

**Original Paper**

# The Impact of Integrating Nephrologists into the Postoperative Cardiac Intensive Care Unit: A Cohort Study

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**Key Words**

Acute kidney injury · Intensive care unit · Early nephrology

**Abstract**

**Background/Aims:** We evaluated the potential preventive effect of Nephrology On-Site (i.e. nephrologists integrated into the postoperative cardiac intensive care unit, ICU, team) versus Nephrology On-Demand (i.e. nephrology consultation depending on intensivist criteria) in the ICU on in-hospital outcomes. **Methods:** This was a retrospective cohort study comparing outcomes during 2 consecutive time periods: from March 1, 2009 to February 28, 2010 with Nephrology On-Demand, and from March 1, 2010 to February 28, 2011 with Nephrology On-Site. Adult patients admitted to the postoperative cardiac ICU in an academic hospital in Mexico City were eligible. Patients with chronic kidney disease stage 5 or minimally invasive procedures were excluded. **Results:** We analyzed 1,096 patients, 558 and 538 in the respective periods. The patients were  $52.4 \pm 16.2$  years old, 56.1% were males, 17.2% had diabetes and 37.6% had hypertension. Further, the patients' median Euroscore was 5 (3–5) and their median Thakar score was 3 (2–4). With Nephrology On-Site, we observed a lower incidence of acute kidney injury [AKI; 25.7 vs. 31.9%,  $p = 0.02$ ; adjusted OR 0.71 (0.53–0.95),  $p = 0.02$ ], lower in-hospital mortality among patients with severe AKI [34.1 vs. 55.9%,  $p = 0.06$ ; adjusted OR 0.33 (0.12–0.95),  $p = 0.04$ ] and higher renal recovery [61.0 vs. 35.3%,  $p = 0.03$ ; adjusted OR 3.57 (1.27–10.11),  $p = 0.02$ ]. No differences were found in the length of stay at the ICU and mechanical ventilation. **Conclusion:** Integrating nephrologists into the postoperative cardiac ICU team was associated with a lower incidence of AKI. Patients who developed severe AKI had lower in-hospital mortality and higher renal recovery.

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## Introduction

The incidence of acute kidney injury (AKI) after cardiac surgery defined using the internationally accepted RIFLE [1] and AKIN [2] criteria has been reported to be around 30%, contributing to a mortality rate of 30–50% as the AKI stage increases [3–7].

Delayed recognition leads to delayed implementation of therapeutic strategies, reducing their chance to be effective [8]. Moreover, nephrologists are usually not involved until severe AKI is established. The impact of early nephrology consultation has been evaluated either by the time since admission to the intensive care unit (ICU) [9], by the increase in serum creatinine (SCr) at consultation [10] or by standardized interventions once AKI has been recognized through an automatic detector [11], showing general better outcomes. However, the preventive role of nephrologists in routinely screening postoperative cardiac patients even prior to the development of AKI has not been explored yet.

The aim of our study was to compare in-hospital outcomes before and after the integration of nephrologists into the postoperative cardiac ICU team.

## Methods

### *Study Design and Population*

We conducted a retrospective cohort study comparing outcomes during 2 consecutive time periods in a single center – the Instituto Nacional de Cardiología Ignacio Chavez, a tertiary referral public academic hospital in Mexico City. We included all adult patients who underwent open cardiac or thoracic large vessel surgery from March 1, 2009 to February 28, 2011. Patients were excluded in case of preexisting chronic kidney disease stage 5, preoperative AKI, minimally invasive procedures, and death in the operating room or during the first 24 h after ICU admission. Patients were eliminated if information about in-hospital outcomes was missing.

The 2 study periods were from March 1, 2009 to February 28, 2010 when nephrology consultation depended on intensivist criteria (Nephrology On-Demand) and from March 1, 2010 to February 28, 2011 when nephrologists were integrated into the postoperative cardiac ICU team (Nephrology On-Site). The Nephrology On-Site scheme required the daily presence of nephrology fellows actively participating in clinical rounds with intensivists, followed by a clinical round with the attending nephrologists. The main tasks among others were to identify and control risk factors, assess fluid balances, adjust drug dosing and nutritional support, and monitor the RIFLE/AKIN criteria. Decisions on extracorporeal renal support therapy (RST) were taken by consensus within the ICU team on an individual basis without standardized criteria.

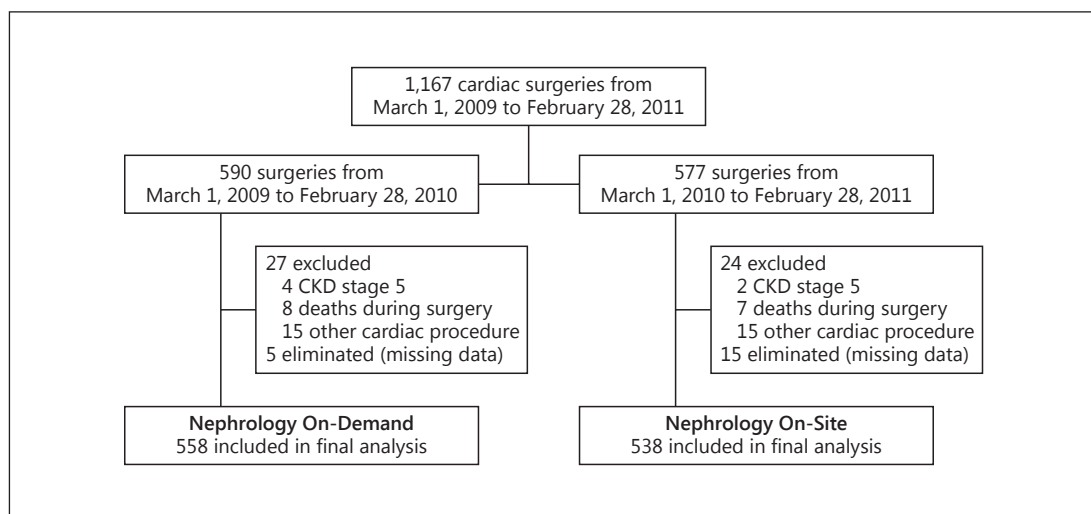
### *Data Collection and Outcomes*

Pre-, trans- and postoperative data were systematically collected from the patients' medical records. Data on urine output, fluid balance and medications were not available. No specific medical record was generated that could delimit nephrology interventions from the rest of the postoperative cardiac ICU interventions. The main outcome was the incidence of AKI. Secondary outcomes were in-hospital mortality, length of stay at the ICU, length of mechanical ventilation and renal recovery at hospital discharge.

### *Definitions*

AKI was defined by the RIFLE/AKIN criteria based only on SCr within 7 days after cardiac surgery. The baseline SCr level was the lowest value within the previous 3 months before surgery or at hospitalization. Severe AKI was defined as RIFLE-I/AKIN-2 or RIFLE-F/AKIN-3. Chronic kidney disease was defined as an estimated glomerular filtration rate of  $<60$  ml/min/1.73 m<sup>2</sup> calculated by the CKD-EPI equation [12].

Renal recovery at hospital discharge was defined as follows: (1) no recovery if patients continued on RST or if the AKI stage was equal to the maximum AKI; (2) complete recovery if SCr was less than 26.5  $\mu$ mol/l above the baseline value, and (3) partial recovery if the AKI stage was lower than the maximum AKI but the SCr level was at least 26.5  $\mu$ mol/l higher than the baseline value.



**Fig. 1.** Flow chart of study participants.

### Statistical Analysis

Quantitative variables were expressed as means  $\pm$  standard deviations or medians with 25th–75th percentiles and compared between the 2 time periods using Student’s t test or the Mann-Whitney U test. Categorical variables were expressed as proportions and compared using the  $\chi^2$  or Fisher’s exact test. A p value  $<0.05$  was considered statistically significant. Survival analysis was done by Kaplan-Meier curves and log-rank test to compare the 2 study periods. Multivariate regression models were performed using the ‘stepwise forward’ method. Patients with partial and complete recoveries were taken as one group in the multivariate analysis, which was repeated with and without death as a no-recovery group. An event/parameter ratio of  $\geq 10$  was targeted. Subgroup analysis was performed for AKI by preoperative risk, and for mortality and renal recovery by AKI stage. SPSS v16 was used for calculations.

## Results

During the study, 1,167 adult patients were admitted to the postoperative cardiac ICU. The final cohort included 1,096 patients, 558 in the first and 538 in the second period (fig. 1). Baseline and surgery-related characteristics are shown in table 1.

### Outcomes

#### Acute Kidney Injury

A significantly lower incidence of AKI was observed with Nephrology On-Site, mainly due to a reduction in patients with RIFLE-R/AKIN-1 (table 2). The number needed to treat was 16 patients. Survival analysis demonstrated a significant increase in survival free from AKI (fig. 2a), and multivariate analysis confirmed a lower adjusted OR for AKI (table 3). In the subgroup analysis, the benefit was higher in patients with intermediate risk (Thakar score 3–4; table 3). The onset of AKI was delayed from  $1.69 \pm 1.34$  to  $2.06 \pm 1.53$  days ( $p = 0.024$ ).

A nonsignificantly higher proportion of patients with severe AKI received RST with Nephrology On-Site (fig. 3). At the start of RST, a trend towards lower values for SCr [ $176.8$  ( $159.1$ – $221.0$ ) vs.  $212.2$  ( $159.1$ – $282.9$ )  $\mu\text{mol/l}$ ,  $p = 0.08$ ] and urea nitrogen [ $19.3$  ( $14.6$ – $24.6$ ) vs.  $26.4$  ( $13.9$ – $33.9$ )  $\text{mmol/l}$ ,  $p = 0.07$ ] was found with Nephrology On-Site.

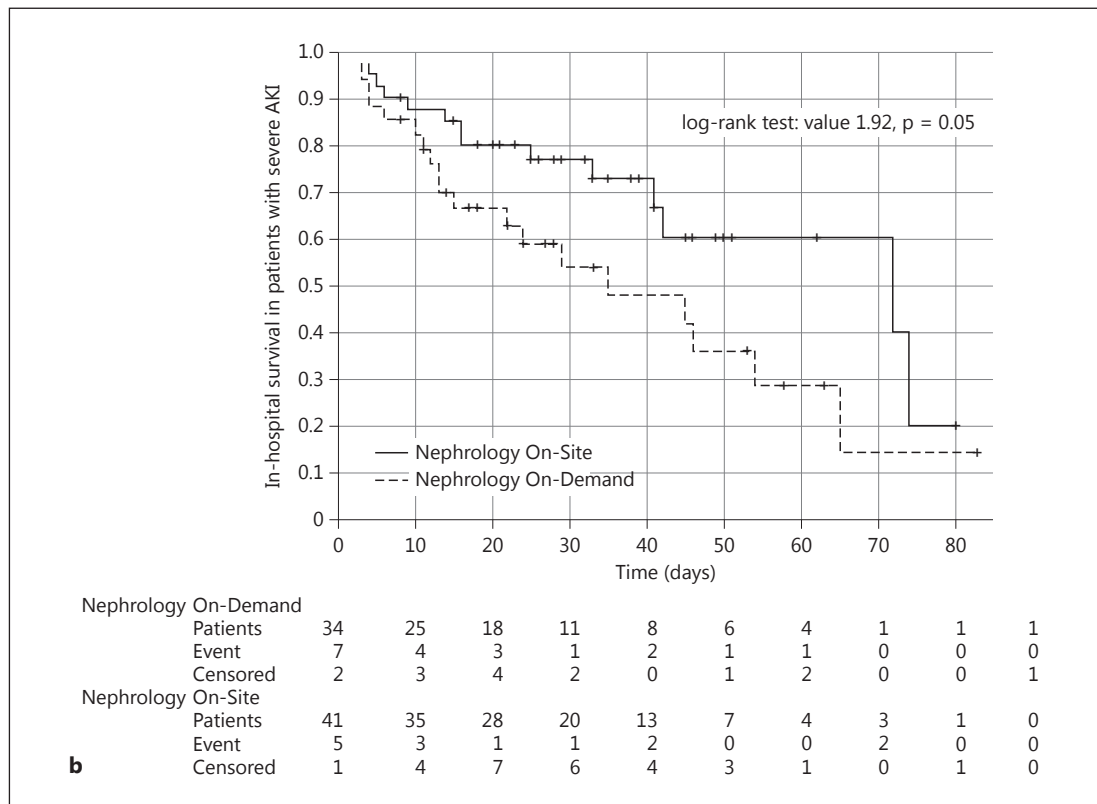
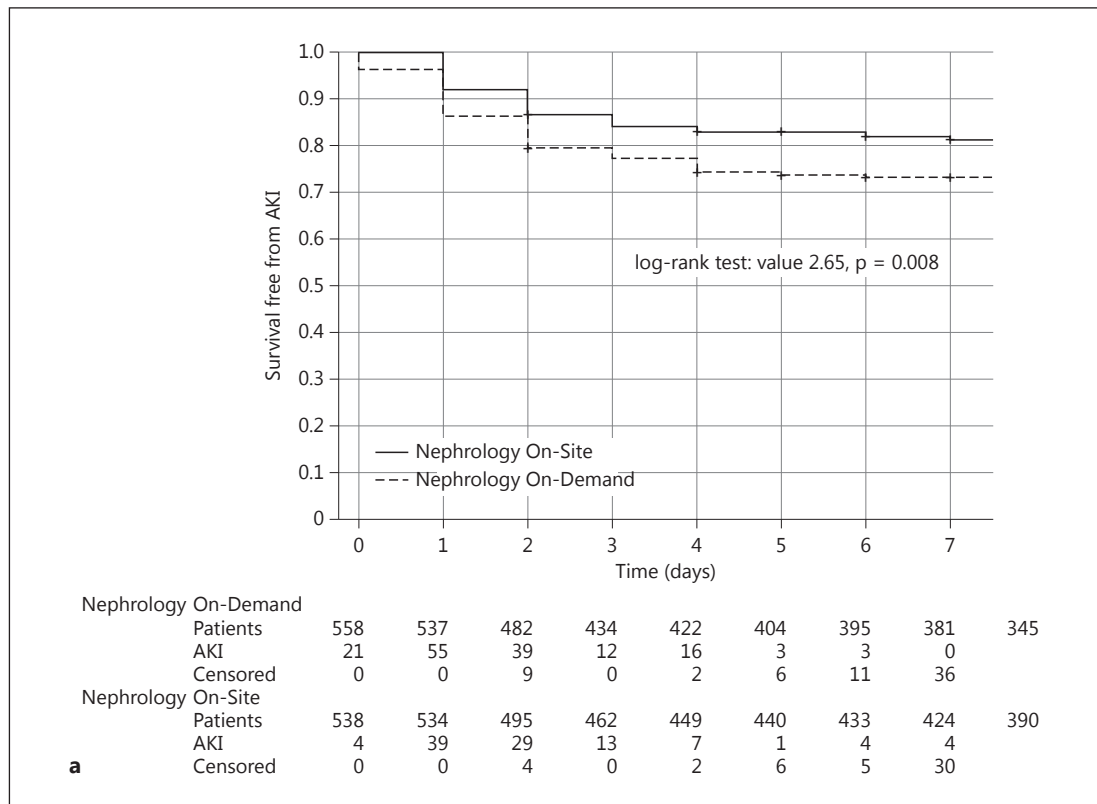
**Table 1.** Baseline and surgery-related characteristics

	Total (n = 1,096)	Nephrology On-Demand (n = 558)	Nephrology On-Site (n = 538)	p value
<i>General characteristics</i>				
Age, years	52.4±16.2	52.1±16.1	52.7±16.3	0.53
Weight, kg	68.4±12.7	69.3±12.4	67.6±13.0	0.03
Male gender	615 (56.1)	313 (56.1)	302 (56.1)	0.99
<i>Comorbidities</i>				
Diabetes mellitus	189 (17.2)	92 (15.5)	97 (18.0)	0.50
Hypertension	412 (37.6)	210 (37.6)	202 (37.5)	0.98
Chronic pulmonary disease	21 (1.9)	16 (2.9)	5 (0.9)	0.02
Cerebrovascular disease	50 (4.6)	24 (4.3)	26 (4.8)	0.67
Previous myocardial infarction	268 (24.5)	140 (25.1)	128 (23.8)	0.62
Previous cardiac surgery	135 (12.3)	59 (10.6)	76 (14.1)	0.07
Endocarditis	62 (5.7)	30 (5.4)	32 (5.9)	0.68
New York Heart Association III–IV	123 (11.2)	58 (10.4)	65 (12.1)	0.38
Critical preoperative state	120 (10.9)	63 (11.3)	57 (10.6)	0.71
Left ventricular ejection fraction, %	56.9±11.8	57.2±11.3	56.7±12.5	0.47
Pulmonary arterial pressure, mm Hg	35 (27–47)	36.50 (29–50)	35 (25–45)	<0.001
Glomerular filtration rate, ml/min/1.73 m <sup>2</sup>	91.7±22.7	88.8±23.6	94.7±21.5	<0.001
Chronic kidney disease	83 (7.6)	54 (9.7)	29 (5.0)	0.007
Baseline SCr, µmol/l	79.6 (61.9–97.2)	88.4 (70.7–106.1)	70.7 (61.9–88.4)	<0.001
<i>Scores</i>				
Euroscore	5 (3–5)	5 (3–6)	5 (3–7)	0.14
Thakar score	3 (2–4)	3 (2–4)	3 (2–4)	0.77
<i>Surgical variables</i>				
Emergency surgery	151 (13.8)	80 (14.0)	71 (13.2)	0.58
Type of surgery				0.22
Valvular	623 (56.8)	310 (55.6)	313 (58.2)	
Coronary bypass	234 (21.4)	118 (21.1)	116 (21.6)	
Coronary bypass + valvular	49 (4.5)	25 (4.5)	24 (4.5)	
Congenital heart disease	42 (3.8)	17 (3.0)	25 (4.6)	
Surgery on the aorta <sup>a</sup>	67 (6.1)	30 (5.4)	37 (6.9)	
Others <sup>b</sup>	148 (13.5)	88 (15.7)	60 (11.1)	
Extracorporeal circulation on-pump	973 (88.8)	480 (86.0)	493 (91.6)	0.003
Pump time, min	102 (75–136)	98 (73–129)	105 (76–140)	0.01
Aortic clamp time, min	70 (50–97)	68 (50–95.7)	72 (51–98)	0.48
Bleeding, ml	740 (500–1,140)	730 (500–1,132.5)	740 (506.5–1,147.5)	0.48
<i>Complications</i>				
Infection	121 (11.0)	70 (12.5)	51 (9.5)	0.11
Severe infection <sup>c</sup>	65 (5.9)	32 (5.7)	33 (6.1)	0.08
Reoperation	204 (18.6)	114 (20.4)	90 (16.8)	0.12

Data are shown as n (%), mean ± standard deviation or median (25th–75th percentiles).

<sup>a</sup> Could be in addition to other surgery. <sup>b</sup> Including surgery on the pericardium, isolated atrial septum defect and myxoma.

<sup>c</sup> Including pneumonia, mediastinitis and catheter-associated bacteremia.



**Fig. 2.** Kaplan-Meier curves comparing the two study periods. **a** Survival free from AKI (censoring for deaths) and **b** in-hospital survival in patients with severe AKI.

**Table 2.** Comparison of in-hospital outcomes

	Total (n = 1,096)	Nephrology On-Demand (n = 558)	Nephrology On-Site (n = 538)	p value
<i>AKI</i>	316 (28.8)	178 (31.9)	138 (25.7)	0.02
RIFLE-R/AKIN-1	241 (22.0)	144 (25.8)	97 (18.0)	0.002
Severe AKI	75 (6.8)	34 (6.1)	41 (7.6)	0.32
RST	56 (5.1)	23 (4.1)	33 (6.1)	0.13
<i>Renal recovery from AKI<sup>a</sup></i>	216 (68.4)	124 (69.7)	92 (66.7)	0.57
RIFLE-R/AKIN-1	179 (74.3)	112 (77.8)	67 (69.1)	0.13
Severe AKI	37 (49.3)	12 (35.3)	25 (61.0)	0.03
RST	23 (41.1)	6 (26.1)	17 (51.5)	0.06
<i>Mortality</i>	76 (6.9)	46 (8.2)	30 (5.6)	0.08
AKI	50 (15.8)	29 (16.3)	21 (15.2)	0.80
RIFLE-R/AKIN-1	17 (7.1)	10 (6.9)	7 (7.2)	0.94
Severe AKI	33 (44.0)	19 (55.9)	14 (34.1)	0.06
RST	30 (53.6)	16 (69.6)	14 (42.4)	0.04

Data are shown as n (%). <sup>a</sup> Including complete and partial renal recovery.

**Table 3.** Multivariate logistic regression analysis for the effect of Nephrology On-Site on in-hospital outcomes, and subgroup analysis

Outcome	Unadjusted OR	p value	Adjusted OR	p value
<i>AKI</i>	0.80 (0.63–1.02)	0.07	0.71 (0.53–0.95) <sup>c</sup>	0.02
Thakar score 0–2			0.62 (0.36–1.08) <sup>c</sup>	0.09
Thakar score 3–4			0.56 (0.36–0.88) <sup>c</sup>	0.01
Thakar score ≥5			1.18 (0.59–2.39) <sup>c</sup>	0.64
<i>Renal recovery<sup>a</sup></i>	0.74 (0.47–1.15)	0.18	1.00 (0.61–1.65) <sup>d</sup>	0.99
Severe AKI			3.57 (1.27–10.11) <sup>d</sup>	0.02
RST			3.78 (1.01–14.21) <sup>d</sup>	0.04
<i>Renal recovery<sup>a,b</sup></i>	0.72 (0.46–1.12)	0.14	0.94 (0.57–1.54) <sup>d</sup>	0.81
Severe AKI			3.27 (1.16–9.17) <sup>d</sup>	0.03
RST			3.48 (0.92–13.13) <sup>d</sup>	0.07
<i>Mortality</i>	0.96 (0.76–1.21)	0.72	0.49 (0.29–0.83) <sup>e</sup>	0.008
AKI			0.67 (0.34–1.32) <sup>e</sup>	0.25
Severe AKI			0.33 (0.12–0.95) <sup>e</sup>	0.04
RST			0.35 (0.10–1.18) <sup>e</sup>	0.09

<sup>a</sup> Partial or complete recovery. <sup>b</sup> Deaths included in the no-recovery group. <sup>c</sup> Adjusted for age, weight, reoperation, mechanical ventilation for more than 24 h, hypertension, previous myocardial infarction, baseline SCr and Thakar score; however, extracorporeal circulation time was tested but it did not affect the model. <sup>d</sup> Adjusted for severe infections, baseline SCr and Thakar score. <sup>e</sup> Adjusted for weight, severe infections and Euroscore.

### Renal Recovery

Renal recovery was higher with Nephrology On-Site among patients with severe AKI or RST (table 2). Similar results were obtained when death was included in the no-recovery group (data not shown). These results were confirmed in the multivariate analysis (table 3).

### Mortality

Nephrology On-Site was associated with a trend towards lower in-hospital mortality, mainly among patients with severe AKI or RST (table 2). The survival curve showed a noticeable difference at 30 and 60 days in patients with severe AKI (fig. 2b). Again, these observations were reinforced in the multivariate analysis (table 3).

### Length of Stay and Mechanical Ventilation

The general length of stay at the ICU was 4 (3–6) days, which increased to 5 (3–7) days with RIFLE-R/AKIN-1, to 13 (8–23) days with severe AKI and to 15 (9.3–25.8) days with RST. There was no difference between the 2 periods. The length of mechanical ventilation was higher with Nephrology On-Site [2 (2–4) vs. 2 (1–3) days,  $p < 0.001$ ], although it lost its significance in patients with AKI. The duration of mechanical ventilation in patients with RIFLE-R/AKIN-1 was 3 (2–4) versus 3 (2–5) days ( $p = 0.06$ ), and in those with severe AKI it was 9 (5–20) versus 8 (3–13) days ( $p = 0.20$ ), respectively. A shift from 1 day to 2 days of mechanical ventilation was noticed in a subgroup of patients with Nephrology On-Site who had a higher proportion of previous cardiac surgery and valvular, combined and on-pump techniques (data not shown). The extreme values prevented the transformation of these variables to fit a normal distribution, so multivariate linear regression analysis was not performed.

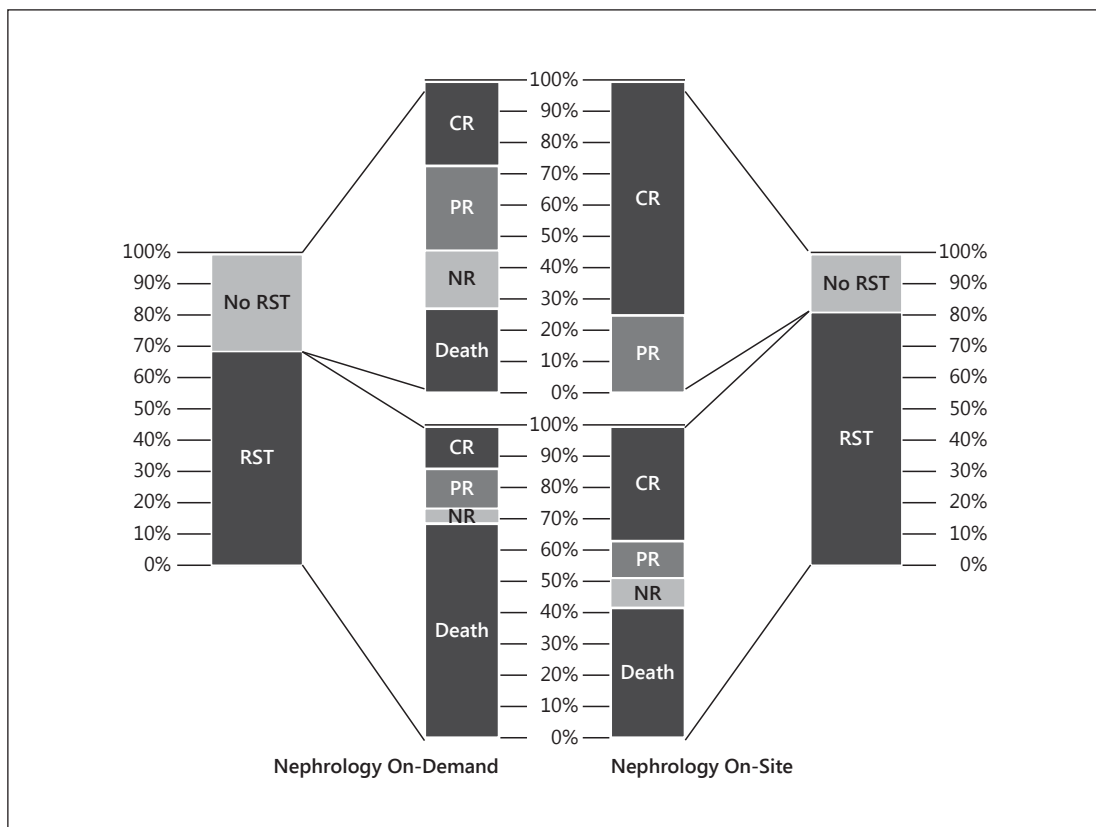
## Discussion

We found a significant reduction in AKI with Nephrology On-Site and a baseline AKI incidence with Nephrology On-Demand of 31.9%, similar to previous studies [3, 4]. As expected, a higher positive impact was achieved in patients with intermediate risk. Of note, the cutoff points for Thakar scores [13] were adapted to our population, RST in 1.54% corresponding to scores 0–2, in 3.52% to scores 3–4 and in 18.56% to scores  $\geq 5$ .

With Nephrology On-Site, there was a trend towards a higher proportion of patients receiving RST, together with some parameters of earlier dialysis, even though the proportion of severe AKI was lower than previously reported [4]. Although no specific criterion was used to determine when RST should be started, in our center patients with RIFLE-I/AKIN-2 plus an unfavorable state (mechanical ventilation, hemodynamic instability, inputs higher than urine output or sepsis) are usually considered for RST, and with Nephrology On-Site this recognition process might be accelerated. While patients requiring RST usually are at higher risk, as an early intervention it could contribute to better outcomes [14]. Nevertheless, in our study, lower values of SCr and urea nitrogen at the start of RST were not associated with renal recovery or survival. It is clear that supporting patients with less severe AKI and those who are maybe less critically ill would produce a bias towards survival benefit; however, our outcomes among patients with severe AKI were better with Nephrology On-Site with or without RST (fig. 3). Without RST, 45.4% of patients either died or did not recover renal function with Nephrology On-Demand versus 0.0% with Nephrology On-Site (fig. 3). In this sense, daily evaluation not only consists of starting RST early, but also of preventing and choosing the right patient for RST.

Nephrology On-Site resulted in an increased renal recovery rate in patients with severe AKI or RST. In a study by Hobson et al. [15], 45% of patients undergoing cardiothoracic





**Fig. 3.** Renal outcomes and mortality in patients with severe AKI in both periods. CR = Complete renal recovery; PR = partial renal recovery; NR = no renal recovery.

surgery were dialysis dependent at hospital discharge and 21% had complete recovery, while in our study 15.4% of patients were dialysis dependent in both periods; further, we found a nonsignificant increase in the complete recovery rate from 42.9 to 63.2%. It is remarkable that, even though in both study periods patients who required RST were treated by nephrologists, the collaborative work might have contributed to adjust the therapeutic strategies more efficiently for multiorgan benefits.

There was a reduction in mortality among patients with severe AKI from 55.9 to 34.1%, and among patients with RST from 69.6 to 42.4%, which is similar to results reported in the literature [3, 6, 16]. Of course, it is to be expected that high-quality centers with baseline mortality rates lower than ours might obtain a lower proportional benefit with the same approach.

Equivalent approaches in the literature have shown similar survival benefits. Higher adherence to evidence-based practice and lower rates of complications have been reported when intensivists are present in the ICU 24 h a day versus only on demand [17]. In systematic reviews, the constant presence of intensivists or obligatory consultation in the ICU has shown relative reduction rates for mortality of around 13.8–60% [18, 19], while in our center this relative reduction was between 36.3 and 39.0% among the different AKI stages. This survival benefit seems plausible as it is associated with both a lower incidence of AKI and a higher renal recovery rate.

In our study, some variables associated with worse outcomes had an unbalanced distribution. Lower body weight, severe infections such as pneumonia as well as mechanical venti-



lation were more prevalent in patients with Nephrology On-Site, which could have played a major role reducing benefits in the length of stay at the ICU. Specifically, surgical techniques might explain the increase in mechanical ventilation.

The limitations of our study include that missing information on urine output and fluid balance could have influenced AKI diagnosis; however, as preventing and treating fluid overload were part of the tasks, we hypothesize that with Nephrology On-Demand the percentage of fluid overload could have been higher, and therefore also the real incidence of AKI. In this sense, 29.6% of patients with Nephrology On-Demand had a reduction in their SCr levels from baseline to day 1 versus 21.9% with Nephrology On-Site ( $p = 0.004$ ), which could indicate a positive fluid balance. Other missing information such as medications including diuretics and severity scores would have allowed us to assess factors potentially associated with our results. As this was a retrospective/observational study, we could not control or assess the specific interventions with Nephrology On-Site; we recognize that these positive results could have been achieved by the original ICU team, simply by the effect of being observed and becoming sensitive towards renal care, but in our perspective this is still a positive achievement. Although not originally planned, we could have influenced the anesthesiology team towards avoiding the intraoperative use of NSAIDs and diuretics, which could explain the change in the early onset of AKI.

In the 90's, Ronco and Bellomo [20] emphasized that the complexity of AKI in ICUs demands nephrologists and intensivists to join forces in a new academic structure called Critical Care Nephrology, and an example of this new structure is the 'Vicenza Model' [21]. This multidisciplinary multitarget model with low costs by only optimizing human resources can have benefits beyond clinical outcomes through promoting cordial relationships among specialists, improving academic programs for fellows and potentiating research work.

## Conclusion

Integrating nephrologists into the postoperative cardiac ICU team decreased the incidence of AKI and promoted renal recovery and survival among patients with severe AKI. This approach can have multiple benefits for academic centers.

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## Disclosure Statement

All authors declare no conflict of interest.

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