Incidence and Outcomes of Acute Kidney Injury Requiring Renal Replacement Therapy in Patients on Percutaneous Mechanical Circulatory Support with Impella-CP for Cardiogenic Shock

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Abstract
Background: Acute kidney injury (AKI) complicating cardiogenic shock is associated with increased mortality. We hypothesize that renal replacement therapy (RRT) improves survival in cardiogenic shock supported by Impella-CP (Abiomed, Danvers, MA) complicated by AKI.

Methods: A retrospective chart review identified 34 patients on Impella-CP for cardiogenic shock between January 2015 and December 2017. AKI was defined as an increase in serum creatinine≥0.3 mg/dL from baseline. Three groups were analyzed: AKI plus RRT, AKI minus RRT, and no AKI. Pre-existing dialysis patients were excluded. The only indication for RRT was AKI not responding to diuretics. Thirty-day mortality was analyzed.

Results: There were 13 patients with no AKI, 9 with AKI plus RRT groups, and 12 with AKI minus RRT. Thirty-day mortality was similar between no AKI and AKI plus RRT groups [30.8% (4/13) vs. 22.2% (2/9), p=0.48; relative risk [RR] 2.25 (95% confidence interval [CI] 0.22-22.1)]. Thirty-day mortality was higher in AKI minus RRT group compared to the no AKI group [75.0% (9/12) vs. 30.8% (4/13); p=0.03; RR 6.75 (95% CI 1.16-39.2)].

Conclusion: In cardiogenic shock patients on Impella-CP, AKI minus RRT is associated with a higher 30-day mortality compared to patients without AKI and/or patients with AKI plus RRT. Short-term mortality may improve in cardiogenic shock patients with AKI who are treated with RRT.

Introduction
Despite advances in technology for the treatment of cardiogenic shock, mortality has not dramatically improved [1]. Cardiogenic shock primarily occurs in the setting of acute myocardial infarction. Standard of care in acute myocardial infarction complicated by cardiogenic shock is revascularization [2]. Percutaneous mechanical circulatory support devices, such as the Impella-CP (Abiomed, Danvers, MA), are used as supportive therapy in cardiogenic shock, despite limited randomized clinical data [3]. Registry data do suggest

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improved outcomes in cardiogenic shock supported by the Impella-CP [4]. However, there is a paucity of data surrounding outcomes in cardiogenic shock patients with concurrent acute kidney injury (AKI), managed with or without renal replacement therapy (RRT). Cardiogenic shock complicated by AKI requiring RRT is associated with increased mortality [5,6]. We hypothesize that early RRT for AKI in cardiogenic shock patients on Impella-CP improves survival.

**Materials And Methods**

Our cohort was a single-center retrospective study including all patients on Impella-CP for cardiogenic shock admitted to Albany Medical Center between January 2015 and December 2017. Data were obtained by a retrospective review of the electronic medical record. Cardiogenic shock was defined as elevated serum lactate (>2.0 mmol/L) and hypotension requiring inotrope/vasopressors to maintain a mean arterial blood pressure above 65 mmHg. Eligible patients were classified based on AKI at presentation (increase in serum creatinine>0.3 mg/dL from baseline); those with AKI were further categorized by RRT. This led to three groups: no AKI, AKI minus RRT, and AKI plus RRT. RRT included hemodialysis or continuous RRT.

The exclusion criteria included pre-existing hemodialysis-dependent patients, unknown baseline renal function, or patients lost to follow-up within 30 days. The chi-square test was utilized to compare 30-day mortality of AKI plus RRT and no AKI groups as well as AKI minus RRT and no AKI groups. Continuous variables (lab parameters at presentation) were compared between groups using ANOVA. Type 1 error was prespecified at less than or equal to 5%. The protocol was approved by the Institutional Review Board at Albany Medical Center.

**Results**

Between January 2015 and December 2017, 34 patients with cardiogenic shock on Impella-CP met the inclusion criteria for our study. There were 15 patients with no AKI, 9 had AKI plus RRT, and 12 had AKI minus RRT. The only indication for RRT was AKI not responding to diuretics (oliguria or anuria). Baseline characteristics and lab parameters at presentation are included in Tables 1, 2.
<table>
<thead>
<tr>
<th></th>
<th>no-AKI</th>
<th>AKI without RRT*</th>
<th>AKI+RRT*</th>
<th>p-value</th>
</tr>
</thead>
<tbody>
<tr>
<td>Hemoglobin (g/dL)</td>
<td>13.2±0.529</td>
<td>12.1±0.583</td>
<td>12.9±1.46</td>
<td>0.448</td>
</tr>
<tr>
<td>Serum creatinine (mg/dL)</td>
<td>0.97±0.071</td>
<td>2.10±0.320</td>
<td>2.55±0.918</td>
<td>0.010</td>
</tr>
<tr>
<td>Serum lactate (mmol/L)</td>
<td>5.82±4.01</td>
<td>4.73±0.628</td>
<td>4.75±0.329</td>
<td>0.904</td>
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<tr>
<td>Serum HCO₃ (mEq/L)</td>
<td>20.9±1.42</td>
<td>19.7±1.40</td>
<td>19.7±2.78</td>
<td>0.773</td>
</tr>
<tr>
<td>INR</td>
<td>1.96±0.529</td>
<td>1.70±0.416</td>
<td>1.67±0.509</td>
<td>0.929</td>
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<tr>
<td>Arterial blood gas</td>
<td></td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>pH</td>
<td>7.20±0.053</td>
<td>7.28±0.024</td>
<td>7.18±0.055</td>
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<tr>
<td>pCO₂</td>
<td>47.0±5.24</td>
<td>41.8±2.63</td>
<td>51.0±9.43</td>
<td>0.687</td>
</tr>
<tr>
<td>pO₂</td>
<td>146±46.9</td>
<td>163±41.2</td>
<td>95.2±18.4</td>
<td>0.526</td>
</tr>
</tbody>
</table>

**TABLE 1: Lab parameters at presentation (±SEM)**

SEM, standard error of the mean; AKI, acute kidney injury; RRT, renal replacement therapy; INR, international normalized ratio.

*Renal replacement therapy (hemodialysis or continuous renal replacement therapy).
Prior PCI | 7 (33.3) | 6 (46.2) | 0.459
---|---|---|---
Tobacco use in prior 12 months | 7 (33.3) | 5 (38.5) | 0.825
Medication use prior to presentation
ACEi/ARB | 10 (47.6) | 6 (46.2) | 0.985
Beta blocker | 11 (52.4) | 5 (38.5) | 0.515
Calcium channel blocker | 0 (0) | 3 (23.1) | 0.044
Diuretics | 10 (47.6) | 4 (30.8) | 0.341
Aspirin | 11 (52.4) | 5 (38.5) | 0.515
NSAID | 1 (4.76) | 0 (0) | 0.283
Digoxin | 0 (0) | 1 (7.69) | 0.168
Statin | 12 (57.1) | 5 (38.5) | 0.316
Metformin | 2 (9.52) | 3 (23.1) | 0.455

TABLE 2: Baseline characteristics
AKI, acute kidney injury; HTN, hypertension; DM, diabetes mellitus; CVA/TIA, cerebrovascular accident/transient ischemic attack; GFR, glomerular filtration rate; PCI, percutaneous coronary intervention; ACEi/ARB, angiotensin-converting enzyme inhibitor/angiotensin receptor blocker; NSAID, non-steroidal anti-inflammatory drug.

AKI was associated with higher 30-day mortality compared to patients with no AKI in our cardiogenic shock cohort [52.4% (11/21) vs. 30.8% (4/13), p=0.01; relative risk [RR] 6.16 (95% confidence interval [CI] 1.41-26.7)]. Thirty-day mortality was similar in patients with no AKI and AKI plus RRT [30.8% (4/13) vs. 22.2% (2/9), p=0.48; RR 2.25 (95% CI 0.22-22.1)]. Patients with AKI minus RRT had a higher 30-day mortality than patients with no AKI [75.0% (9/12) vs. 30.8% (4/13); p=0.03; RR 6.75 (95% CI 1.16-39.2)]. Outpatient use of calcium channel blockers and hemoglobin greater than 11 g/dL were associated with decreased likelihood of AKI at presentation (Table 1). Serum creatinine at presentation was higher in patients who received RRT for AKI; serum lactic acid, hemoglobin, and arterial pH were similar across all groups (Table 2).

Discussion
AKI in severe system illness and particularly cardiogenic shock is associated with poor outcomes [7,8]. AKI in cardiogenic shock has been postulated to be caused by venous congestion and reduced renal perfusion [9].

Our single-center retrospective study evaluated outcomes of cardiogenic shock in patients on Impella-CP complicated by AKI with and without RRT. We found that patients with AKI had a similar 30-day mortality when receiving RRT compared to patients without AKI at presentation.

It seems intuitive that early recognition of AKI in cardiogenic shock and aggressive treatment with RRT may improve outcomes, but data are conflicting. Aside from cardiogenic shock...
populations, the Impella-CP device is shown to be protective against AKI in patients undergoing high-risk percutaneous coronary intervention [10]. In a single-center study, patients on extracorporeal membrane oxygenation for cardiogenic shock with associated AKI did not have improved short-term survival with RRT [11]. Whether this cohort was more critically ill than the patients we studied is not known.

The limitations of our study are acknowledged: small sample size, retrospective examination, single-center review, and wide confidence intervals.

Conclusions
AKI is an independent predictor of mortality in cardiogenic shock. Based on our single-center, retrospective comparison of cardiogenic shock supported by Impella-CP, 30-day mortality was similar between patients with no AKI and those who received RRT for AKI. Early initiation of RRT for AKI coupled with improved left ventricular unloading with Impella-CP may offset the detrimental effects of AKI in cardiogenic shock. Our study provides preliminary evidence to warrant further investigation on early RRT for AKI in cardiogenic patients supported with Impella-CP.

Additional Information

Disclosures

Human subjects: Consent was obtained by all participants in this study. Albany Medical Center IRB issued approval 5072. Submitted retrospective study was conducted after institutional IRB approval. Animal subjects: All authors have confirmed that this study did not involve animal subjects or tissue. Conflicts of interest: In compliance with the ICMJE uniform disclosure form, all authors declare the following: Payment/services info: All authors have declared that no financial support was received from any organization for the submitted work. Financial relationships: All authors have declared that they have no financial relationships at present or within the previous three years with any organizations that might have an interest in the submitted work. Other relationships: All authors have declared that there are no other relationships or activities that could appear to have influenced the submitted work.

References