li-censes/by-nc/4.0/) which permits unrestricted noncommercial use, distribution, and reproduction in any medium, provided the original work is properly cited.



and congestive heart failure [7,8].

Modalities of RRT in the setting of AKI include continuous renal replacement therapy (CRRT) and intermittent hemodialysis (iHD). Whereas iHD achieves a rapid clearance of small uremic toxins within a short time, CRRT provides osmotic stability by the gradual elimination of fluid and toxins over a long period [9]. Although previous studies have not demonstrated the difference in survival or renal recovery outcome between the two modalities, CRRT is recommended as an initial modality for patients with hemodynamic instability [10-13].

Once patients with AKI achieve hemodynamic stability and fluid balance with CRRT, clinicians may consider a transition to iHD [10,14]. Sometimes, the unaffordable cost of CRRT or the need for patient mobilization and rehabilitation demands the transition to iHD [15,16]. Although RRT modality transitions are a common occurrence in the intensive care unit (ICU), data regarding adverse events, such as intradialytic hypotension (IDH) or arrhythmia after transition, are still limited.

IDH is attributable to overzealous ultrafiltration and impaired vascular resistance in decreased cardiovascular reserves [17]. IDH can occur in 10%–80% of patients with AKI, which may be dependent on the definition [18-20]. IDH is associated with high mortality in patients with AKI regardless of its modalities [19,21]. However, it remains undetermined whether IDH after a transition from CRRT to iHD is associated with adverse outcomes in patients with AKI. The present study addressed this issue using a cohort undergoing a transition and identified risk factors related to IDH occurrence.

MATERIALS AND METHODS

The study was approved by the Institutional Review Board of the Seoul National University Hospital (No. H-2110-085-1262) and complied with the Declaration of Helsinki. The requirement for informed consent was waived by the Review Board.

Patient and Data Collection

The study was retrospective in nature using a cohort of 2,397 patients who underwent CRRT due to AKI at Seoul National University Hospital from June 2010 to December 2020. A total of 1,108 patients underwent a transition from CRRT to iHD because of the recovery phase in patient status. When the volume control became easy due to resolved oliguria and need of vasoactive agents was reduced, the transition between modalities was recommended according to the Kidney Disease Improving Global guideline [10]. Patients who were under 18 years

old (n=11) and patients without information on iHD (n=556) were excluded. Accordingly, 541 patients were analyzed in the present study (Figure 1).

Baseline information at the time of transition was collected, such as age, sex, body mass index, cause of AKI (e.g., septic and nonseptic), the time interval between CRRT and iHD, systolic blood pressure, diastolic blood pressure, mean arterial pressure, pulse rate, duration of iHD, ultrafiltration volume, urine output, mechanical ventilation, and use of vasopressors. Blood pressures were measured using noninvasive blood pressure monitoring system. Sequential Organ Failure Assessment (SOFA) was calculated according to the original formula [22]. Dialysis settings, such as ultrafiltration target, blood flow rate, and session time, could be different between patient according to their status. All patients performed high-flux hemodialysis. Either unfractionated heparin or nafamostat was used as an anticoagulant during CRRT. The Charlson comorbidity index was calculated to consider the effect of various comorbidities, such as acute myocardial infarction, congestive heart failure, peripheral vascular disease, cerebral vascular accident, dementia, pulmonary disease, connective tissue disorder, peptic ulcer, liver disease, diabetes, paraplegia, chronic kidney disease, malignancy, and human immunodeficiency virus infection [23].

The patients were categorized into two groups according to the presence of IDH during the 1st session of iHD. Because there is no consensus on defining IDH when patients have AKI rather than end-stage kidney disease, we referred to methods used in previous studies as follows [24]: discontinuation of dialysis as a result of hemodynamic instability plus a nadir systolic blood pressure <90 mm Hg (termed IDH-A) or a decrease in systolic blood pressure \geq 30 mm Hg (termed IDH-B).



Figure 1. Flowchart of selection criteria. CRRT: continuous renal replacement therapy; AKI: acute kidney injury; iHD: intermittent hemodialysis.

Outcomes

The primary outcome was all-cause mortality after transition until hospital discharge or death. Additionally, weaning from RRT as a cessation of RRT for >7 days was evaluated during the above timeframe.

Statistical Analysis

Categorical and continuous variables are expressed as proportions and means±standard deviations when they were normally distributed and as medians with interquartile ranges when they were not normally distributed, respectively. The normality of the distribution was analyzed using the Kolmogorov-Smirnov test. The chi-square test or Fisher's exact test was used to compare categorical variables. The Student t test or the Mann-Whitney U-test was used for continuous variables with or without a normal distribution. Kaplan-Meier survival curves were drawn and compared between groups using a logrank test. The odds ratios (ORs) and confidence intervals for mortality and kidney outcomes were calculated using logistic regression model. The adjusted variables were selected based on their significant association with mortality. To identify risk factors for IDH, backward stepwise selection was further applied. All statistical analyses were performed using IBM SPSS ver. 27 (IBM (IBM Corp.) and R software (ver. 3.5.1; R core team). A P-value of less than 0.05 was considered statistically significant.

RESULTS

Patient Characteristics

The mean patient age was 64 ± 15 years, and 39.5% of the patients were female. The proportion of patients with septic AKI was 50.6\%. IDH occurred in 197 patients (36.4%). Among them, 123 patients (22.7%) had IDH-A alone, 138 patients (25.5%) had IDH-B alone, and 64 patients (11.8%) had both. The IDH prevalence was different according to the ICU subtypes: 39.0% in medical ICU, 28.5% in surgical ICU, 35.6% in cardiopulmonary ICU, and 38.2% in emergency ICU.

The IDH group was more likely to have low systolic blood pressure, high pulse rate, low urine output, high SOFA score and the need for mechanical ventilation and vasopressors than the non-IDH group. Before hemodialysis, 22.0% of patients and 9.2% after hemodialysis used vasopressors, respectively. Other baseline characteristics are shown in Table 1.

Relationship between IDH and Mortality

During a median follow-up period of 25 days (interquartile range, 13–54 days), 160 patients (28.3%) died. The mortality incidence was 5.7 deaths per 1,000 person-days. When a univariate Cox regression model was applied, several variables, such as cause of AKI, pulse rate, SOFA score, platelet count, and total bilirubin, were associated with in-hospital mortality in the present cohort (Table 2), and these were included in subsequent multivariate regression models as adjusting factors. Figure 2 shows Kaplan-Meier survival curves according to IDH occurrence; the groups with IDH-A, IDH-B, or both had lower survival rates than the non-IDH group. The low survival rates in patients with IDH compared to those without IDH remained consistent after adjusting for multiple variables (Table 3).

Relationship between IDH and Weaning from RRT

Figure 3 depicts curves that show weaning from RRT between the groups with and without IDH, and the weaning rate was higher in the non-IDH group than in the IDH group (P<0.001). In a subgroup analysis, the association between IDH and low weaning from RRT was prominent when patients were male and had septic AKI or high SOFA scores (Figure 4). When logistic regression was applied, the iHD sessions delayed successful weaning from RRT (OR, 0.62; 95% confidence interval, 0.43–0.90) compared with sessions on those without IDH.

Factors Related to IDH

When a multivariate logistic regression model was applied, several factors were associated with IDH occurrence, such as low systolic blood pressure, high pulse rate, low urine output, and the use of mechanical ventilation and vasopressors. Regarding IDH-A, hypoalbuminemia was further noted to correlate with IDH occurrence (Table 4).

DISCUSSION

It is well known that IDH occurrence confers a high risk of mortality in patients with end-stage kidney disease; however, this relationship has not been well established for AKI patients requiring iHD. The present study used a cohort of patients with AKI who needed a transition from CRRT to iHD. IDH occurrence after transition was associated with subsequent high risks of mortality and nonweaning from RRT. Several factors were identified to be associated with IDH occurrence, and the findings will help clinicians identify patients at risk of hemody-



Table 1. Baseline patient characteristics

Variable	Overall (n=541)	Non-IDH (n=344)	IDH (n=197)	P-value
Age (yr)	64.2±15.3	63.8±15.9	64.9±14.0	0.416
Female (%)	39.5	38.4	42.6	0.330
Body mass index (kg/m ²)	23.2±4.5	23.3±4.5	23.5±4.7	0.573
Septic acute kidney injury (%)	49.7	47.4	53.8	0.150
Transition after 24 hr (%)	25.5	27.8	22.3	0.219
Systolic blood pressure (mm Hg)	123.8±21.3	125.4±18.6	120.6±25.0	0.020
Diastolic blood pressure (mm Hg)	68.3±13.5	68.0±13.1	68.8±14.3	0.524
Mean arterial pressure (mm Hg)	86.8±13.7	87.1±12.8	86.1±15.2	0.401
Pulse rate (beats/min)	91.1±18.7	87.9±16.9	96.4±20.1	<0.001
Hemodialysis duration (hr)	3.0±0.3	3.0±0.3	3.0±0.2	0.213
Ultrafiltration volume (L)	1.0 (0.5–1.8)	1.0 (0.5–1.8) 1.1 (0.5–1.8)		0.467
Urine output (ml/day)	159.0 (35.0–546.0)	195.0 (45.0–684.0)	105.0 (30.0–312.0)	0.001
Mechanical ventilation (%)	29.9	24.1	39.1	<0.001
Use of vasopressors (%)	22.0	16.2	32.0	<0.001
SOFA score	8.9±2.8	8.6±2.7	9.4±2.7	<0.001
Charlson comorbidity index	3 (2–5)	3 (2–5)	3 (2–4)	0.641
Previous history (%)				
Myocardial infarction	10.9	11.9	9.1	0.318
Congestive heart failure	22.2	24.7	19.8	0.161
Diabetes	34.0	34.6	33.0	0.706
Stroke	13.7	15.4	10.7	0.102
Liver cirrhosis	4.4	4.7	4.1	0.748
Active malignancy	32.7	32.0	34.0	0.628
Blood finding				
Hemoglobin (g/dl)	9.3±1.4	9.4±1.4	9.3±1.3	0.599
Platelet (×10 ³ /ml)	81.0 (49.0–120.0)	82.0 (51.8–123.3)	78.0 (43.0–118.0)	0.162
Blood urea nitrogen (mmol/l)	29.0 (20.0–40.0)	29 (19.5–41.0)	28 (21.0–39.0)	0.463
Creatinine (mg/dl)	1.7 (1.3–2.4)	1.7 (1.3–2.5)	1.8 (1.3–2.4)	0.938
Sodium (mmol/l)	135.8±3.4	135.9±3.3	135.8±3.5	0.337
Potassium (mmol/l)	4.0±0.5	4.0±0.5	3.9±0.5	0.302
Chloride (mmol/l)	100.8±3.5	100.9±3.5	100.8±3.6	0.808
Calcium (mg/dl)	8.6±0.7	8.6±0.7	8.5±0.6	0.117
Phosphate (mg/dl)	3.1 (2.5–3.8)	3.1 (2.5–3.8)	3.1 (2.5–3.9)	0.234
Total bilirubin (mg/dl)	1.8 (0.9–5.0)	1.7 (0.9–4.5)	2.0 (0.9–5.8)	0.210
Albumin (g/dl)	2.8 (2.6-3.2)	2.9 (2.6-3.2)	2.8 (2.5-3.1)	0.060

Values are presented as mean±standard deviation or median (interguartile range).

IDH: intradialytic hypotension; SOFA: Sequential Organ Failure Assessment.

namic instability, such as IDH after the transition to iHD.

The prevalence of IDH on severe AKI varied from 30% to 90% depending on the definition of AKI and its timing [19,21,25]. Several factors may further increase the risk of IDH in patients with AKI receiving iHD compared to those with end-stage kid-ney disease receiving maintenance iHD. These include fluid overload due to hemodynamic instability, nutritional support, and intravenous antibiotics. Furthermore, IDH can occur because of an impaired response to the physiological stress of

RRT, such as increased vascular resistance or cardiac reserve due to critical illness [26]. The occurrence of IDH might be related to the delivered dose of RRT and fluid removal being insufficient, all of which influence patient outcomes.

A previous prospective cohort study of patients with AKI requiring RRT, wherein IDH was defined for each session of RRT as either the occurrence of mean arterial pressure <65 mm Hg or a reduction of \geq 20% from the starting mean arterial pressure, demonstrated that more frequent exposure to IDH led to



Variable	Odds ratio (95% Cl)	P-value
Age (per 10 yr)	1.03 (0.906–1.159)	0.693
Female (vs. male)	0.70 (0.476–1.039)	0.077
Body mass index (per 1 kg/m ²)	1.01 (0.967–1.053)	0.675
Septic acute kidney injury (vs. nonseptic)	2.18 (1.480–3.197)	<0.001
Transition after 24 hr (vs. within 24 hr)	1.25 (0.973–1.604)	0.081
Systolic blood pressure (per 10 mm Hg)	0.88 (0.804–0.963)	0.005
Diastolic blood pressure (per 10 mm Hg)	1.01 (0.878–1.160)	0.900
Mean arterial pressure (per 10 mm Hg)	0.91 (0.791–1.041)	0.166
Pulse rate (per 10 beat/min)	1.29 (1.159–1.432)	0.001
Hemodialysis duration (per 1 hr)	0.96 (0.521–1.748)	0.880
Ultrafiltration volume (per 1 L)	1.03 (0.842–1.248)	0.806
Urine output (per 100 ml/day)	0.96 (0.923-1.000)	0.049
Mechanical ventilation (%)	2.28 (1.534–3.383)	<0.001
Use of vasopressors (vs. none)	1.61 (1.041–2.473)	0.032
SOFA score (per 1 score)	1.23 (1.146–1.327)	<0.001
Charlson comorbidity index (per 1 score)	1.07 (0.988–1.154)	0.097
Blood finding		
Hemoglobin (per 1 g/dl)	0.88 (0.765–1.020)	0.090
Platelet (per ×10³/ml)	0.99 (0.988–0.996)	0.001
Blood urea nitrogen (per 1 mmol/l)	1.00 (0.987–1.010)	0.754
Creatinine (per 1 mg/dl)	0.89 (0.740-1.075)	0.230
Sodium (per 1 mmol/l)	0.99 (0.938–1.049)	0.776
Potassium (per 1 mmol/l)	1.02 (0.707–1.458)	0.933
Chloride (per 1 mmol/l)	0.99 (0.937–1.042)	0.662
Calcium (per 1 mg/dl)	1.00 (0.771–1.303)	0.985
Phosphate (per 1 mg/dl)	1.07 (0.893–1.290)	0.450
Total bilirubin (per 1 mg/dl)	1.08 (1.049–1.104)	<0.001
Albumin (per 1 g/dl)	0.62 (0.415-0.934)	0.022

Cl: confidence interval; SOFA: Sequential Organ Failure Assessment.





Figure 2. Kaplan-Meier survival curves between patients with and without intradialytic hypotension (IDH). IDH was defined as a discontinuation of dialysis because of hemodynamic instability plus a nadir systolic blood pressure <90 mm Hg (IDH-A) or a decrease in systolic blood pressure \geq 30 mm Hg (IDH-B).

Figure 3. Kaplan-Meier curves of the weaning rate from renal replacement therapy (RRT). IDH: intradialytic hypotension.





Figure 4. Forest plot of subgroup analyses for the odds ratio (OR) of the renal replacement therapy weaning rate in the group with intradialytic hypotension compared with that of the group without intradialytic hypotension. CI: confidence interval; AKI: acute kidney injury; SOFA: Sequential Organ Failure Assessment.

TILO	<u> </u>	C 1 1 1/1			1 141	11 14 A ID11
Table 3.	Odds ratios	of in-hospital	mortality in	patients with IDH	compared with	those without IDH
	00001000	01 111 1100001001		porcienco mici ibri	companed mich	

Outcome	Unadjusted OR (95% CI)	P-value	Adjusted OR (95% Cl) ^{a)}	P-value
IDH	3.33 (2.258–4.914)	<0.001	2.57 (1.632–4.040)	<0.001
IDH-A ^{b)}	3.33 (2.182–5.076)	<0.001	2.71 (1.600-4.643)	<0.001
IDH-B ^{b)}	2.46 (1.631–3.694)	<0.001	2.35 (1.443–3.834)	0.001

IDH: intradialytic hypotension; OR: odds ratio; CI: confidence interval; SOFA: Sequential Organ Failure Assessment.

a) Adjusted for age, sex, septic acute kidney injury, systolic blood pressure, pulse rate, urine output, mechanical ventilation, use of vasopressors, SOFA score, Charlson comorbidity index, hemoglobin, platelet, total bilirubin, and albumin; b) IDH was defined as a discontinuation of dialysis because of hemodynamic instability plus a nadir systolic blood pressure <90 mm Hg (IDH-A) or a decrease in systolic blood pressure \geq 30 mm Hg (IDH-B).

Table 4. Factors associated with IDH in logistic regression with backward stepwise selection

Variable	IDH		IDH-A ^{a)}		IDH-B ^{a)}	
variable	OR (95% Cl)	P-value	OR (95% CI)	P-value	OR (95% CI)	P-value
Age (per 10 yr)	1.10 (0.970–1.257)	0.135	1.15 (0.973–1.350)	0.103	1.12 (0.972–1.282)	0.118
Female (vs. male)	1.23 (0.836–1.802)	0.295	1.25 (0.787–1.999)	0.341	1.47 (0.972–2.213)	0.068
Systolic blood pressure (per 10 mm Hg)	0.92 (0.834–1.003)	0.058	0.68 (0.594–0.768)	< 0.001	1.15 (1.033–1.274)	0.010
Pulse rate (per 10 beats/min)	1.33 (1.191–1.485)	<0.001	1.35 (1.187–1.534)	< 0.001	1.31 (1.170–1.475)	0.001
Transition after 24 hr (vs. within 24 hr)	1.13 (0.873–1.471)	0.348	1.15 (0.826–1.556)	0.397	0.99 (0.746–1.311)	0.937
Hemodialysis duration (per 1 hr)	0.61 (0.314–1.191)	0.148	0.67 (0.298–1.526)	0.355	0.63 (0.304–1.303)	0.213
Ultrafiltration volume (per 1 L)	1.10 (0.898–1.354)	0.348	1.05 (0.831–1.336)	0.665	0.99 (0.794–1.232)	0.989
Urine output (per 100 ml/day)	0.96 (0.921–0.998)	0.038	0.97 (0.925–1.023)	0.259	0.96 (0.964–1.007)	0.096
Mechanical ventilation (%)	1.71 (1.121–2.553)	0.012	1.39 (0.848–2.284)	0.191	1.40 (0.900–2.178)	0.136
Use of vasopressor (vs. none)	1.91 (1.212–3.015)	0.005	2.84 (1.719–4.691)	< 0.001	1.72 (1.053–2.794)	0.030
Albumin (per 1 g/dl)	0.71 (0.470–1.055)	0.089	0.51 (0.311–0.845)	0.009	1.00 (0.656–1.524)	0.998

IDH: intradialytic hypotension; OR: odds ratio; CI: confidence interval.

a) IDH was defined as a discontinuation of dialysis because of hemodynamic instability plus a nadir systolic blood pressure <90 mm Hg (IDH-A) or a decrease in systolic blood pressure \ge 30 mm Hg (IDH-B).

<mark>∧CC</mark>√

a higher risk of mortality outcome regardless of RRT modality [19]. Another observational study with CRRT alone showed that IDH occurring within the first hour of CRRT initiation, defined as a decline in blood pressure from baseline, was associated with an increased risk of mortality [21]. The present study reconfirmed the association of patient outcome with IDH, although the definition and timing of IDH differ between studies.

One previous study suggested a high risk of mortality when IDH occurs during the transition period (n=213) [25]. The present study expanded the sample size and added sufficient laboratory data at the time of iHD initiation to the analyses. Furthermore, the present study considered an RRT weaning outcome. Baseline hypoalbuminemia indicated the risk of IDH, which might reflect pathological conditions, such as organ failure, nutritional deficiency, and inflammation, in critically ill patients [27,28]. Because serum albumin primarily determines osmotic pressure and has a decisive effect on intravascular volume, hypoalbuminemia would result in hemodynamic instability while on iHD. In a prospective randomized controlled trial, the administration of albumin during iHD resulted in few episodes of IDH and improved the fluid removal rate compared to the nonadministration of albumin [29].

The occurrence of IDH delayed RRT weaning and thus prolonged the dependence of patients with ischemic AKI on RRT [30]. Hot lesions in ischemic tubules were further observed in kidney biopsies of patients dependent on RRT, which suggests that IDH would cause an additional ischemic injury [31]. In a randomized controlled trial comparing CRRT and iHD after AKI, IDH was associated with mortality and a lower probability of renal recovery for both modalities [32].

Although the study is informative, there are certain limitations to be addressed. Because of the retrospective design, unmeasured bias and confounders could have interfered with the present analyses. Continuous fluctuations in biochemical parameters and alterations during practice could be related to mortality but were not considered in the study. The present study did not categorize nonseptic patients into specific causes, which could further segment risk factors of IDH. The cause of death was not taken into consideration, which could affect the results.

The occurrence of IDH after transitioning from CRRT is associated with an increased risk of hospital mortality and dependence on RRT. Certain factors, such as low blood pressure, high pulse rate, low urine output, use of mechanical ventilation and vasopressors, and hypoalbuminemia, are associated with IDH occurrence. The findings provide a basis for future studies to clarify the clinical implications of IDH and strategies to prevent it in the context of transitions between RRT modalities in patients with AKI.

CONFLICT OF INTEREST

No potential conflict of interest relevant to this article was reported.

FUNDING

None.

ACKNOWLEDGMENTS

The results presented in this paper have not been published previously in whole or part, except in abstract format at The Korean Society of Nephrology (KSN) 2022.

ORCID

Seong Geun Kim Donghwan Yun Jinwoo Lee Yong Chul Kim Dong Ki Kim Kook-Hwan Oh Kwon Wook Joo Yon Su Kim Seung Seok Han https://orcid.org/0000-0002-8831-9838 https://orcid.org/0000-0001-6566-5183 https://orcid.org/0000-0003-0656-9364 https://orcid.org/0000-0003-3215-8681 https://orcid.org/0000-0002-5195-7852 https://orcid.org/0000-0001-9525-2179 https://orcid.org/0000-0001-9941-7858 https://orcid.org/0000-0003-3091-2388 https://orcid.org/0000-0003-0137-5261

AUTHOR CONTRIBUTIONS

Conceptualization: SGK, KWJ, SSH. Data curation: JL, YCK, YSK. Formal analysis: SGK, DY, DKK, KHO. Methodology: DY, DKK, KHO, KWJ. Project administration: YSK. Writing–original draft: SGK. Writing–review & editing: SSH.

REFERENCES

- 1. Cerdá J, Lameire N, Eggers P, Pannu N, Uchino S, Wang H, et al. Epidemiology of acute kidney injury. Clin J Am Soc Nephrol 2008;3:881-6.
- 2. Hsu RK, McCulloch CE, Dudley RA, Lo LJ, Hsu CY. Tempo-

ral changes in incidence of dialysis-requiring AKI. J Am Soc Nephrol 2013;24:37-42.

- **3.** Uchino S, Kellum JA, Bellomo R, Doig GS, Morimatsu H, Morgera S, et al. Acute renal failure in critically ill patients: a multinational, multicenter study. JAMA 2005;294:813-8.
- 4. Tolwani A. Continuous renal-replacement therapy for acute kidney injury. N Engl J Med 2012;367:2505-14.
- Lo LJ, Go AS, Chertow GM, McCulloch CE, Fan D, Ordoñez JD, et al. Dialysis-requiring acute renal failure increases the risk of progressive chronic kidney disease. Kidney Int 2009;76:893-9.
- 6. Duran PA, Concepcion LA. Survival after acute kidney injury requiring dialysis: long-term follow up. Hemodial Int 2014;18 Suppl 1:S1-6.
- Chawla LS, Amdur RL, Shaw AD, Faselis C, Palant CE, Kimmel PL. Association between AKI and long-term renal and cardiovascular outcomes in United States veterans. Clin J Am Soc Nephrol 2014;9:448-56.
- 8. Wu VC, Wu CH, Huang TM, Wang CY, Lai CF, Shiao CC, et al. Long-term risk of coronary events after AKI. J Am Soc Nephrol 2014;25:595-605.
- 9. Bellomo R, Kellum JA, La Manna G, Ronco C. 40 Years of continuous renal replacement therapy. Karger; 2018.
- **10.** Khwaja A. KDIGO clinical practice guidelines for acute kidney injury. Nephron Clin Pract 2012;120:c179-84.
- 11. Schefold JC, von Haehling S, Pschowski R, Bender T, Berkmann C, Briegel S, et al. The effect of continuous versus intermittent renal replacement therapy on the outcome of critically ill patients with acute renal failure (CONVINT): a prospective randomized controlled trial. Crit Care 2014;18:R11.
- 12. Chawla LS, Bellomo R, Bihorac A, Goldstein SL, Siew ED, Bagshaw SM, et al. Acute kidney disease and renal recovery: consensus report of the Acute Disease Quality Initiative (ADQI) 16 Workgroup. Nat Rev Nephrol 2017;13:241-57.
- Nash DM, Przech S, Wald R, O'Reilly D. Systematic review and meta-analysis of renal replacement therapy modalities for acute kidney injury in the intensive care unit. J Crit Care 2017;41:138-44.
- 14. Karkar A, Ronco C. Prescription of CRRT: a pathway to optimize therapy. Ann Intensive Care 2020;10:32.
- Brownback CA, Fletcher P, Pierce LN, Klaus S. Early mobility activities during continuous renal replacement therapy. Am J Crit Care 2014;23:348-51.
- 16. Srisawat N, Lawsin L, Uchino S, Bellomo R, Kellum JA, BEST Kidney Investigators. Cost of acute renal replacement therapy in the intensive care unit: results from The Beginning and Ending Supportive Therapy for the Kidney (BEST Kidney) study. Crit

Care 2010;14:R46.

17. Sars B, van der Sande FM, Kooman JP. Intradialytic hypotension: mechanisms and outcome. Blood Purif 2020;49:158-67.

- 18. VA/NIH Acute Renal Failure Trial Network, Palevsky PM, Zhang JH, O'Connor TZ, Chertow GM, Crowley ST, et al. Intensity of renal support in critically ill patients with acute kidney injury. N Engl J Med 2008;359:7-20.
- **19.** Silversides JA, Pinto R, Kuint R, Wald R, Hladunewich MA, Lapinsky SE, et al. Fluid balance, intradialytic hypotension, and outcomes in critically ill patients undergoing renal replacement therapy: a cohort study. Crit Care 2014;18:624.
- **20.** Tonelli M, Astephen P, Andreou P, Beed S, Lundrigan P, Jindal K. Blood volume monitoring in intermittent hemodialysis for acute renal failure. Kidney Int 2002;62:1075-80.
- 21. Shawwa K, Kompotiatis P, Jentzer JC, Wiley BM, Williams AW, Dillon JJ, et al. Hypotension within one-hour from starting CRRT is associated with in-hospital mortality. J Crit Care 2019;54:7-13.
- 22. Jones AE, Trzeciak S, Kline JA. The Sequential Organ Failure Assessment score for predicting outcome in patients with severe sepsis and evidence of hypoperfusion at the time of emergency department presentation. Crit Care Med 2009;37:1649-54.
- 23. Sundararajan V, Henderson T, Perry C, Muggivan A, Quan H, Ghali WA. New ICD-10 version of the Charlson comorbidity index predicted in-hospital mortality. J Clin Epidemiol 2004;57:1288-94.
- 24. Flythe JE, Xue H, Lynch KE, Curhan GC, Brunelli SM. Association of mortality risk with various definitions of intradialytic hypotension. J Am Soc Nephrol 2015;26:724-34.
- **25.** Beaubien-Souligny W, Yang Y, Burns KE, Friedrich JO, Meraz-Muñoz A, Clark EG, et al. Intra-dialytic hypotension following the transition from continuous to intermittent renal replacement therapy. Ann Intensive Care 2021;11:96.
- **26.** Douvris A, Zeid K, Hiremath S, Bagshaw SM, Wald R, Beaubien-Souligny W, et al. Mechanisms for hemodynamic instability related to renal replacement therapy: a narrative review. Intensive Care Med 2019;45:1333-46.
- 27. Kaysen GA, Dubin JA, Müller HG, Rosales L, Levin NW, Mitch WE, et al. Inflammation and reduced albumin synthesis associated with stable decline in serum albumin in hemodialysis patients. Kidney Int 2004;65:1408-15.
- 28. Gabay C, Kushner I. Acute-phase proteins and other systemic responses to inflammation. N Engl J Med 1999;340:448-54.
- **29.** Macedo E, Karl B, Lee E, Mehta RL. A randomized trial of albumin infusion to prevent intradialytic hypotension in hospitalized hypoalbuminemic patients. Crit Care 2021;25:18.
- 30. Manns M, Sigler MH, Teehan BP. Intradialytic renal haemo-



dynamics: potential consequences for the management of the patient with acute renal failure. Nephrol Dial Transplant 1997;12:870-2.

31. Kelleher SP, Robinette JB, Miller F, Conger JD. Effect of hemorrhagic reduction in blood pressure on recovery from acute renal failure. Kidney Int 1987;31:725-30.

32. Augustine JJ, Sandy D, Seifert TH, Paganini EP. A randomized controlled trial comparing intermittent with continuous dialysis in patients with ARF. Am J Kidney Dis 2004;44:1000-7.