



Impact of hypertension on short- and long-term survival of patients who underwent emergency surgery for type A acute aortic dissection

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Background: To explore the clinical characteristics and prognosis of patients with hypertension underwent emergency surgery for type A acute aortic dissection (TA-AAD).

Methods: The present study enrolled 712 consecutive patients diagnosed with TA-AAD and received aortic repair surgery at our hospital between January 2014 to December 2018. Clinical characteristics of enrolled patients were retrospectively reviewed. Patients were grouped by preexisting hypertension history and matched with propensity scores matching method. Patients' clinical characteristics were compared and analyzed before and after propensity scoring. To identify predictors for long-term mortality rate, Kaplan-Meier survival estimation and Cox proportional hazard analysis were performed.

Results: A total of 492 patients (69.1% of all patients in the cohort) were included in the hypertensive group and they had increased age and weight compared to patients in the non-hypertensive group. Between two groups, preoperative leukocyte count and serum creatinine level were found significant different ($P < 0.05$). After propensity scoring, 128 pairs (256 patients) were successfully matched. Our analysis showed that there was no significant difference of ventilation duration, 30-day mortality rate, intensive care unit stay and hospitalization time between two groups. However, our data suggested that hypertensive patients presented with less intra-operative aortic valve involvement. There was a significant difference in long-term survival rate ($P = 0.037$) between two groups. Cox regression analysis demonstrated that hypertension was an independent risk factor [hazard ratio (HR), 3.040; 95% confidence interval (CI), 1.124–8.227; $P = 0.029$].

Conclusions: Our data suggested that TA-AAD patients complicated with hypertension had increased age and weight compared to non-hypertensive patients. Concomitant hypertension identified upon hospital administration was an independent risk factor for long-term survival in TA-AAD patients while did not influence the 30-day mortality rate.

Keywords: Type A acute aortic dissection; hypertension; early outcomes; propensity score matching; long-term outcomes

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Introduction

Type A acute aortic dissection (TA-AAD) is a life-threatening disease, characterized by abrupt onset and catastrophic outcomes. The mortality rate of acute aortic dissection increases by 1% to 2% per hour after the first 24 to 48 h of symptom onset (1) and ranges from 8% to 34% even after prompt surgeries (2). Several risk factors associated with spontaneous dissection have been identified and hypertension is considered the most important one and presented in about 65–75% of all patients (3). In general population, hypertension accounts for 54% of the population-attributable risk of developing acute aortic dissection (4).

Surgical repair is the 1st line treatment for TA-AAD patients. Repair of TA-AAD can be restrained to replacement of the proximal arch sparing the aortic arch branch vessels or total arch replacement. Study showed that the prevalence of hypertension in thoracic aortic dissection patients increased in the past few years (5). However, even though the importance of hypertension in preoperative risk evaluation has been well examined, whether hypertension could influence short- and long-term survival rates after surgery has not been well studied. Therefore, the present study aimed to investigate the impact of pre-existing hypertension on short- and long-term mortality rates in patients with TA-AAD after emergency surgery. We present the following article in accordance with the STROBE reporting checklist (available at <http://dx.doi.org/10.21037/jtd-20-2336>).

Methods

Patients

A total of 712 consecutive patients diagnosed with TA-AAD and received aortic repair surgery at our hospital between January 2014 to December 2018 were initially screened for the study. The diagnosis of aortic dissection was made using computed tomography angiography upon admission to the hospital. This study was reviewed and approved by the institutional research ethics committee of Nanjing Drum Tower Hospital (No. BL2014004). Informed written consent was waived owing to the retrospective nature of the study. The study was conducted in accordance with the Declaration of Helsinki (as revised in 2013).

Exclusion criteria included a history of preoperative renal replacement therapy or death occurred within 24 hours after surgery. All patients were assigned to two groups based

on if they were also diagnosed with hypertension upon hospital admission.

Date collection

A standardized form was used to collect clinical information. Data included patient demographics, history of hypertension and aortic dissection disease, blood pressure (BP) at admission, aortic dissection features (blood supply of renal artery), peri-operational parameters including preoperative pericardial effusion, operation parameters and postoperative parameters like drainage volume 24 hours after surgery, mechanical ventilation time, 30-day mortality, length of intensive care unit (ICU) stay, and hospitalization time were collected.

Outcome variables

The primary endpoint was long-term survival rates with a median follow-up of 29 months. Secondary endpoints were early postoperative clinical characteristics and surgery complications.

Surgical techniques

A standard median sternotomy was routinely performed on patients with TA-AAD. After systemic heparinization, cardiopulmonary bypass (CPB) was established by arterial cannulation of the right axillary or femoral artery or the ascending aorta and venous cannulation of the right atrium. When patients were cooled to 18–20 °C (nasopharyngeal temperature) or 22–24 °C (bladder temperature), the blood circulation was stopped, and the flow rate was reduced to 3–5 mL/kg/min. After clamping of the ascending aorta, cardiac arrest was accomplished with cold cardioplegic solution. Bentall procedure or ascending aorta replacement was performed depending on the conditions of aortic regurgitation. Total arch replacement method was traditionally selected in the following scenarios: major dissection tear around the aortic arch or proximal descending aorta, or diameter of the aortic arch is greater than 4 cm on contrast computed tomography. Once the distal and transverse anastomoses were completed, antegrade perfusion was reinitiated. Systemic warming was then initiated, and the proximal reconstruction marked the completion of a standard CPB surgery. All patients were then transferred to ICU for routine monitoring.

Follow-up

Routine evaluation of enrolled patients' general health status was obtained through contacting with patients and their relatives through phone calls which was conducted every one year after discharging from the hospital. The follow-up information included BP, management of hypertension, current medication and deaths (if any, cause of death and date were recorded). All follow-up data was collected. The median follow-up period was 29 months, and follow-up information was obtained in 92.1% (569 of 618) of all enrolled patients.

Diagnosis of hypertension

For TA-AAD patients, the diagnosis of hypertension was made if the patient was diagnosed with hypertension by a physician previously or presented with at least 3 times of elevated BP (systolic BP greater than 140 mmHg and diastolic BP greater than 90 mmHg) on different days during hospitalization (6).

Statistical analysis

Data was analyzed with SPSS software (version 25.0, IBM Corp, Armonk, NY). Continuous and categorical variables were expressed as mean \pm standard deviation. The student *t*-test was used for Gaussian distributed continuous variables and the Mann-Whitney *U* test for non-Gaussian distributed variables. Categorical variables were compared with the χ^2 test or Fisher exact test.

To reduce selection bias, a one-to-one propensity score matching (PSM) method was applied to analyze the short-term outcomes, with hypertension as the dependent variable. And it was conducted between groups highlighted in the univariate analysis (preoperative characteristics that showed significant differences between two groups). PSM was based on one-to-one nearest neighbor matching method with a tolerance level on the maximum propensity score distance (calipers of width 0.1 standard deviations of the logit of the propensity score).

Kaplan-Meier survival estimation was applied for long-term survival analysis, whereas the Log-Rank test was applied for the calculation of significances for follow-up period. Univariate and multivariate Cox proportional hazard analysis were performed to determine independent predictors for late mortality, variables with a *P* value less than 0.1 on univariate analyses were included in the

multivariate model. All analyses were two-sided, and any *P*<0.05 was considered statistically significant.

Results

Demographic and clinical features

After excluding 3 patients requiring dialysis and 15 patients with death within 24 hours after the operation, a total of 712 patients were included in this study. Among these patients, 694 patients (97.5%) received emergency operation and 492 patients (69.1%) were diagnosed with hypertension. Patients' preoperative characteristics were presented in *Table 1*. Compared to normotensive patients, patients with hypertension were elder and heavier (*P*<0.001) and presented with elevated preoperative leucocyte and serum creatinine (sCr) levels.

In terms of operative variables, hypertensive patients required less intraoperative aortic valve management and experienced shorter aortic cross-clamp duration. While the posterior parallel circulation time and the deep hypothermic circulatory arrest duration were significantly longer in the hypertensive group, as shown in *Table 2*. In addition, normotensive patients were more likely to receive aortic valve management (*P*<0.001).

In order to better compare patients with similar baseline characteristics, propensity matching was performed in the entire cohort based on age, body mass index (BMI), preoperative leucocyte count and preoperative sCr. After matching, a total of 256 patients (128 pairs) remained for further analysis.

Short-term outcome

As indicated in *Table 3*, hypertensive patients had a significantly higher incidence of developing in-hospital complications compared to patients without hypertension. The 30-day mortality rate before propensity matching was 14.4% in the hypertension group and 8.6% in the normotensive patients (*P*=0.032). Other parameters, such as average ICU stay and hospital stay showed no significant differences between two groups.

However, there was no significant differences discovered between hypertensive and normotensive patients regarding 30-day mortality (9.4% *vs.* 10.9%, *P*=0.697) after propensity-matching suggested that the increased mortality observed in hypertensive group previously was largely due to the patients' baseline differences. Furthermore, our data suggested that there was no difference in variables like

Table 1 Comparison of preoperative variables before and after matching

Characteristics	Overall cohort			PSM cohort		
	Normotension (n=220)	Hypertension (n=492)	P value	Normotension (n=128)	Hypertension (n=128)	P value
Demographics						
Age (year)	49.6±14.6	53.8±12.3	<0.001	51.0±15.0	52.1±12.0	0.497
Male (%)	160 (72.7)	362 (73.6)	0.813	93 (72.7)	96 (75.0)	0.670
BMI (kg/m ²)	23.9±4.3	25.9±5.1	<0.001	24.1±4.3	25.0±5.4	0.129
Medical history						
Diabetes mellitus (%)	3 (1.4)	9 (1.8)	0.763	0 (0)	5 (3.9)	0.060
Previous cardiac surgery (%)	9 (4.1)	27 (5.5)	0.432	8 (6.3)	10 (7.8)	0.625
Previous coronary artery disease (%)	4 (1.8)	23 (4.7)	0.065	4 (3.1)	4 (3.1)	1.000
Cerebrovascular disease (%)	6 (2.7)	18 (3.7)	0.525	3 (2.3)	3 (2.3)	1.000
LVEF (%)	54.4±8.0	55.5±5.7	0.440	53.3±8.8	52.7±7.8	0.813
Pericardial effusion (%)	10 (4.5)	18 (3.7)	0.574	4 (3.1)	5 (3.9)	1.000
Duration of hypertension (y)	0	10.3±6.3	<0.001	0	9.8±5.6	<0.001
BP at admission						
SBP (mmHg)	117.4±11.6	147.3±21.6	<0.001	118.0±10.7	148.2±22.9	<0.001
DBP (mmHg)	72.4±8.1	87.9±12.6	<0.001	72.9±8.5	88.1±13.2	0.003
Mean BP (mmHg)	87.4±7.7	107.7±14.5	<0.001	87.9±7.5	108.1±15.1	<0.001
Preoperative laboratory data						
WBC (10 ⁹ /L)	10.5±4.1	11.2±4.2	0.041	10.5±4.1	10.9±4.1	0.396
sCr (μmol/L)	90.1±53.8	110.5±90.9	<0.001	89.0±60.0	102.6±82.6	0.132
PLT (10 ⁹ /L)	151.6±72.6	142.0±64.3	0.093	155.0±72.2	154.2±65.5	0.926
Fibrinogen (g/L)	2.6±1.3	2.6±1.3	0.453	2.6±1.2	2.6±1.3	0.977
Triglyceride (mmol/L)	1.1±1.0	1.3±1.1	0.057	1.1±1.1	1.2±0.9	0.439
Preoperative CTA						
Left renal artery: true lumen (%)	121 (65.4)	243 (63.3)	0.621	76 (70.4)	66 (63.5)	0.285
Right renal artery: true lumen (%)	139 (75.1)	292 (76.0)	0.813	80 (74.1)	78 (75.0)	0.877

Data presented as n (%); mean ± standard deviation. PSM, propensity score matching; BMI, body mass index; LVEF, left ventricular ejection fraction; ACEI, angiotensin-converting enzyme inhibitors; ARB, angiotensin receptor blocker; BP, blood pressure; SBP, systolic blood pressure; DBP, diastolic blood pressure; WBC, white blood cell; SCr, serum creatinine; PLT, platelet; CTA, computed tomography angiography.

ventilation duration, ICU and hospital stay time between two matched groups.

Long-term outcome

Seventy-three patients (14.8%) in the hypertensive group

and 21 patients (9.5%) in the normotensive group died during the hospitalization period. As a result, a total of 419 hypertensive patients and 199 normotensive patients were included in the follow-up. We identified 32 (6.5%) deaths that occurred in the hypertension group and 9 (4.1%) in the normotensive group. The causes of late death are presented

Table 2 Comparison of operative variables before and after matching

Characteristics	Overall cohort			PSM cohort		
	Normotension (n=220)	Hypertension (n=492)	P value	Normotension (n=128)	Hypertension (n=128)	P value
TAR (%)	117 (53.2)	254 (51.6)	0.701	63 (49.2)	72 (56.3)	0.260
CABG/MVR/MVP/TVP (%)	26 (11.8)	42 (8.5)	0.169	17 (13.3)	13 (10.2)	0.437
Aortic valve (%)	108 (49.1)	108 (22.0)	<0.001	58 (45.3)	29 (22.7)	<0.001
CPB time (min)	245.3±66.5	239.5±68.9	0.296	238.0±62.7	243.7±79.0	0.522
Posterior parallel circulation time (min)	51.4±28.4	58.1±34.4	0.008	48.5±21.1	54.2±25.1	0.055
Aortic cross-clamp time (min)	176.0±56.7	165.1±58.6	0.021	171.4±55.7	171.2±66.6	0.976
DHCA time (min)	26.9±13.4	29.0±12.4	0.046	26.9±13.4	28.7±12.6	0.286

Data presented as n (%); mean ± standard deviation. PSM, propensity score matching; TAR, total arch replacement; CABG, coronary artery bypass graft; MVR, mitral valve replacement; MVP, mitral valvuloplasty; TVP, tricuspid valvuloplasty; CPB, cardiopulmonary bypass; DHCA, hypothermic circulatory arrest.

Table 3 Comparison of postoperative variables before and after matching.

Characteristics	Overall cohort			PSM cohort		
	Normotension (n=220)	Hypertension (n=492)	P value	Normotension (n=128)	Hypertension (n=128)	P value
Drainage volume 24 hours after surgery (mL)	808.6±833.9	721.8±607.0	0.182	828.1±935.0	696.9±547.2	0.184
Ventilation time (hour)	33.5±49.3	35.5±53.5	0.704	36.4±56.7	33.1±45.0	0.656
30-day mortality (%)	19 (8.6)	71 (14.4)	0.032	14 (10.9)	12 (9.4)	0.697
ICU stay time (day)	6.1±5.4	7.3±12.8	0.212	5.7±5.2	5.6±4.6	0.770
Hospital stay time (day)	21.6±11.1	22.7±13.2	0.301	21.3±13.2	23.9±13.7	0.128

Data presented as n (%); mean ± standard deviation. PSM, propensity score matching; ICU, intensive care unit.

Table 4 Causes of late mortality (n=41)

Cause of late mortality	Overall cohort	
	Normotension (n=9)	Hypertension (n=32)
Aortic rupture	2	7
Cardiac failure	2	4
Neurological	1	3
Respiratory failure		1
Malignancy	1	1
Suicide	1	
Unknown	2	16

in *Table 4*. It was important to point out that 1 normotensive patient committed suicide six months after discharge and was excluded from the study. 48 (7.8%) patients were lost to follow-up after hospital discharge, and these patients were excluded from the subsequent long-term survival analysis. The preoperative, operative and postoperative characteristics of the 49 patients mentioned above were comparable with the rest of the population (569 patients) (*Table 5*). Comparison between patients who survived (529 patients) with who did not (40 patients) suggested that the incidence of hypertension in patients who deceased during follow-up was higher and the survived patients were more likely to take regular anti-hypertensive medication (*Table 6*).

Table 5 Comparison of variables between lost to follow-up and the rest of patients

Variables	Available (n=569)	Lost to follow-up (n=49)	P value
Preoperative variables			
Demographics			
Age (year)	51.5±13.1	51.1±11.0	0.763
Male (%)	420 (73.8)	37 (75.5)	0.795
BMI (kg/m ²)	25.2±5.2	26.5±4.1	0.219
Medical history			
Hypertension (%)	383 (67.3)	36 (73.5)	0.376
Diabetes mellitus (%)	11 (1.9)	0 (0)	1.000
Previous cardiac surgery (%)	31 (5.4)	0 (0)	0.162
Previous coronary artery disease (%)	18 (3.2)	0 (0)	0.386
Cerebrovascular disease (%)	19 (3.3)	0 (0)	0.388
LVEF (%)	55.0±6.4	56.2±8.3	0.555
Pericardial effusion (%)	19 (3.3)	1 (2.0)	1.000
BP at admission			
SBP (mmHg)	137.0±23.6	138.6±22.0	0.647
DBP (mmHg)	82.4±13.3	86.4±13.6	0.053
Mean BP (mmHg)	100.6±15.7	103.8±15.0	0.166
Preoperative laboratory data			
WBC (10 ⁹ /L)	10.9±4.1	10.6±3.7	0.589
sCr (μmol/L)	102.1±84.9	95.0±53.2	0.563
PLT (10 ⁹ /L)	146.6±65.1	145.1±65.5	0.880
Fibrinogen (g/L)	2.6±1.3	2.7±1.3	0.835
Triglyceride (mmol/L)	1.3±1.0	1.4±1.6	0.312
Preoperative CTA			
Left renal artery: true lumen (%)	313 (63.7)	24 (63.2)	0.966
Right renal artery: true lumen (%)	371 (75.6)	29 (76.3)	1.000
Operative variables			
TAR (%)	298 (52.4)	27 (55.1)	0.714
CABG/MVR/MVP/TVP (%)	44 (7.7)	8 (16.3)	0.055
Aortic valve (%)	176 (30.9)	15 (30.6)	0.963
CPB time (min)	236.0±245.9	245.9±60.5	0.299
Posterior parallel circulation time (min)	54.1±29.2	61.3±38.0	0.113
Aortic cross-clamp time (min)	165.6±55.4	170.7±52.2	0.537
DHCA time (min)	28.1±12.9	27.3±13.8	0.664

Table 5 (continued)

Table 5 (continued)

Variables	Available (n=569)	Lost to follow-up (n=49)	P value
Postoperative variables			
Drainage volume 24 hours after surgery (mL)	700.7±601.0	725.0±629.3	0.792
Ventilation time (hour)	34.3±49.2	27.2±24.6	0.421
ICU stay time (day)	6.9±11.9	6.1±4.9	0.633
Hospital stay time (day)	23.7±11.4	24.5±14.6	0.654

Data presented as n (%); mean ± standard deviation. BMI, body mass index; LVEF, left ventricular ejection fraction; BP, blood pressure; SBP, systolic blood pressure; DBP, diastolic blood pressure; WBC, white blood cell; PLT, platelet; SCr, serum creatinine; CTA, computed tomography angiography; TAR, total arch replacement; CABG, coronary artery bypass graft; MVR, mitral valve replacement; MVP, mitral valvuloplasty; TVP, tricuspid valvuloplasty; CPB, cardiopulmonary bypass; DHCA, hypothermic circulatory arrest; ICU, intensive care unit.

Table 6 Comparison between patients who survived with who did not in the follow-up period

Characteristics	Follow-up survivals (n=529)	Follow-up death (n=40)	P value
Hypertension (%)	350 (66.2)	33 (82.5)	0.034
Regular treatment of hypertension (%)	182 (34.4)	5 (12.5)	0.004
Antihypertensive medications			
Beta-blocker (%)	132 (25.0)	5 (12.5)	0.076
Calcium channel blocker (%)	102 (19.3)	4 (10.0)	0.146
Diuretic (%)	58 (11.0)	1 (2.5)	0.108
ACEI or ARB (%)	74 (14.0)	3 (7.5)	0.247
Others (%)	50 (9.5)	7 (17.5)	0.105
Uncontrolled hypertension (%)	157 (29.7)	15 (37.5)	0.299

Data presented as n (%). ACEI, angiotensin-converting enzyme inhibitors; ARB, angiotensin receptor blocker.

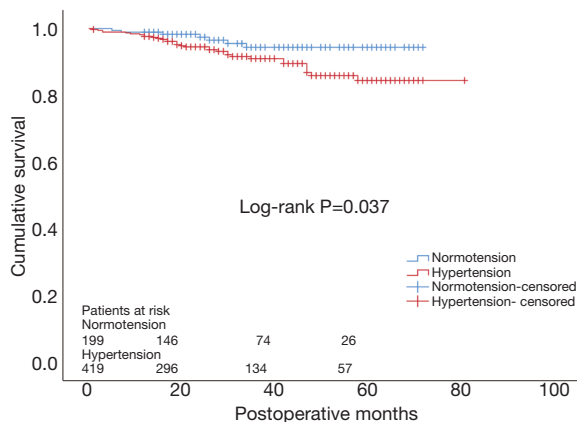


Figure 1 Kaplan-Meier estimate for late survival of patients operated on for type A acute aortic dissection according to the presence of hypertension.

Furthermore, a significant difference of long-term survival rate was discovered by Kaplan-Meier test between hypertensive and normotensive two groups ($P=0.037$) (Figure 1).

Following univariate analysis, parameters including age above 65 years ($P=0.050$), female gender ($P=0.095$), hypertension ($P=0.044$), overweight (BMI greater than 24) (7) ($P=0.096$), duration of hypertension ($P=0.004$), diastolic BP ($P=0.020$) and mean BP ($P=0.045$) at admission, postoperative regular treatment of hypertension ($P=0.015$), postoperative usage of beta-blockers ($P=0.073$) were selected for further multivariate analysis. Multivariable Cox regression analysis showed that hypertension [hazard ratio (HR), 3.040; 95% confidence interval (CI), 1.124–8.227; $P=0.029$], postoperative regular treatment of hypertension (HR, 0.227; 95% CI, 0.087–0.589; $P=0.002$) and usage of beta-blockers after discharge (HR, 0.328; 95% CI, 0.126–

0.853; $P=0.022$) were proved to be independent predictors for long-term survival (Table 7).

Discussion

It has been known that hypertension is one of the most important risk factors for aortic dissection (4). However, the predictive value of hypertension in patients of TA-AAD underwent cardiac surgeries remained understudied. Therefore, we examined the association between preexisting hypertension and mortality rates in our hospital. In this study, we demonstrated that complicating with hypertension did not increase patients' mortality rate within 30 days after surgery. However, the long-term survival rate of hypertensive patients was significantly lower compared to normotensive patients. In addition, hypertension was identified as an independent predictor for increased long-term mortality rate.

Compared to previous studies that trying to decipher the influence of blood pressure on patients with TA-AAD (8,9), one of the strengths of this study was that we applied PSM methods on the cohort before further analysis. As a result, confounders that might bias the results were excluded between two groups. The patient cohort analyzed in our study comprised of consecutively enrolled patients who underwent surgical repair of TA-AAD. In our cohort, 492 patients (69.1%) were complicated with hypertension, a percentage that is consistent with previous studies (5,10-13). Our clinical profile depictions suggested that hypertensive patients were older and heavier than normotensive patients, similar to previous reports (8,9,14). Unsurprisingly, hypertensive patients were associated with significantly elevated preoperative sCr, which was also consistent with other reports (9,15). Elevated preoperative leucocyte levels were also identified in hypertensive patients, which was consistent with prior studies that demonstrated an association between leucocyte count and hypertension incidence (16,17). This finding suggested that inflammation may play a role in the development of hypertension. Most studies started their survival analysis on hospital day 1 thereby included in-hospital deaths. It is important to notice that we only count deaths that happened after hospital discharge in this study which might give us an underestimated mortality rate when comparing to other studies.

Another important finding in our study was that hypertensive patients received less aortic valve management. However, normotensive patients tend to be younger in our study and we could not rule out the influence of age on

disease manifestations and procedure selections. As a matter of fact, reports from IRAD investigators suggested that younger patients more often underwent root surgeries (18).

In the present study, we found that hypertension was not associated with increasing peri-operative mortality rate. Several reports analyzing the influence of preoperative factors on the survival of patients with TA-AAD found a similar result that hypertension was not an independent predictor of higher operative mortality (8,19-21). One reason might be that the patients' BP was closely monitored and proper anti-hypertensive medications were prescribed during hospitalization period. Another reason might be that the area of greatest aortic shear stress was surgically repaired and the risk of developing a new separation of the aortic vessel layers was minimized.

Our findings suggested hypertension was associated with long-term survival in TA-AAD patients who received surgeries. We also found that patients who had never been treated or taking antihypertensive medication irregularly after surgery had an increased long-term mortality compared to whose hypertension was properly controlled. Uncontrolled BP could lead to cardiac failure and aortic rupture, which were the main reasons for late death as well as other studies (21). Unfortunately, it is difficult to achieve in some patients. From a nationwide database of 276,197 subjects (aged 40-75 years), there were 123,063 hypertensive patients, and only 77,379 (63%) received proper anti-hypertensive drugs. Another research found that only 598 out of 848 (70.5%) hypertensive patients received regular medication in China (9). A previous report demonstrated that an increase in BP of 26 mm Hg was equivalent to an increase of the aortic diameter of 1 centimeter (22). An 25 years follow-up study of 252 patients who underwent repair of TA-AAD suggested that the reoperation probability was markedly decreased with improved systolic BP control (11). Tsai and associates examined 303 consecutive patients with TA-AAD concluded that patients who died within 3 years of surgery had increased systolic BP compared with those who survived (130 vs. 122, $P<0.01$) (13). Consistent with previous studies, our data suggested that strict medication adherence and BP control after discharge is pivotal in the management of patients experienced aortic dissection surgery repair.

Contrary to our conclusions, Merkle and colleagues (8) demonstrated that hypertensive patients were not associated with worse long-term survival compared to normotensive patients. However, compared to their study, we involved more cases and more sophisticated statistical analysis like PSM and Cox regression analysis to eliminate potential bias

Table 7 Univariate and multivariate analysis on factors for late mortality in hospital survivors

Characteristics	Univariate analysis			Multivariate analysis		
	HR	95% CI	P value	HR	95% CI	P value
Age(≥65y)	2.051	1.001–4.203	0.050		N/A	0.460
Female gender	1.727	0.910–3.275	0.095		N/A	0.084
Hypertension	2.317	1.025–5.238	0.044	3.040	1.124–8.227	0.029
BMI ≥24	1.738	0.906–3.333	0.096		N/A	0.156
Duration of hypertension	1.057	1.018–1.098	0.004	1.042	0.994–1.092	0.087
Regular treatment of hypertension	0.314	0.123–0.801	0.015	0.227	0.087–0.589	0.002
DBP at admission	1.028	1.004–1.052	0.020		N/A	0.278
Mean BP at admission	1.021	1.000–1.042	0.045		N/A	0.428
Beta blocker	0.424	0.166–1.082	0.073	0.328	0.126–0.853	0.022
Diuretic	0.244	0.034–1.777	0.164		N/A	
SBP at admission	1.010	0.996–1.023	0.160		N/A	
Uncontrolled hypertension	1.047	0.741–2.670	0.296		N/A	
Calcium channel blocker	0.503	0.179–1.413	0.192		N/A	
ACEI or ARB	0.573	0.177–1.859	0.354		N/A	
Other medication	1.786	0.790–4.037	0.164		N/A	
Previous coronary artery disease	1.514	0.365–6.238	0.567		N/A	
Cerebrovascular disease	2.537	0.782–8.231	0.121		N/A	
sCr	1.001	0.998–1.003	0.559		N/A	
WBC	0.944	0.868–1.027	0.182		N/A	
PLT	0.998	0.993–1.003	0.458		N/A	
Triglyceride	0.806	0.527–1.232	0.319		N/A	
LVEF	0.987	0.905–1.077	0.768		N/A	
Pericardial effusion	2.523	0.777–8.189	0.123		N/A	
TAR	0.719	0.351–1.472	0.367		N/A	
CPB time	1.000	0.995–1.004	0.923		N/A	
Aortic valve	1.009	0.520–1.955	0.980		N/A	
DHCA time	0.980	0.957–1.004	0.105		N/A	
Drainage volume 24 hours after surgery	1.000	0.999–1.001	0.901		N/A	
Ventilation time	1.000	0.992–1.007	0.971		N/A	
ICU stay time	1.003	0.986–1.019	0.761		N/A	
Hospital stay time	1.000	0.973–1.028	0.994		N/A	

HR, hazard ratio; CI, confidence interval; BMI, body mass index; DBP, diastolic blood pressure; SBP, systolic blood pressure; ACEI, angiotensin-converting enzyme inhibitors; ARB, angiotensin receptor blocker; TAR, total arch replacement; SCr, serum creatinine; WBC, white blood cell; PLT, platelet; LVEF, left ventricular ejection fraction; CPB, cardiopulmonary bypass; DHCA, hypothermic circulatory arrest; ICU, intensive care unit; N/A, not available.

in baseline characteristics, which might render us a better strength. And it is important to point out that ethnics might also play an important role. In addition, the observation in a cohort consisted of 232 consecutive patients treated surgically for TA-AAD followed up to 4 years supported our findings (21). Numerous studies, including us, proved that in order to prevent late adverse events, frequent follow-up and appropriate medication are essential for patients who have undergone aortic repair for TA-AAD.

Regular beta-blockers treatment after hospital discharge was associated with decreased long-term mortality in our study. Current evidence regarding the beneficial effect of beta-blockers may be numerous, while there is little data after aortic dissection repair. In a prospective cohort of 3,102 patients, Chan *et al.* reported that patients regularly taking beta-blockers after cardiac surgery exhibit a substantially lower mortality rate during long-term follow-up (23). Thus, in patients with hypertension, strong consideration should be given to the prescription of beta-blockers after aortic dissection repair surgery.

Study limitations

Our research presented with several limitations. Firstly, our study was a retrospective study conducted in a single center, with a cohort that might not necessarily representable for the general population. In addition, our study had a median follow-up period of 29 months. A longer follow-up period might provide additional insights into the impact of hypertension on disease's long-term outcome. Another points that worth mentioning are that the survival analysis excluded patients lost to follow-up and PSM method was not applied to the long-term follow-up analysis. Moreover, during the follow-up, the exact cause of death in some patients could not be verified, which also might affect our analysis.

Conclusions

Hypertensive patients who underwent surgeries for TA-AAD presented with similar perioperative outcomes but increased long-term mortality rate compared to normotensive patients. Hypertension was identified as an independent risk factor for late mortality. Our study provided insights in re-evaluating the importance of hypertension in TA-AAD patients. Future studies included more patients with prolonged follow-up period are needed to verify our findings.

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Footnote

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Ethical Statement: The authors are accountable for all aspects of the work in ensuring that questions related to the accuracy or integrity of any part of the work are appropriately investigated and resolved. The ethics committee of Nanjing Drum Tower hospital approved this retrospective study (No. BL2014004) and waived the need for individual informed consent due to the nature of this study. The study was conducted in accordance with the Declaration of Helsinki (as revised in 2013).

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