Impact of renal dysfunction on surgical outcomes in patients with aortic dissection

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Abstract
Preoperative renal dysfunction is associated with mortality in patients who undergo coronary artery bypass graft and valve surgery. Thus, urgent surgical repair should be considered for all patients with TAAD. In-hospital mortality for patients with renal dysfunction exhibited a lower 90-day survival rate than did patients without the condition (P = 0.005).

Preoperative renal dysfunction may have a critical role in the surgical outcomes of patients with TAAD. Additional large-scale investigations are warranted.

Abbreviations: AKI = acute kidney injury, CABG = coronary artery bypass graft, CI = confidence interval, CKD = chronic kidney disease, COPD = chronic obstructive pulmonary disease, Cr = creatinine, CT = computed tomography, eGFR = estimated glomerular filtration rate, ICU = intensive care unit, IRAD = International Registry of Acute Aortic Dissections, NA = not applicable, TAAD = type A aortic dissection.

Keywords: prognosis, renal dysfunction, risk factors, type A aortic dissection (TAAD)

1. Introduction
The mortality rate of patients with acute unoperated type A aortic dissection (TAAD) increases 1% to 2% every hour within the first 2 days of an acute event. Thus, urgent surgical repair should be considered for all patients with TAAD. In-hospital mortality for patients managed surgically was found to be 27%, compared with 56% for those managed medically.[11] Refractory pain, an age of over 70 years, and the absence of chest pain on admission have been reported as predictors of death. Aortic rupture was the most common cause of death identified in the International Registry of Acute Aortic Dissections (IRAD).[2] In the past decade, neither the symptoms nor physical findings of TAAD have changed substantially. Advancements in diagnostic computed tomography (CT) have significantly reduced in-hospital mortality by facilitating early surgical intervention.[3] Evidence suggests that acute kidney injury (AKI) following dissection worsens the prognosis and prolongs hospitalization.[4,5] However, the potential effects of preoperative renal dysfunction have not been clearly explored in the literature. Recently, preoperative organ malperfusion was proved to affect the outcome in these patients.[6] Therefore, our investigation evaluated the impact of preoperative renal dysfunction on mortality in patients with TAAD.

2. Methods
2.1. Study participants and design
The data of patients with TAAD treated between January 2005 and December 2010 were extracted for analysis from a prospectively collected database. This study design was approved by the institutional review board of Chang Gung Memorial Hospital (201601407B), and the need for individual consent was waived. We excluded patients who had undergone prior cardiac surgery, had end-stage renal disease, or died within 1 day after...
surgery. All patients had received contrast CT before surgery. The final cohort comprised 159 consecutive patients who received dissection repair in a single tertiary referral hospital. The patients were divided into 2 groups according to their serum creatinine (Cr) levels.

2.2. Data collection and definition

The baseline characteristics and demographic data of the patients were extracted from the database. The laboratory data comprised the preoperative values recorded on the operation date. Patient surgical details were also obtained from the database for analysis. Outcomes in terms of de novo dialysis, mortality, major complications, blood transfusions, and hospital stay were compared between the 2 renal function groups. Postoperative care was standardized in our intensive care unit (ICU) by 2 critical care specialists. Shock was defined as systolic pressure less than 90 mmHg.

2.3. Statistical analysis

Continuous variables were summarized as mean and standard deviation and were compared using the Student t test between the 2 groups defined according to preoperative Cr (<1.5 vs ≥1.5). The distribution of categorical variables in each group was compared using Fisher exact test. Associations of characteristics and operative conditions with risk of in-hospital mortality were assessed using univariate and multivariable logistic regression models. Variables with a P value less than .20 in the univariate models were subsequently introduced into the multivariable model with a stepwise selection procedure. Finally, Kaplan-Meier survival curves of 90-day mortality were plotted together with the log-rank test to compare the 2 groups. All statistical tests were 2-tailed and a P value less than .05 was considered significant. Data were analyzed using SPSS 22.0 software (IBM SPSS, Armonk, NY: IBM Corp).

3. Results

3.1. Study population characteristics and surgical details

One hundred and fifty-nine patients were examined in this study. The mean age was 57.5 years (SD = 13.2 years), and 27.0% of the patients were female. Thirty-two of the patients (20.1%) had preoperative Cr of 1.5 mg/dL or more. The baseline preoperative demographic and clinical characteristics are listed in Table 1. The 2 groups did not differ significantly in their baseline characteristics, namely age, sex, diabetes mellitus, hypertension, silent stroke, cardiac ejection fraction, chronic obstructive pulmonary disease (COPD), chronic kidney disease (CKD) and cirrhosis. Renal artery involvement was found in 36.1% in our cohort. The patients with Cr less than 1.5 mg/dL were more likely to have renal artery involvement than the others (40.9% vs 16.1% P = .012, respectively). Laboratory data for hemoglobin, platelet count, and prothrombin time also did not differ significantly between the groups, with the exception of Cr level.

The surgical details are shown in Table 2. They indicate that the patients with Cr less than 1.5 mg/dL were more likely to receive aortic arch repair (P = .043), which might reflect a more aggressive surgical strategy adopted for patients with a stable preoperative condition. No significant differences were present for additional coronary artery bypass graft, intraoperative bypass time, clamp time, arrest time, or brain protection strategy.

3.2. Patient outcomes in different groups

In-hospital mortality was significantly higher in the group whose Cr was 1.5 mg/dL or more (34.4% vs 12.6%, P = .007). Other associated outcomes are listed in Table 3. However, no significant difference was observed in postoperative ventilator time, tracheostomy rate, ICU stay, hospital stay, unexpected bleeding check, or wound infection. In the first 48 hours after surgery, the transfusion amount did not differ significantly between the groups. The incidence of de novo dialysis in the patients with Cr of 1.5 mg/dL or more before surgery was also significantly higher.

Table 1
Preoperative demographic data and clinical characteristics.

| Variable                  | All patients (n = 159) | Cr < 1.5 mg/dL (n = 127) | Cr ≥ 1.5 mg/dL (n = 32) | P Value |
|---------------------------|------------------------|--------------------------|-------------------------|---------|
| Demographic data          |                        |                          |                         |         |
| Age (y)                   | 57.5±13.2              | 56.5±12.6                | 61.5±14.6               | .055    |
| Female sex, n (%)         | 43 (27.0)              | 36 (28.3)                | 7 (21.9)                | .513    |
| Body weight (kg)          | 73.9±13.3              | 73.7±13.7                | 74.5±12.1               | .773    |
| Diabetes mellitus, n (%)  | 13 (8.2)               | 9 (7.1)                  | 4 (12.5)                | .301    |
| Hypertension, n (%)       | 113 (71.1)             | 90 (70.9)                | 23 (71.9)               | 1.000   |
| CKD, n (%)                | 123 (77.8)             | 95 (74.8)                | 28 (90.3)               | .089    |
| Silent stroke, n (%)      | 15 (9.4)               | 11 (8.7)                 | 4 (12.5)                | .505    |
| Ejection fraction (%)     | 59.1±10.0              | 59.4±10.1                | 57.6±9.9                | .372    |
| Smoking, n (%)            | 68 (42.8)              | 53 (41.7)                | 15 (46.9)               | .690    |
| COPD, n (%)               | 4 (2.5)                | 4 (3.1)                  | 0 (0.0)                 | .584    |
| Liver cirrhosis, n (%)    | 1 (0.6)                | 1 (0.8)                  | 0 (0.0)                 | 1.000   |
| Hemopericardium, n (%)    | 61 (38.4)              | 46 (36.2)                | 15 (46.9)               | .311    |
| Shock, n (%)              | 6 (3.8)                | 4 (3.2)                  | 2 (6.3)                 | .348    |
| Lab data                  |                        |                          |                         |         |
| Cr (mg/dL)                | 1.21±0.42              | 1.05±0.22                | 1.84±0.44               | <.001   |
| Hemoglobin (g/dL)         | 13.4±1.8               | 13.5±1.7                 | 13.3±1.9                | .547    |
| Platelet (x10^4/µL)       | 180.0±46.8             | 183.4±65.4               | 166.8±61.3              | .197    |
| Prothrombin time (second) | 1.10±0.14              | 1.10±0.12                | 1.13±0.19               | .261    |

CKD = chronic kidney disease, COPD = chronic obstructive pulmonary disease.
these literatures, they investigated renal dysfunction as complication or postoperative outcome. A retrospective study by Kato et al. showed that TAAD is a risk factors for AKI after aortic dissection, and AKI increased the all-cause mortality. Another study by Pismis et al showed that thoracic aortic endograft has a significant rate of renal dysfunction. Some literatures revealed renal ischemia as risk factor of surgical outcome of acute TAAD. A retrospective review by Kazui et al demonstrated that renal ischemia is one of preoperative dissection-related complications and comorbidities, which significantly affect early and late survival rates after surgical treatment of acute TAAD. Our study focuses on the impact of pre-operative renal function to surgical outcome of patient with aortic dissection, which has not been investigated thoroughly. In the present study, we identified preoperative Cr level as an independent predictor for in-hospital mortality. The observed mortality rate of 17.0% is similar to those reported in prior publications.

3.3. Association between Cr and In-hospital mortality

To determine the risk factors for mortality, baseline characteristics and operative factors were analyzed using univariate logistic regression models; the results are shown in Table 4. The following variables were found to be significant: preoperative Cr of 1.5 mg/dL or more, preoperative shock, number of transfused platelets, prothrombin time, and intraoperative bypass time. The multivariable model indicated that preoperative renal function (odds ratio [OR], 3.79; 95% confidence interval [CI], 1.64–8.77), preoperative hypotension (OR, 8.75; 95% CI, 2.83–27.02), and bypass time (OR, 1.008; 95% CI, 1.003–1.013) were independently associated with risk of in-hospital mortality. Figure 1 presents the 90-day Kaplan–Meier survival curves for both groups, revealing that the patients with a higher Cr level had a lower 90-day survival rate (P = 0.005).

4. Discussion

The relationship of renal dysfunction and surgery for aortic dissection have been discussed in many literatures. In most of

| Table 2 |
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| Operative and periprocedure data. |
| Variable | All patients (n = 159) | Cr < 1.5 mg/dL (n = 127) | Cr ≥ 1.5 mg/dL (n = 32) | P Value |
| --- | --- | --- | --- | --- |
| Surgical method, n (%) | | | | |
| Bentall | 11 (6.9) | 9 (7.1) | 2 (6.3) | 1.000 |
| Aortic valve replacement | 4 (2.5) | 4 (3.1) | 0 (0.0) | 0.584 |
| Aortic arch | 16 (10.1) | 16 (12.6) | 0 (0.0) | 0.043 |
| Additional surgery, n (%) | | | | |
| CABG | 5 (3.1) | 4 (3.1) | 1 (3.1) | 1.000 |
| Intraoperative data (min) | | | | |
| Bypass time | 275.0 ± 85.1 | 277.3 ± 87.8 | 266.2 ± 53.2 | 0.493 |
| Clamp time | 175.9 ± 49.1 | 178.0 ± 52.0 | 167.6 ± 45.5 | 0.288 |
| Arrest time | 58.6 ± 25.9 | 60.2 ± 28.2 | 52.5 ± 12.1 | 1.133 |
| Brain protection strategy, n (%) | | | | |
| Selective antegrade cerebral perfusion | 100 (62.9) | 84 (66.1) | 16 (50.0) | 1.104 |
| Retrograde cerebral perfusion | 59 (37.1) | 43 (33.9) | 16 (50.0) | 1.104 |

CABG = coronary artery bypass graft.

| Table 3 |
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| Postoperative outcomes. |
| Variable | All patients (n = 159) | Cr < 1.5 mg/dL (n = 127) | Cr ≥ 1.5 mg/dL (n = 32) | P Value |
| --- | --- | --- | --- | --- |
| In-hospital mortality, n (%) | 27 (17.0) | 16 (12.6) | 11 (34.4) | 0.007 |
| Ventilator time (h) | 104.0 ± 41.2 | 101.1 ± 88.0 | 115.6 ± 103.7 | 0.423 |
| ICU stay (d) | 7.8 ± 6.5 | 7.8 ± 6.5 | 7.8 ± 6.9 | 0.999 |
| Hospital stay (d) | 25.1 ± 19.7 | 25.3 ± 19.5 | 24.4 ± 21.1 | 0.817 |
| Transient ischemic attack, n (%) | 4 (2.5) | 4 (3.1) | 0 (0.0) | 0.584 |
| De novo dialysis (%) | 18 (11.3) | 9 (7.1) | 6 (18.8) | 0.045 |
| Tracheostomy, n (%) | 6 (3.8) | 4 (3.1) | 2 (6.3) | 0.348 |
| Unexpected bleeding check, n (%) | 56 (32.5) | 31 (24.4) | 15 (46.8) | 0.351 |
| Wound infection, n (%) | 12 (7.5) | 10 (7.9) | 2 (6.3) | 1.000 |
| Transfusion 0–48 h (l) | 7.0 ± 5.0 | 7.1 ± 5.0 | 6.9 ± 5.3 | 0.894 |
| Packed red blood cells | 6.9 ± 4.9 | 6.8 ± 5.0 | 7.4 ± 4.3 | 0.628 |
| Fresh frozen plasma | 12.9 ± 8.3 | 12.9 ± 8.4 | 13.1 ± 7.8 | 0.917 |

ICU = intensive care unit.
### Table 4
Associated factors of risk of in-hospital mortality.

| Characteristics                  | Univariate model |          |          |          | Multivariable model |          |          |
|----------------------------------|------------------|----------|----------|----------|---------------------|----------|----------|
|                                  | OR               | 95% CI   | P Value  | OR       | 95% CI              | P Value  |
| Preop Cr ≥ 1.5 mg/dL             | 2.80             | 1.30–6.04| .009     | 3.79     | 1.64–8.77           | .002     |
| Age (yr)                         | 1.01             | 0.98–1.04| .465     |          |                     |          |
| Female sex                       | 0.63             | 0.24–1.68| .358     |          |                     |          |
| Body weight (kg)                 | 1.01             | 0.98–1.04| .509     |          |                     |          |
| Diabetes mellitus                | 1.35             | 0.41–4.50| .622     |          |                     |          |
| Hypertension                     | 0.52             | 0.24–1.12| .092     |          |                     |          |
| Silent stroke                    | 1.05             | 0.32–3.51| .934     |          |                     |          |
| Ejection fraction (%)            | 0.99             | 0.96–1.03| .585     |          |                     |          |
| Smoking                          | 1.36             | 0.64–2.91| .424     |          |                     |          |
| COPD                             | 1.87             | 0.52–13.84| .540   |          |                     |          |
| Liver cirrhosis                  | NA               | NA       | NA       |          |                     |          |
| Hemopericardium                  | 0.84             | 0.38–1.88| .674     |          |                     |          |
| Shock                            | 10.35            | 3.41–31.40| <.001   | 8.75     | 2.83–27.02          | <.001    |
| Cr, mg/dL                        | 1.93             | 0.98–3.81| .059     |          |                     |          |
| Hemoglobin (g/dL)                | 1.13             | 0.91–1.42| .276     |          |                     |          |
| Platelet (× 10^3/µL)             | 0.99             | 0.98–1.00| .012     |          |                     |          |
| Prothrombin time (s)             | 19.08            | 2.78–130.43| .003   |          |                     |          |
| Surgical method                  |                  |          |          |          |                     |          |
| Bentall                          | 1.07             | 0.25–4.54| .923     |          |                     |          |
| Aortic valve replacement         | NA               | NA       | NA       |          |                     |          |
| Aortic arch                      | 2.12             | 0.85–5.28| .108     |          |                     |          |
| Additional surgery               |                  |          |          |          |                     |          |
| CABG                             | 1.20             | 0.16–8.94| .862     |          |                     |          |
| Intra-operative data             |                  |          |          |          |                     |          |
| Bypass time                      | 1.006            | 1.002–1.01| .007   | 1.008    | 1.003–1.013         | .002     |
| Clamp time                       | 1.005            | 0.999–1.01| .130   |          |                     |          |
| Arrest time                      | 1.009            | 0.998–1.02| .103   |          |                     |          |
| Brain protection strategy        |                  |          |          |          |                     |          |
| Selective antegrade cerebral perfusion | Ref. | Ref. | Ref. | | | |
| Retrograde cerebral perfusion    | 1.05             | 0.48–2.29| .907     |          |                     |          |

CABG = coronary artery bypass graft, CI = confidence interval, COPD = chronic obstructive pulmonary disease, NA = not applicable, OR = odds ratio.

* included variables with a P value of <.2 in the univariate analyses and employed a stepwise selection method.

Figure 1. Kaplan-Meier survival curves for 90-day mortality according to the serum creatinine level.
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