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Systematic total arch replacement with thoraflex hybrid graft in acute type A aortic dissection: A single centre experience

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Introduction: In the last two decades, a more aggressive approach has been encouraged to treat patients with acute type A aortic dissection (ATAAD), extending the repair to the aortic arch and proximal descending thoracic aorta with the frozen elephant trunk (FET) implantation. Here, we report our single-centre experience with the FET technique for the systematic treatment of emergency type A aortic dissection.

Materials and methods: Between December 2017 and January 2022, 69 consecutive patients were admitted with ATAAD; of those, 66 patients (62.9 \pm 10.2 years of age, 81.8% men) underwent emergency hybrid aortic arch and FET repair with the multibranched Thoraflex hybrid graft and were enrolled in the study. Primary endpoints were 30 days- and in-hospital mortality. Secondary endpoints were postoperative morbidity and follow-up survival. To better clarify the impact of age on surgical outcomes, we have divided the study population into two groups: group A for patients <70 years of age (47 patients), and group B for patients \geq 70 years (19 patients). Time-to-event analysis has been conducted using the Log-rank test and is displayed with Kaplan-Meier curves. A multiple Cox proportional Hazard model was developed to identify predictors of long-term survival with a stepwise backward/forward selection process.

Results: 30-days- and in-hospital mortality were 10.6 and 13.6%, respectively. Stroke occurred in three (4.5%) patients. Two (3.0%) patients experienced spinal cord ischemia. We did not find any statistically significant difference between the two groups in terms of main post-operative outcomes. The multivariable Cox proportional hazard model showed left ventricular ejection fraction (HR: 0.83, 95% CI: 0.79–0.92, p < 0.01), peripheral vascular disease (HR: 15.8, 95% CI: 3.9–62.9, p < 0.01), coronary malperfusion (HR: 0.10, 95% CI: 0.01–0.77,

p = 0.03), lower limbs malperfusion (HR: 5.1, 95% CI: 1.10–23.4, p = 0.04), and cardiopulmonary bypass time (HR: 1.02, 95% CI: 1–1.04, p = 0.01) as independent predictors of long term mortality.

Conclusions: Frozen elephant trunk repair to treat emergency type A aortic dissection appears to be associated with good early and mid-term clinical outcomes even in the elderly.

KEYWORDS

aortic arch surgery, FET, frozen elephant trunk, hybrid arch surgery, acute type A aortic dissection

Introduction

Acute type A aortic dissection (ATAAD) is one of the most dangerous and fatal cardiovascular emergencies. Aortic rupture and subsequent cardiac tamponade as well as systemic malperfusion are the most common cause of death as a result of this potentially deadly condition. In most of the cases, emergency surgery is the only therapeutic choice (1, 2). Although substantial improvements in surgical repair techniques and post-operative management have enhanced outcomes for this subset of patients, the largest registries still show high in-hospital mortality of 15–20% (3–6). Traditionally, a less aggressive approach limited to ascending aorta with or without hemiarch replacement has been adopted to reduce the surgical risk (7, 8). However, pathologic remodeling with dilation of the residual dissected aorta and subsequent risk of rupture requiring further intervention in the future may occur. Nowadays, the frozen elephant trunk (FET) technique has become a valuable alternative to treat aortic disease when the arch and the thoracic aorta are involved, both in elective and emergency settings (9-12). Particularly, in acute aortic dissection, the use of FET can lead to expansion and stabilization of the true lumen and can cover eventual additional tears in the stented part of the aortic arch or proximal descending thoracic aorta (DTA) (13, 14). Despite these potential advantages, only a restricted number of institutions have employed this procedure to treat aortic dissection (15-33).

The aim of this study is to review our systematic experience with arch reconstruction and the FET technique for patients presenting with acute type A dissection. A secondary aim is to evaluate the impact of age on the postoperative outcome after this type of surgery.

Materials and methods

Study population and definitions

This is a single-center, retrospective, observational study based on prospectively collected data obtained from institutional cardiac surgery dataset at the University Hospital San Giovanni di Dio and Ruggi d'Aragona, Salerno, Italy. The study was conducted in accordance with the principles of the Declaration of Helsinki. Institutional board approval was obtained for the study, and patient consent was waived. Between December 2017 and January 2022, 69 consecutive patients presented at our unit with the diagnosis of ATAAD. The Glasgow Coma Scale was used to assess the level of consciousness, and patients were classified into the following 3 subgroups based on their Glasgow Coma Scale scores: severe, 3-8; moderate, 9-12; and mild, 13-15 (34). We did not perform AAAD repair in patients whose severe coma persisted for more than 10 h. There was only 1 non-operative management patient in this study period. Two patients were treated with a more conservative approach (ascending aorta and hemiarch replacement) due to extremely poor general conditions at presentation. All the other 66 patients underwent repair with the FET technique using Thoraflex hybrid (Terumo Aortic, Scotland) prosthesis and were enrolled in the present study. Fifty-four (81.8%) were male and the mean age was 62.8 (± 10.1) years. All the surviving patients were followed-up until May 2022. Patient demographics are summarized in Table 1. To better clarify the impact of age on surgical outcomes, we have divided the study population into two groups: group A for patients <70 years of age (47 patients) and group B for patients \geq 70 years (19 patients, Table 1). Emergency surgery was defined as surgery conducted within 24 h of

Abbreviations: AKI, acute kidney injury; ATAAD, acute type A aortic dissection; AVR, aortic valve replacement; CABG, coronary artery bypass grafting; CI, confidence interval; CKD, chronic kidney disease; COPD, chronic obstructive pulmonary disease; CPB, cardiopulmonary bypass; CVA, cerebro-vascular event; CVVH, continuous veno-venous haemofiltration; DSINEs, distal stent graft induced new entries; DTA, descending thoracic aorta; eGFR, estimated glomerular filtration rate; FET, frozen elephant trunk; HCA, hypothermic circulatory arrest; LCA, left common carotid artery; LOS, length of stay; LSA, left subclavian artery; SD, standard deviation; TEVAR, thoracic endovascular aortic repair.

TABLE 1 Preoperative characteristics.

Characteristics	Overall (66)	Group A (47)	Group B (19)	<i>p</i> -value
Age (years)	62.9 (10.2)	58 (7.9)	75 (3)	< 0.001
Male gender	54 (81.8)	41 (87.2)	13 (68.4)	0.089
COPD	13 (19.7)	11 (23.4)	2 (10.5)	0.3
History of hypertension	57 (86.4)	41 (87.2)	16 (84.2)	0.7
CKD	6 (9.1)	6 (12.7)	0	na
History of cancer	3 (4.5)	2 (4.2)	1 (5.3)	>0.9
Diabetes mellitus	7 (10.6)	4 (8.5)	3 (15.7)	0.4
Peripheral vascular disease	8 (12.1)	4 (8.5)	4 (21.0)	0.2
Previous cardiac surgery	3 (4.5)	1 (2.1)	2 (10.5)	0.2
Previous cerebrovascular accident	1 (1.5)	1 (2.1)	0	na
Left ventricle ejection fraction (perc.)	57 (6.3)	57.4 (7.3)	59.1 (4.2)	0.28
Bicuspid aortic valve	1 (1.5)	1 (2.1)	0	na
Marfan syndrome	1 (1.5)	1 (2.1)	0	na
Coronary malperfusion at presentation	10 (15.2)	6 (12.8)	4 (21.0)	0.5
Preoperative hemoglobin level (g/dl)	12.7 (1.7)	13.1 (1.6)	11.9 (1.8)	0.017
eGFR	68.1 (27.5)	67.3 (27.7)	70.1 (27.8)	0.14
Cerebral malperfusion at presentation	8 (12.1)	4 (8.5)	4 (21.0)	0.2
Abdominal malperfusion at presentation	11 (16.7)	9 (19.1)	2 (10.5)	0.5
Lower limb ischaemia at presentation	7 (10.6)	6 (12.8)	1 (5.3)	0.7
Penn classification				0.8
Aa	32 (48.5)	23 (48.9)	9 (47.3)	
Ab	21 (31.8)	16 (34.0)	5 (26.3)	
Ac	6 (9.1)	4 (8.5)	2 (10.5)	
Abc	7 (10.6)	4 (8.5)	3 (15.8)	
Preoperative lactate level	1.68 (1.48)	1.62 (1.33)	1.84 (1.8)	0.7

Group A: age at surgery <70 years; group B: age at surgery ≥ 70 years; COPD, Chronic Obstructive Pulmonary Disease; CKD, Chronic Kidney Disease; eGFR: estimated glomerular filtration rate. Data are reported as mean (SD) for numerical variables and as count (%) for categorical variables.

unscheduled admission. Patients were considered to have a chronic obstructive pulmonary disease (COPD) if they had any of the following conditions: long-term use of bronchodilators or steroids for lung disease before admission; outpatient visits including a diagnosis of COPD; and preoperative lung function test with evidence of an obstructive pattern, Patients were considered to have diabetes if they had any of the following conditions: receipt of insulin or oral hypoglycemic medications before admission; outpatient visits including a diagnosis of diabetes mellitus on two occasions; or a previous inpatient stay with a discharge diagnosis of diabetes mellitus. Cardiogenic shock was defined as a preoperative systolic blood pressure <90 mmHg or a cardiac index <2.0 l/min/m2 at arrival. Malperfusion syndromes were defined as symptoms due to disrupted blood flow to the coronary arteries, central nervous system, or peripheral arteries. Any patient with neurological symptoms or syncope that was apparently caused by cardiogenic shock was excluded from the classification of cerebral malperfusion (35).

Thoraflex hybrid FET graft and size selection

The Thoraflex hybrid is a vascular graft designed for complex aortic arch surgery. It consists of a proximal not-stented tubular gelatin-coated Dacron graft and a distal polyestermade stent-graft with a self-expandable nitinol stent, which is deployed anterogradely into the aortic arch/DTA during circulatory arrest. The hybrid prosthesis has four integrated lateral branches: three on the dorsal side for the single reimplantation of supra-aortic vessels and one on the ventral side for systemic perfusion. Between the two portions, there is a sewing collar that makes the distal anastomosis of the prosthesis to the aortic arch wall easier. The Thoraflex hybrid comes in different sizes: the proximal part diameter varies from 22 to 32 mm, and the stented part from 24 to 40 mm. Two different distal lengths are available: 100 and 150 mm. The combination of the varied sizes and lengths allows us to tailorize the graft to the patient's anatomy and pathologic condition. In our series, the decision on stented graft size was figured out by the total aortic diameter and relative diameters of the true and false lumen at the level of the landing zone, based on the exact evaluation of preoperative CT angiogram of the aorta for the entire cohort of patients. No oversizing was performed to reduce the risk of rupture or distal stent graft-induced new entries (DSINEs). To minimize the risk of spinal cord ischemia, we only implanted the 100 mm length stented graft.

Surgical technique

Our "debranching first" FET technique (Figure 1) has already been described in earlier reports (36, 37). The usual incision was normally extended in a small bilateral supraclavicular cervicotomy to improve access to the epiaortic vessels. The right subclavian artery (isolated from anterior mediastinum) or the right axillary artery (isolated in the right sub clavicular region) were routinely employed as the arterial site for central cannulation. In all cases, an end-to-side Dacron vascular graft (8 or 10 mm, depending on the native vessel size) was interposed to avoid direct cannulation of the artery. The right atrium and right superior pulmonary vein were cannulated for venous return and left ventricle venting. A homemade 4-branched perfusion circuit was used for extracorporeal circulation. Cooling to 26, 28°C for hypothermic circulatory arrest was employed in all cases. During the first cooling phase, from a beating heart, the left common carotid artery (LCCA) and the left subclavian artery (LSA) were isolated and prepared for selective cannulation with the interposition of a vascular Dacron graft. The vessel perfusion was sequentially started to achieve complete antegrade "trilateral" cerebral perfusion before clamping the ascending aorta. After debranching completion, at 30°C core temperature, the ascending aorta was cross-clamped and opened and cardioplegia was administered. A single dose of Custodiol^(R) cardioplegia was routinely administered selectively in the coronary ostia. The proximal aortic valve and root reconstruction were performed in different manners as required by the case as shown in Table 2. At a temperature of 26–28°C, the brachiocephalic artery was clamped and the selective antegrade perfusion began at 10-15 ml/kg/min, to stop the systemic circulation. The aortic arch was then opened and inspected. The landing zone (usually zone 2) was reinforced with internal and external teflon strips. At this stage, the distal stent graft of the Thoraflex hybrid was released into the DTA. The strengthened collar of the prosthesis was sutured to the aorta, and, after cannulation of the fourth lateral branch and accurate de-airing, systemic perfusion was resumed, starting to rewarm the body. In the last 19 cases, to reduce lower body ischemia time, a blood flow of around 1,200 ml for spinal and splanchnic perfusion was started through the lateral branch of the Thoraflex hybrid even before completing the collar's anastomosis. The anastomosis between the hybrid prosthesis and sino-tubular junction (either native or prosthetic, depending on the proximal repair) was then completed and the cross-clamp was released. The prosthesiselongated supra-aortic vessels were then termino-terminally reanastomosed to the corresponding branches of the Thoraflex hybrid, starting with the LSA to LCCA and finally at the brachiocephalic artery.

Endpoints

The primary endpoints of the study were 30-days and inhospital mortality, defined as death due to any cause during the postoperative course at 30 days and until discharge, respectively.

Secondary endpoints included postoperative stroke (defined as clinical and radiological evidence of a new postoperative cerebrovascular event -CVA-), spinal cord injury, return to the operating room for cardiac causes, renal failure requiring replacement therapy, respiratory insufficiency requiring prolonged ventilation and/or tracheostomy, deep sternal wound infection involving sternal bone and/or mediastinal structures, recurrent laryngeal nerve palsy, and in-hospital length of stay.

Statistical analysis

Data are presented as mean and standard deviation (SD) for continuous numerical variables and as count and percentages for categorical variables. Numerical variables have been compared using the Student t-test or Mann-Whitney U test, while categorical variables have been compared with the chi-square test or Fisher exact test as appropriate. Time-to-event analysis has been conducted using the Log-rank test and displayed with Kaplan-Meier curves. A multiple Cox proportional Hazard model was developed to identify predictors of long-term survival with a stepwise backward/forward selection process. The alpha error was set at 0.05 for significance and all tests are two-sided. Missing values were screened before analysis: every variable with more than 5% of the missing value was eliminated from the final dataset. The remaining variables were imputed using simple imputation models. The statistical analysis was conducted with R version 3.6.0 (2019-04-26) (38).

Results

The mean follow-up was 19.7 ± 17.4 months. The distributions of baseline characteristics for the overall population are presented in Table 1. Three (4.5%) patients had already undergone cardiac surgery. Thirty-six patients (54.5%) were presented with peripheral malperfusion. Particularly, 10 (15.2%) patients were presented with coronary malperfusion, 8 (12.1%) patients with cerebral malperfusion, 11 (16.7%) with abdominal malperfusion, and 7 (10.6%) patients with peripheral



on the proximal repair) is then completed (I) and the cross-clamp is released. The prosthesis-elongated supra-aortic vessels are then termino-terminally re-anastomosed to the corresponding branches of the Thoraflex hybrid (L), starting with the LSA to LCCA and finally to the

brachiocephalic artery (M). Finally, the anonymous vein is reconstructed with the interposition of an 8 mm Dacron vascular graft (O).

limb ischemia. According to the Penn classification (39) for the overall study cohort, 32 (48.5 %) patients were in class Aa, 21 (31.8%) were in class Ab, 6 (9%) patients were in class Ac, and 7 (10.6%) patients were in class Abc. There were no significant differences in terms of significant preoperative comorbidities and clinical presentation between the groups, A (<70 yrs old) and B (≥ 0 yrs old). The Penn classification did not differ between the two groups with 49 vs. 47% patients in class Aa, 34 vs. 26% in class Ab, 8.5 vs. 11% in class Ac, and 8.5 vs. 16% in class Abc (group A vs. group B respectively, p = 0.8). However, patients in group A were more frequently male (87 vs. 68%, p = 0.089) and had higher preoperative hemoglobin

levels (13.1 \pm 1.6 vs. 11.9 \pm 1.8, p = 0.017). No deaths were recorded during the procedure. Operative characteristics and their distributions among the two groups are shown in Table 2. Concomitant procedures were needed in 32 (48.5%) patients: 11 (16.7%) coronary artery bypass grafting, 4 (6.0%) aortic valve replacement, 17 (25.7%) aortic root surgery, such as 12 (18.2%) Florida sleeve procedures and 5 (7.6%) modified Bentall. There was no difference between the two groups in terms of cardiopulmonary bypass time (205.1 \pm 43.5 min in group A vs. 203 \pm 50.3 min in group B, P = 0.7) and aortic cross-clamp time (118 \pm 35.8 min in group A vs. 108 \pm 36.9 min in group B, P = 0.2) However, the mean moderate hypothermic

	Overall (66)	Group A (47)	Group B (19)	<i>p</i> -value
CPB time (min)	204.5 (45.2)	205.1 (43.5)	203 (50.3)	0.7
Aortic cross-clamp time (min)	115.1 (36.1)	118 (35.8)	108.1 (36.9)	0.2
HCA time (min)	28.7 (6.9)	30 (6.1)	25.5 (7.9)	0.046
Intraoperative lactate level (peak)	5.7 (2.8)	5.4 (2.2)	6.5 (3.8)	0.6
Concomitant procedures				
CABG	11 (16.6)	6 (12.8)	5 (26.3)	0.3
AV replacement	5 (7.6)	4 (8.5)	1(5.3)	0.4
Aortic root surgery				
Florida Sleeve	12 (18.2)	9 (19.1)	3 (15.8)	0.9
Modified Bentall	5 (7.6)	3 (6.4)	2 (10.5)	0.6

TABLE 2 Operative characteristics.

Group A: age at surgery <70 years; group B: age at surgery ≥ 70 years; CPB, Cardiopulmonary Bypass; HCA, Hypothermic Circulatory Arrest (including selective antegrade cerebral and visceral perfusion); CABG, Coronary Artery Bypass Graft; AV, Aortic Valve. Data are reported as mean (SD) for numerical variables and as count (%) for categorical variables.

TABLE 3 Post-operative outcomes.

	Overall (66)	Group A (47)	Group B (19)	<i>p</i> -value
30-days mortality	7 (10.6)	3 (6.4)	4 (21.0)	0.1
In-hospital mortality	9 (13.6)	4 (8.5)	5 (26.3)	0.11
ITU LOS (days)	15 (16.4)	15.3(16.9)	14.4 (15.4)	>0.9
In-hospital LOS (days)	27.7 (21.7)	27.5 (20.7)	28.3 (24.6)	0.78
Return to operating room	6 (9.1)	5 (10.6)	1 (5.3)	0.7
Prolonged ventilation	26 (39.3)	15 (31.9)	11 (57.9)	0.05
Respiratory failure	39 (59.1)	28 (59.6)	11 (57.9)	0.9
Tracheostomy	19 (28.8)	12 (25.5)	7 (36.8)	0.4
Pericardial effusion requiring drainage	15 (22.7)	11 (23.4)	4 (21.0)	>0.9
Pleural effusion requiring drainage	20 (30.3)	14 (29.8)	6 (31.6)	0.8
Deep sternal wound infection	7 (10.6)	4 (8.5)	3 (15.7)	0.4
Recurrent laryngeal nerve palsy	6 (9.1)	5 (10.6)	1 (5.3)	>0.9
AKI requiring CVVH	20 (30.3)	16 (34.0)	4 (21.0)	0.3
Spinal cord injury/paraplegia	2 (3.0)	1 (2.1)	1 (5.3)	0.069
Permanent CVA	3 (4.5)	1 (2.1)	2 (10.5)	0.2
Lower limb ischaemia	2 (3.0)	1 (2.1)	1 (5.3)	0.5

Group A: age at surgery <70 years; group B: age at surgery ≥ 70 years; ITU, Intensive Therapy Unit; LOS, Length of stay; CO, Cardiac Output; AKI, Acute Kidney Injury; CVVH, Continuous Veno-Venous Haemofiltration; CVA, Cerebrovascular Accident. Data are reported as mean (SD) for numerical variables and as count (%) for categorical variables.

circulatory arrest time with a selective antegrade trivascular cerebral perfusion was significantly higher in group A (30 \pm 6.1 min in group A vs. 25.5 \pm 7.9 min in group B, P = 0.046). Postoperative outcomes are summarized in Table 3. 30 days mortality was 10.6% for the entire cohort of patients (6.4% in group A vs 21% in group B, P = 0.1). Cumulative in-hospital mortality was 13.6% (8.5% in group A vs. 26% in group B, P = 0.11). No differences were found in terms of postoperative CVA (2.1% in group A vs. 10.5% in group B, P = 0.2), spinal cord injury (2.1% in group A vs. 5.3% in group B, P = 0.069) and AKI requiring hemodialysis (34% in group A vs. 21% in group B, P = 0.3). Also, the incidence of reopening for bleeding (10.6 vs. 5.3%, P=0.7), deep sternal wound infection (8.5 vs.

15.8%), respiratory failure requiring tracheostomy (25.5 vs 36.8%, P=0.4), and laryngeal nerve palsy (10.6 vs. 5.3%) did not differ among the two groups.

Overall survival for the entire cohort at 3, 6, 12, and 24 months was 85, 80.3, 76.5, and 74.4% respectively (Figure 2). Survival rates by group were 91 vs. 73.7% at 3 months, 85.6 vs. 68% at 6 months, 79.9 vs. 68% at 12 months, and 77 vs. 68% at 24 months, all group A vs. group B, respectively (Figure 3).

The multivariable Cox proportional hazard model (Table 4) showed left ventricular ejection fraction (HR: 0.83, 95% CI: 0.79– 0.92, p < 0.01), peripheral vascular disease (HR: 15.8, 95% CI: 3.9–62.9, p < 0.01), coronary malperfusion (HR: 0.10, 95% CI: 0.01-0.77, p =0.03), lower limbs malperfusion (HR: 5.1, 95% CI:



1.10–23.4, p = 0.04), and cardiopulmonary bypass time (HR: 1.02, 95% CI: 1 – 1.04, p = 0.01) as independent predictors of long term mortality.

All survived patients underwent clinical and imaging followup until May 2022. According to our protocol, a postoperative CT angiogram was planned at 3, and 12 months after surgery and once yearly for the following 2 years. During this period, 11 patients required further aortic interventions: 10 patients (eight in group A and two in group B) underwent endovascular extension of the stent graft (TEVAR) using Relay Plus (Terumo Aortic, Scotland) covered endograft due to partial false lumen thrombosis/negative aortic remodeling as defined by Shesthra et al. (12). One patient developed a pseudoaneurysm at the level of proximal anastomosis which was successfully treated by positioning a vascular plug percutaneously under fluoroscopy via the right femoral artery. Freedom from aortic reinterventions for the entire cohort at 6, 12, and 24 months was 97, 92.2, and 80.6%, respectively (Figure 4); if considered by the group, freedom from aortic re-intervention rate was 97.9 vs. 100% at 6 months, 88.8 vs. 100% at 12 months and 78.1 vs. 88.2% at 24 months, all group A vs. group B, respectively (Figure 5).

Discussion

There is an ongoing discussion about the most appropriate management of the aortic arch in the context of ATAAD, and particularly whether the FET technique should be consistently and systematically employed to treat these patients regardless of the primary involvement of the arch itself in the pathologic process. Our series proves that systematic use of the FET technique in an emergency setting to treat patients with ATAAD is feasible with good short-term and mid-term outcomes. In our study, 30-days and in-hospital mortality, CVA, and spinal cord injury were overall satisfactory and in line with the range reported in a contemporary review of literature recently published by our group where the FET technique was used as a surgical procedure in such complex setting (15). These results are even superior if compared to the data reported by recent



reports of international registries where a more conservative approach with ascending aorta and hemiarch replacement was employed (3-5, 9).

Our experience with FET using Thoraflex hybrid in an emergency setting has been overall satisfactory; technical success was achieved in all patients. We did not pay attention to the physiological learning curve of this systematic approach in an acute setting with bad surgical results even in the first part of our experience. Correct cerebral perfusion during the hypothermic circulatory arrest is one of the main aspects determining neurological outcomes. As already reported, we believe that our peculiar "debranching first" strategy, as well as the trilateral selective antegrade cerebral perfusion during a moderate hypothermic circulatory arrest, is important to explain our particularly satisfactory results in terms of neurological complications, with a low overall incidence of permanent stroke (4.5%) compared to data from similar studies (36). This peculiar strategy allows uniform cerebral perfusion throughout the operation, except for the brief time needed for LCCA and LSA anastomosis, thus minimizing the cerebral ischemia time.

In our opinion, the claimed technical difficulties of this technique should not stop surgeons from adopting this surgical strategy in an acute setting, especially if a specialized centre with a high volume of aortic surgery is identified on a regional basis to receive and treat acute aortic syndrome (40–42). From a technical point of view, we believe that the FET hybrid prostheses, such as Thoraflex hybrid help to reduce the risk of aortic arch surgery in this context, by avoiding the need of performing a challenging distal anastomosis deeply at the level of the proximal DTA, where the risk of bleeding or rupture mainly due to the fragile dissected aortic wall is extremely high. Several authors, adopting systematically this strategy in an emergency setting, have stressed the importance of a hybrid prosthesis to face this challenging scenario (43).

Moreover, many patients affected by ATAAD present with distal aorta malperfusion due to the true lumen compression by the false lumen; in these cases, the FET technique can potentially TABLE 4 Multiple Cox PH model for long-term survival.

Left ventricular ejection fraction 0.85 0.78–0.92	< 0.01
Peripheral vascular disease 15.8 3.9–62.9	< 0.01
Coronary malperfusion 0.10 0.01–0.77	0.03
Lower limb malperfusion 5.1 1.10–23.4	0.04
Cardiopulmonary bypass time 1.02 1.00-1.05	0.01



re-expand the compressed true lumen and cover supplementary entry tears at the level of the proximal DTA, which may well sustain pressurization of the false lumen, thus reverting the malperfusion and improving surgical outcomes (44).

It is well-known that aortic dissection is a progressive disease and the natural history of the distal false lumen of the aorta following a primary procedure of ascending aorta or hemiarch replacement may lead to degeneration, aneurysm, and/or rupture, thus requiring additional extensive interventions. Data from large registries and recent literature suggest that in 60 to 90% of cases, a negative remodeling of the distal aorta takes place, to the extent of requiring second-stage endovascular or open surgical completion. In acute dissection, the FET technique can reduce DTA dilatation by bringing both coverages of secondary entry tears located in the proximal DTA and obliteration of the false lumen at the proximal DTA, thus reducing aortic-related deaths and the requirement for challenging distal aortic reinterventions (45–49). In our series, only 11 patients have required further aortic interventions: 10 patients underwent endovascular extension of the stent graft (TEVAR) using Relay Plus (Terumo Aortic, Scotland) covered endograft due to partial false lumen thrombosis, and 1 patient developed a pseudoaneurysm at the level of proximal anastomosis which was successfully treated by positioning a vascular plug percutaneously under fluoroscopy *via* the right femoral artery. This aspect supports the idea that the rate of aortic reintervention is lower if compared to conventional conservative treatment. Also, the rate of success for those patients was 100%, mainly due to less complex endovascular procedures when a hybrid



vascular prosthesis has been implanted during the first surgical treatment.

In a recent paper, Beckmann et al. (50) have reported the results of a series of patients treated for non-urgent total aortic arch replacement using the FET technique. In their study worse perioperative mortality and morbidity as well as long-term survival was proved in septuagenarians than in younger patients. For this reason, to better clarify the impact of age on our cohort of patients, we have divided our study population into two groups: group A <70 years old and group B \geq 70 years old. Our results showed no significant differences both for primary and secondary outcomes, thus supporting the thesis that in an acute setting, the age above 70 years old on its own does not impact negatively on perioperative outcomes and demonstrating that the use of FET in this setting is safe and effective even in the elderly population.

Although deeper evidence on long-term benefits is still needed, we believe that patients presenting with ATAAD should be at least considered for surgical treatment using the FET technique. Surely, the condition of the patient must be taken into account in deciding which approach should be adopted to treat this condition. In fact, even if the anatomy of the dissection is suitable, total arch replacement may be not indicated, such as in significantly elderly patients or in particularly poor clinical conditions with signs and symptoms of systemic malperfusion (18). In this regard, our subanalysis based on Penn classification showed that patients in class Ac and Abc, presenting with poor haemodynamic conditions, had worse outcomes if compared to patients in class Aa and/or Ab; particularly, 30 days and inhospital mortality were significantly higher in class Ac and Abc (see Supplementary Table E1).

Study limitations

The main limitation of this study is related to the retrospective and single-center nature of the study.

Moreover, the small number of patients included in this study represents a limitation in terms of statistical power and possible generalization of the results.

Conclusion

Hybrid aortic arch and FET repair with the Thoraflex Hybrid graft to systematically treat emergency type A aortic dissection appears to be associated with good early- and mid-term clinical outcomes even in the elderly. Our perfusional strategy with a single pump and head vessel reimplantation as the last step has proved to offer highly effective cerebral protection while optimizing the heart and body ischemic time. Moreover, by inducing distal false lumen obliteration and thrombosis, the FET technique is likely to reduce the need for secondary procedures. Further studies to better clarify late complications are still needed.

Author's note

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Data availability statement

The raw data supporting the conclusions of this article will be made available by the authors, without undue reservation.

Ethics statement

The studies involving human participants were reviewed and approved by University Hospital San Giovanni di Dio e Ruggi d'Aragona. Written informed consent for participation was not required for this study in accordance with the national legislation and the institutional requirements.

Author contributions

Conception and design: PC, GM, VB, PM, and SI. Administrative support: EM, VB, RC, MM, EM, and MD. Provision of study materials or patients: PC, RL, MM, MC, and FC. Collection and assembly of data: PC, GM, MM, MC, MD,

References

1. El-Hamamsy I, Ouzounian M, Demers P, McClure S, Hassan A, Dagenais F, et al. Canadian thoracic aortic collaborative (CTAC). State-of-the-art surgical management of acute type A aortic dissection. *Can J Cardiol.* (2016) 32:100–9. doi: 10.1016/j.cjca.2015.07.736

2. Czerny M, Schoenhoff F, Etz C, Englberger L, Khaladj N, Zierer A, et al. The impact of pre-operative malperfusion on outcome in acute type a aortic dissection: results from the GERAADA registry. *J Am Coll Cardiol.* (2015) 65:2628–35. doi: 10.1016/j.jacc.2015.04.030

3. Berretta P, Patel HJ, Gleason TG, Sundt TM, Myrmel T, Desai N, et al. IRAD experience on surgical type A acute dissection patients: results and predictors of mortality. *Ann Cardiothorac Surg.* (2016) 5:346–51. doi: 10.21037/acs.2016.05.10

4. Matsushita A, Tabata M, Fukui T, Sato Y, Matsuyama S, Shimokawa T, et al. Outcomes of contemporary emergency open surgery for type A acute aortic dissection in elderly patients. *J Thorac Cardiovasc Surg.* (2014) 147:290–4. doi: 10.1016/j.jtcvs.2012.11

5. Pape LA, Awais M, Woznicki EM, Suzuki T, Trimarchi S, Evangelista A, et al. Presentation, diagnosis, and outcomes of acute aortic dissection: 17-year trends from the international registry of acute aortic dissection. *J Am Coll Cardiol.* (2015) 66:350–8. doi: 10.1016/j.jacc.2015.05.029

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Conflict of interest

The authors declare that the research was conducted in the absence of any commercial or financial relationships that could be construed as a potential conflict of interest.

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Supplementary material

The Supplementary Material for this article can be found online at: https://www.frontiersin.org/articles/10.3389/ fcvm.2022.997961/full#supplementary-material

6. Peterss S, Pichlmaier M, Curtis A, Luehr M, Born F, Hagl C. Patient management in aortic arch surgery†. *Eur J Cardiothorac Surg.* (2017) 51:i4-i14. doi: 10.1093/ejcts/ezw337

7. Westaby S, Saito S, Katsumata T. Acute type A dissection: conservative methods provide consistently low mortality. *Ann Thorac Surg.* (2002) 73:707–13. doi: 10.1016/S0003-4975(01)03449-X

8. Kazui T, Yamashita K, Washiyama N, Terada H, Bashar AH, Suzuki T, Ohkura K. Impact of anaggressive surgical approach on surgical outcome in type A aortic dissection. *Ann Thorac Surg.* (2002) 74:S1844–7. doi: 10.1016/s0003-4975(02)04155-3

9. Di Bartolomeo R, Pantaleo A, Berretta P, Murana G, Castrovinci S, Cefarelli M, et al. Frozen elephant trunk surgery in acute aortic dissection. *J Thorac Cardiovasc Surg.* (2015) 149:S105–9. doi: 10.1016/j.jtcvs.2014.07.098

10. Shrestha M, Fleissner F, Ius F, Koigeldiyev N, Kaufeld T, Beckmann E, Martens A, Haverich A. Total aortic arch replacement with frozen elephant trunk in acute type A aortic dissections: are we pushing the limits too far? *Eur J Cardiothorac Surg.* (2015) 47:361–6. doi: 10.1093/ejcts/ezu185

11. Watanuki H, Ogino H, Minatoya K, Matsuda H, Sasaki H, Ando M, et al. Is emergency total arch replacement with a modified elephant trunk technique

justified for acute type A aortic dissection? Ann Thorac Surg. (2007) 84:1585-91. doi: 10.1016/j.athoracsur.2007.06.045

12. Shrestha M, Bachet J, Bavaria J, Carrel TP, De Paulis R, Di Bartolomeo R, et al. Current status and recommendations for use of the frozen elephant trunk technique: a position paper by the Vascular Domain of EACTS. *Eur J Cardiothorac Surg.* (2015) 47:759–69. doi: 10.1093/ejcts/ezv085

13. Rice RD, Sandhu HK, Leake SS, Afifi RO, et al. Is total arch replacement associated with worse outcomes during repair of acute type a aortic dissection? *Ann Thorac Surg.* (2015) 100:2159–65. doi: 10.1016/j.athoracsur.2015.06.007

14. Di Eusanio M, Berretta P, Cefarelli M, Jacopo A, Murana G, Castrovinci S, et al. Total arch replacement versus more conservative management in type A acute aortic dissection. *Ann Thorac Surg.* (2015) 100:88–94. doi: 10.1016/j.athoracsur.2015.02.041

15. Chivasso P, Mastrogiovanni G, Miele M, Bruno VD, Rosciano A, Montella AP, et al. Frozen elephant trunk technique in acute type A aortic dissection: is it for all? *Medicina*. (2021) 57:894. doi: 10.3390/medicina57090894

16. Harky A, Fok M, Bashir M. Which is the optimal frozen elephant trunk? A systematic review and meta-analysis of outcomes in 2161 patients undergoing thoracic aortic aneurysm surgery using E-vita open plus hybrid stent graft vs. thoraflex hybrid prosthesis. *Braz J Cardiovasc Surg.* (2020) 35:427–36. doi: 10.21470/1678-9741-2019-0220

17. Smith HN, Boodhwani M, Ouzounian M, Saczkowski R, Gregory AJ, Herget EJ, Appoo JJ. Classification and outcomes of extended arch repair for acute Type A aortic dissection: a systematic review and meta-analysis. *Interact Cardiovasc Thorac Surg.* (2017) 24:450–9. doi: 10.1093/icvts/ivw355

18. Berger T, Czerny M. The frozen elephant trunk technique in acute and chronic aortic dissection: intraoperative setting and patient selection are key to success. *Ann Cardiothorac Surg.* (2020) 9:230–2. doi: 10.21037/acs-2019-fet-10

19. Liu K, Zhu C, Zheng X, Wang T, Xu R, Zhu Z, et al. A new aortic arch inclusion technique with frozen elephant trunk for type A aortic dissection. *Ann Surg.* (2020) 271:978–83. doi: 10.1097/SLA.00000000003122

20. Kaneyuki D, Mogi K, Watanabe H, Otsu M, Sakurai M, Takahara Y. The frozen elephant trunk technique for acute retrograde type A aortic dissection: preliminary results. *Interact Cardiovasc Thorac Surg.* (2020) 31:813–9. doi: 10.1093/icvts/ivaa199

21. Inoue Y, Matsuda H, Omura A, Seike Y, Uehara K, Sasaki H, Kobayashi J. Comparative study of the frozen elephant trunk and classical elephant trunk techniques to supplement total arch replacement for acute type A aortic dissection. *Eur J Cardiothorac Surg.* (2019) 56:579–86. doi: 10.1093/ejcts/ezz104

22. Furutachi A, Takamatsu M, Nogami E, Hamada K, Yunoki J, Itoh M, Kamohara K. Early and mid-term outcomes of total arch replacement with the frozen elephant trunk technique for type A acute aortic dissection. *Interact Cardiovasc Thorac Surg.* (2019) 29:753–60. doi: 10.1093/icvts/ivz154

23. Katayama A, Uchida N, Katayama K, Arakawa M, Sueda T. The frozen elephant trunk technique for acute type a aortic dissection: results from 15 years of experience. *Eur J Cardiothorac Surg.* (2015) 47:355–60. doi: 10.1093/ejcts/ezu173

24. Xiao Z, Meng W, Zhu D, Guo Y, Zhang E. Treatment strategies for left subclavian artery during total arch replacement combined with stented elephant trunk implantation. *J Thorac Cardiovasc Surg.* (2014) 147:639–43. doi: 10.1016/j.jtcvs.2013.02.013

25. Yang SM, Xu P, Li CX, Huang Q, Gao HB Li ZF, Chang Q, et al. modified total arch replacement combined with a stented elephant trunk implantation for acute type A dissection under deep hypothermic circulatory arrest and selective antegrade cerebral perfusion. *J Cardiothorac Surg.* (2014) 9:140. doi: 10.1186/s13019-014-0140-6

26. Shi E, Gu T, Yu Y, Yu L, Wang C, Fang Q, et al. Early and midterm outcomes of hemiarch replacement combined with stented elephant trunk in the management of acute DeBakey type I aortic dissection: comparison with total arch replacement. *J Thorac Cardiovasc Surg.* (2014) 148:2125–31. doi: 10.1016/j.jtcvs.2013.10.058

27. Ma WG, Zheng J, Dong SB, Lu W, Sun K, Qi RD, et al. Sun's procedure of total arch replacement using a tetrafurcated graft with stented elephant trunk implantation: analysis of early outcome in 398 patients with acute type A aortic dissection. *Ann Cardiothorac Surg.* (2013) 2:621–8. doi: 10.3978/j.issn.2225-319X.2013.09.06

28. Hoffman A, Damberg ALM, Schälte G, Mahnken AH, Raweh A, Autschbach R. Thoracic stent graft sizing for frozen elephant trunk repair in acute type A dissection. *J Thorac Cardiovasc Surg.* (2013) 145:964–9. doi: 10.1016/j.jtcvs.2012.03.059

29. Uchida N, Katayama A, Tamura K, Sutoh M, Kuraoka M, Ishihara H. Frozen elephant trunk technique and partial remodelling for acute type a aortic dissection. *Eur J Cardiothorac Surg.* (2011) 40:1066–71. doi: 10.1016/j.ejcts.2011. 02.074

30. Chen X, Huang F, Xu M, Wang L, Jiang Y, Xiao L, et al. The stented elephant trunk procedure combined total arch replacement for Debakey I aortic dissection: operative result and follow-up. *Interact Cardiovasc Thorac Surg.* (2010) 11:594–8. doi: 10.1510/icvts.2010.238212

31. Tsagakis K, Pacini D, Di Bartolomeo R, Gorlitzer M, Weiss G, Grabenwoger M, et al. Multicenter early experience with extended aortic repair in acute aortic dissection: is simultaneous descending stent grafting justified? *J Thorac Cardiovasc Surg.* (2010) 140(6 Suppl):S116–20. doi: 10.1016/j.jtcvs.2010.07.066

32. Pochettino A, Brinkman WT, Moeller P, Szeto WY, Moser W, Cornelius K, Bowen FW, Woo YJ, Bavaria JE. Antegrade thoracic stent grafting during repair of acute DeBakey I dissection prevents development of thoracoabdominal aortic aneurysms. *Ann Thorac Surg.* (2009) 88:482–9. doi: 10.1016/j.athoracsur.2009.04.046

33. Jakob H, Tsagakis K, Tossios P, Massoudy P, Thielmann M, Buck T, et al. Combining classic surgery with descending stent grafting for acute DeBakey type I dissection. *Ann Thorac Surg.* (2008) 86:95–101. doi: 10.1016/j.athoracsur.2008.03.037

34. Teasdale G, Maas A, Lecky F, Manley G, Stocchetti N, Murray G. The Glasgow Coma Scale at 40 years: standing the test of time. *Lancet Neurol.* (2014) 13:844–54. doi: 10.1016/S1474-4422(14)70120-6

35. Furukawa T, Uchida N, Takahashi S, Yamane Y, Mochizuki S, Yamada K, et al. Management of cerebral malperfusion in surgical repair of acute type A aortic dissection. *Eur J Cardiothorac Surg.* (2017) 52:327–32. doi: 10.1093/ejcts/ezx056

36. Masiello P, Mastrogiovanni G, Presutto O, Chivasso P, Bruno VD, Colombino M, et al. Frozen elephant trunk procedure for complex aortic arch surgery: the Salerno experience with Thoraflex hybrid. *J Card Surg.* (2022) 37:107–14. doi: 10.1111/jocs.16086

37. Masiello P, Mastrogiovanni G, Chivasso P, Triggiani D, Cafarelli F, Iesu S, et al. modified frozen elephant trunk hybrid device to facilitate supra-aortic trunk anastomosis. *J Card Surg.* (2021) 36:371–3. doi: 10.1111/jocs.15208

38. R Core Team. *R: A Language and Environment for Statistical Computing*. Vienna, Austria: R Foundation for Statistical Computing (2019). Available online at: https://www.Rproject.org/

39. Augoustides JG, Geirsson A, Szeto WY, Walsh EK, Cornelius B, Pochettino A, et al. Observational study of mortality risk stratification by ischemic presentation in patients with acute type A aortic dissection: the Penn classification. *Nat Clin Pract Cardiovasc Med.* (2009) 6:140–6. doi: 10.1038/ncpcardio1417

40. Andersen ND, Ganapathi AM, Hanna JM, Williams JB, Gaca JG, Hughes GC. Outcomes of acute type a dissection repair before and after implementation of a multidisciplinary thoracic aortic surgery program. *J Am Coll Cardiol.* (2014) 63:1796–803. doi: 10.1016/j.jacc.2013.10.085

41. Mariscalco G, Maselli D, Zanobini M, Ahmed A, Bruno VD, Benedetto U, Gherli R, Gherli T, Nicolini F. Aortic centres should represent the standard of care for acute aortic syndrome. *Eur J Prev Cardiol.* (2018) 25(1_suppl):3–14. doi: 10.1177/2047487318764963

42. Chikwe J, Cavallaro P, Itagaki S, Seigerman M, Diluozzo G, Adams DH. National outcomes in acute aortic dissection: influence of surgeon and institutional volume on operative mortality. *Ann Thorac Surg.* (2013) 95:1563–9. doi: 10.1016/j.athoracsur.2013.02.039

43. Di Eusanio M, Petridis FD, Pacini D, Di Bartolomeo R. Facilitated aortic arch repair with the frozen elephant trunk technique. *Eur J Cardiothorac Surg.* (2011) 40:1261–2. doi: 10.1016/j.ejcts.2011.02.017

44. Di Marco L, Pantaleo A, Leone A, Murana G, Di Bartolomeo R, Pacini D. The frozen elephant trunk technique: european association for cardio-thoracic surgery position and bologna experience. *Korean J Thorac Cardiovasc Surg.* (2017) 50:1–7. doi: 10.5090/kjtcs.2017.50.1.1

45. Kimura N, Itoh S, Yuri K, Adachi K, Matsumoto H, Yamaguchi A, Adachi H. Reoperation for enlargement of the distal aorta after initial surgery for acute type A aortic dissection. *J Thorac Cardiovasc Surg.* (2015) 149(2 Suppl):S91-8.e1. doi: 10.1016/j.jtcvs.2014.08.008

46. Geirsson A, Bavaria JE, Swarr D, Keane MG, Woo YJ, Szeto WY, Pochettino A. Fate of the residual distal and proximal aorta after acute type a dissection repair using a contemporary surgical reconstruction algorithm. *Ann Thorac Surg.* (2007) 84:1955–64. doi: 10.1016/j.athoracsur.2007.07.017

47. Fattouch K, Sampognaro R, Navarra E, Caruso M, Pisano C, Coppola G, et al. Long-term results after repair of type a acute aortic dissection according to false lumen patency. *Ann Thorac Surg.* (2009) 88:1244–50. doi: 10.1016/j.athoracsur.2009.06.055

48. Evangelista A, Salas A, Ribera A, Ferreira-González I, Cuellar H, Pineda V, et al. Long-term outcome of aortic dissection with patent false lumen: predictive role of entry tear size and location. *Circulation*. (2012) 125:3133–41. doi: 10.1161/CIRCULATIONAHA.111.090266

49. Tsagakis K, Dohle DS, Jánosi RA, et al. Abstract 19090: Identification and classification of descending aorta re-entry sites in type I aortic dissection. *Circulation*. (2015) 132:S3. doi: 10.1161/circ.132.suppl_3. 19090

50. Beckmann E, Martens A, Kaufeld T, Natanov R, Krueger H, Haverich A, et al. Is total aortic arch replacement with the frozen elephant trunk procedure reasonable in elderly patients? *Eur J Cardiothorac Surg.* (2021) 60:131–7. doi: 10.1093/ejcts/ezab063