Frozen elephant trunk repair of aortic aneurysms: How to reduce the incidence of endoleak and reintervention

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ABSTRACT

Background: Frozen elephant trunk (FET) enables treatment of arch and proximal descending thoracic aorta aneurysms. In treating patients with single-stage FET, the relationship of distal stent size to endoleak and reintervention has remained unexamined.

Methods: In this retrospective analysis of 63 cases in which FET was used to repair arch or thoracic aortic aneurysms between 2008 and 2019, 36 were intended as single-stage procedures. Effective sizing and sealing of distal stents were analyzed by preoperative and postoperative computed tomography angiography (CTA).

Results: During a mean of 25.8 ± 5.7 months of CTA follow-up, 10 of 36 (28%) experienced endoleak, and 3 of 36 (8%) had sac expansion. Ultimately, 5 of 13 (38%) underwent thoracic endovascular aneurysm repair. Patients without endoleak or sac expansion were more likely to have stents with >10% oversize and a >30-mm seal in healthy aorta compared with those experiencing these complications (11 of 23 vs 0 of 13; P = .0031). Conversely, 11 of 36 patients (31%) with adequately oversized and sealed stents developed fewer endoleaks compared with those without (0 of 11 vs 10 of 14; P < .0004). Patients with endoleak or sac expansion had smaller mean distal stent oversize and shorter mean sealing length compared with those without endoleak or sac expansion (2.3 ± 3.9% vs 18 ± 2.9% [P = .0023] and 1 ± 0.7 mm vs 34 ± 6 mm [P = .0005], respectively).

Conclusions: We recommend >10% distal stent oversize and >30-mm sealing length in single-stage frozen elephant trunk repair of aortic aneurysmal disease to prevent endoleak and reintervention. Increasing multidisciplinary collaboration with endovascular surgeons will improve distal stent planning. (JTCVS Techniques 2020;3:13-20)

CENTRAL MESSAGE

We recommend >10% distal stent oversize and >30-mm sealing length in single-stage frozen elephant trunk repair of aortic aneurysmal disease to prevent endoleak and reintervention.

PERPECTIVE

The frozen elephant trunk enables single-stage treatment of arch and proximal descending thoracic aorta aneurysms, but the relationships of distal stent size to endoleak and need for reintervention remain unexamined. We show that patients with adequately oversized and sealed stents developed fewer endoleaks and required fewer reinterventions compared with those with insufficient oversizing or sealing length to the distal stent.

Frozen elephant trunk (FET) repair of the arch and thoracic aorta was introduced as a single-stage solution to extensive aortic arch and proximal thoracic aortic aneurysmal disease. The device consists of a proximal Dacron surgical graft for revascularization of the aortic arch and a distal reinforced stent graft designed to seal in the descending thoracic aorta, connected by a cuff to which the distal arch anastomosis is sutured.

Outcomes from FET procedures appear to compare favorably with those from conventional elephant trunk procedures; a case series from Hannover demonstrated...
decreased in-hospital mortality for FET compared with conventional elephant trunk. However, reintervention rates to the distal aorta in FET procedures are similar to those in conventional elephant trunk procedures, which is unexpected given that one of the advantages of the FET device is to avoid further thoracic aortic procedures by creating a primary distal seal. It is also concerning that FET procedures appear to confer an increased risk of spinal cord ischemia compared with conventional surgery.

Endoleak around the distal stent graft may represent a preventable complication resulting in further intervention to prevent sac expansion and rupture; however, the incidence of endoleak is not reported in many retrospective FET case series. Other case series have reported an incidence of endoleak of 11% to 33%. However, some endoleaks in these series might have been intentional, in a planned second-stage thoracic endovascular aneurysm repair (TEVAR) to achieve a distal seal. The incidence of endoleak does not vary according to the FET device used. Rates of endovascular reintervention to the distal aorta, where reported, are between 11% and 16%. Overall, the long-term performance of distal stent grafts in FET procedures remains unknown.

In infrarenal aortic aneurysms, an appropriate stent graft oversize is known to prevent endoleak, aneurysm sac expansion, and stent graft migration. However, there are little data on the relationship between sizing of the distal stent and later development of endoleak or ongoing sac expansion in FET procedures. Indeed, the recommendations in the position paper from the vascular domain of the European Association for Cardiothoracic Surgery (EACTS) for FET procedures make no mention of appropriate distal stent sizing. In the present study, we aimed to examine the relationship between distal stent graft oversize and sealing length and the incidence of endoleak in FET procedures intended to secure a distal seal in the thoracic aorta.

METHODS

Data were collected retrospectively starting from a database of all FET procedures performed in a single center between 2008 and 2019, containing dates of procedures together with the devices and sizes used. All FET procedures were performed using either the E-Vita Open device (JOTEC, Hechingen, Germany) or the Thoraflex device (Vascutek Terumo, Inchinnan, Scotland). The E-Vita is available in distal stent diameters of 20 to 40 mm with lengths of 130 to 160 mm, and the diameter of the proximal graft to the aortic arch matches that of the distal stent. The Thoraflex device’s distal stent is available in diameters of 24 to 40 mm and lengths of 100 to 150 mm, with varying combinations of diameters of proximal graft diameter and distal stent diameter. The instructions for the Thoraflex device state that when used for aneurysmal pathology, the distal stent diameter should exceed the thoracic aortic diameter by 15% to 25% and the distal stent should seal in >40 mm of healthy aorta. The instructions for the E-Vita device do not state sizing recommendations for aneurysmal disease. The choice of device and size was left to the operating cardiac surgeon; however, during the study period, increased contact with endovascular surgeons, particularly for elective cases, might have increased collaboration in the process of device sizing.

Computed tomography angiography (CTA) imaging pre-FET, the first postoperative scan, and all follow-up scans were analyzed for endoleak. All patients underwent annual surveillance CTA. Multiplanar reconstruction in 3 dimensions was used to measure orthogonal aortic diameters and center line reconstruction software was used to measure aortic length. On pre-FET CTA, a healthy landing zone for the distal stent was identified by finding an area of nonaneurysmal diameter (<40 mm) nontortuous aorta with <2 mm mural thrombus occupying <25% of the aortic circumference. The center lumen line distance of this from the origin of the left subclavian artery was noted. On post-FET CTA, the length of aortic coverage by the distal stent and the diameter of the aorta at this level were measured on the pre-FET scan. This allowed for a comparison between the diameter of the distal stent device and the diameter of the aorta where the distal stent graft landed. The encroachment of the distal stent onto the previously identified appropriate landing zone was measured to calculate the length of the distal seal.

The location of the maximum aortic diameter was used to define the primary location of the aneurysm under treatment (although other parts of the aorta could still have been aneurysmal or ectatic). An endoleak was defined as contrast visible outside the stent graft (in the aneurysm sac or alongside the aorta). An endoleak was considered intended for second-stage TEVAR. Aneurysm sac expansion was defined as an increase in sac size on sequential scans at either the location of the original aneurysm or in the thoracic aorta distal to the stent.

This work was carried out using an existing anonymized prospective database of patients and imaging, collected as part of the organization’s normal audit procedures where no individual can be identified retrospectively. Patient consent for inclusion was taken at the time of entry to the database. Electronic clinic letters from both cardiothoracic and vascular surgery departments were used to identify those patients who underwent further surgery or a non-endoleak distal stent-related complication. Opinions from the treating cardiothoracic surgeons and documentation from multidisciplinary team meetings were used to classify FET procedures as definitive or intended for second-stage TEVAR.

Values are reported as mean ± SEM or median (range) as appropriate. Comparisons of 2 groups were done using the t test for parametric data and the Mann–Whitney U test for nonparametric data. Comparisons of proportions were performed using the χ² tests. Statistical significance was considered to have been achieved at a P value <.05.
RESULTS

Demographics

Out of 63 FET procedures performed for aneurysmal disease between 2008 and 2019, 36 were intended as a single-stage procedure, with no preoperative plan for further open or endovascular surgery to the thoracic aorta to achieve aneurysm exclusion. Most of these procedures were performed electively (26 of 36; 72%), with 7 of 36 (19%) performed urgently or on an emergency basis. In 3 cases, the urgency of the FET procedure was unknown.

FET procedures were performed for aneurysmal disease of the ascending aorta (10 of 36; 27%), innominate artery (2 of 36; 6%), aortic arch (5 of 36; 14%), left subclavian artery (11 of 36; 31%), or descending thoracic aorta (5 of 36; 14%), as defined by the location of the largest aortic diameter. In 3 cases, the location could not be characterized, owing to incomplete preoperative imaging. The median maximum aortic diameter was 65 mm (range, 40 to 90 mm).

The study cohort comprised 17 males (47%) and 19 females (53%) with a median age of 70 years (range, 33 to 83 years) (Table 1). The FET procedure was performed using the Thoraflex device in 25 cases (69%) and with the JOTEC device in 11 cases (31%).

Patients were followed up with CTA for a mean of 25.8 ± 5.7 months after the FET procedure. All patients underwent a postoperative CTA within 4 weeks of the procedure and annual surveillance CTAs thereafter. Patients were followed up in outpatient clinics for a mean of 32 ± 5.8 months postprocedure.

Endoleak and Distal Stent–Related Complications

In 9 of the 36 patients (25%), an endoleak was visualized on the first postoperative CTA, and 1 further endoleak became evident during CTA follow-up at 9 years postprocedure, making a total of 10 endoleaks (28%) postprocedure. All but 1 of the endoleaks were classified as type 1b (arising from the distal stent) based on CTA appearance, with 1 type 2 endoleak from a likely intercostal artery (which did not cause sac expansion or require further intervention). In another 3 patients (8%), expansion of the original sac or aneurysm dilatation immediately distal to the stent became apparent on follow-up CTA (at 3, 5, and 6 years postprocedure) without any evidence of endoleak before this. Other distal stent-related complications were 1 dissection distal to the stent (3%) and 1 in-stent thrombosis (3%) (Figure 1).

Patients with endoleak and those with sac expansion were considered one group, given that the etiology of these complications is likely to be related to the sizing of the distal stent. During follow-up, 5 of 13 patients (38%) with endoleak or aneurysm sac expansion underwent TEVAR to achieve an aneurysm seal, for an overall reintervention rate of 14% (5 of 36). Four TEVAR procedures were performed electively for sac expansion to >65 mm, and 1 TEVAR procedure was performed urgently (within 48 hours of presentation) due to symptomatic rapid sac expansion. There were no ruptures due to endoleak. TEVAR procedures were performed at a median of 12 months (range, 1 to 120 months) after the FET procedure. At the time of this report, 8 of the patients with endoleak or sac expansion remained in yearly CTA surveillance and were asymptomatic with sac sizes below the 65-mm threshold for consideration of treatment.

Distal Stents with Endoleak Were More Likely to Be Undersized or Undersealed

Among the 13 patients with endoleak or sac expansion, 10 (77%) had a distal stent <10% larger than the aortic diameter where the distal end of the stent ultimately landed (undersized). The remaining 3 patients (23%) had an appropriately oversized stent but a <30-mm distal seal in healthy aorta (ie, undersealed). In the 23 patients without endoleak or sac expansion, only 4 (17%) had an undersized distal stent. Patients without endoleak or sac expansion were significantly more likely to have stents that were appropriately oversized and sealed compared with those experiencing these complications (11 of 23 vs 0 of 13; \( P = .0031, \) shaded in green in Figure 2).

### TABLE 1. Demographic data of patients undergoing intended single-stage frozen elephant trunk procedure for aneurysmal disease

| Characteristic | Value |
|---------------|-------|
| Number of patients | 36 |
| Age, y, median (range) | 70 (33-83) |
| Sex, n (%) | Male 17 (47) | Female 19 (53) |
| Urgency, n (%) | Elective 26 (72) | Emergency 7 (19) | Unknown 3 (8) |
| Location of primary aneurysm, n (%) | Ascending aorta 10 (27) | Innominate artery 2 (6) | Aortic arch 5 (14) | Left subclavian artery 11 (31) | Descending thoracic aorta 5 (14) | Unknown 3 (8) |
| Maximum aneurysm diameter, mm, median (range) | 65 (40-90) |
| Pathology, n (%) | Degenerative 31 (86) | Pseudoaneurysm 2 (1) | Connective tissue 3 (1) |
| Device, n (%) | Thoraflex 25 (69) | Jotec 11 (31) |
Patients with endoleak or sac expansion had significantly smaller mean distal stent oversize compared with patients without endoleak or sac expansion (2.3 ± 3.9% vs 18 ± 2.9%; \( P = .0023 \)) (Figure 3, A). Patients with complications had a significantly shorter mean sealing length in healthy aorta than those without complications (1 ± 0.7 mm vs 34 ± 6 mm; \( P = .0005 \)) (Figure 3, B).

**Appropriate Distal Stent Graft Sizing and Seal Decreased the Risk of Endoleak**

In 11 of 36 distal stents (31%), appropriate oversize and sealing length were achieved, whereas 14 of 36 stents (39%) were undersized with a too-short distal seal. Appropriately oversized stents developed significantly fewer sizing-related complications compared with undersized and undersealed stents (0 of 11 vs 10 of 14; \( P < .0004 \)). Ultimately, 3 of the 14 patients with insufficient oversizing and sealing length (21%) underwent TEVAR during the follow-up period (Figures 4 and 5). In 7 of 14 cases with insufficient oversizing and sealing length, the diameter of the thoracic aorta where the distal stent landed was ectatic at >36 mm. In these cases, it would not have been possible to achieve >10% oversize due to the maximum available distal stent diameter of 40 mm and length of 160 mm.

**DISCUSSION**

Our data on the incidence of endoleak after FET procedures is comparable overall to reports from other small case series (11%–35%). However, many case series have a longer follow-up period than ours, suggesting that our rate of endoleak may eventually be higher than expected. These patients are all in yearly CTA surveillance. A higher
incidence of endoleak in our study could be explained by increased sensitivity for endoleak, given that we retrospectively scrutinized surveillance CTA images specifically for this finding. Post-FET CTA surveillance protocols may also differ, altering the opportunities for detection of endoleak during follow-up.

Our rate of endovascular reintervention to the thoracic aorta after FET (14%) is comparable to that in the literature, although published case series do not report whether the FET procedure was intended to create a primary seal in the thoracic aorta or the rationale for device sizing. In addition, as the follow-up period increases, future cases of sac expansion will cross the threshold for TEVAR and require future reintervention. It is difficult to compare our series to other endovascular procedures in the thoracic aorta, given that even in TEVAR procedures, the incidence of type 1b endoleak has not been quantified owing to a lack of published data.

We have clearly demonstrated that appropriate oversizing and sealing length of the distal stent results in decreased postoperative endoleak and need for reintervention. In some of these cases, preoperative selection of a larger distal stent may have prevented endoleak. In a series of 40 FET procedures, Chu and colleagues found that selection of an appropriately oversized distal stent and a 30- to 40-mm landing zone in the thoracic aorta resulted in no type 1b endoleaks (there

![Figure 3](image3.png)

**FIGURE 3.** Frozen elephant trunk distal stents without endoleak or sac expansion had longer distal seal and larger oversize. A, Patients with no distal stent-related complication (endoleak or sac expansion; “No complication”) had a mean aortic diameter oversize of 18% (above the set threshold of 10%) compared with a mean oversize of 2.2% (below the set threshold of 10%) in those with endoleak or sac expansion (“Complication”), demonstrating the importance of distal stent oversize in the prevention of complications. B, Patients without complications (“No complication”) had a mean distal seal of 34 mm (above the set threshold of 30 mm) in healthy aorta, compared with 1 mm (below the set threshold of 30 mm) in those with endoleak or sac expansion (“Complication”), demonstrating the importance of an area of apposition of the distal stent against healthy nonaneurysmal aortic wall.

![Figure 4](image4.png)

**FIGURE 4.** Patients with adequate frozen elephant trunk (FET) distal stent sealing size and length did not experience endoleak. In 11 patients with an appropriately oversized distal stent (distal stent diameter >10% of aortic diameter) and sufficient distal seal (>30 mm of stent apposition with healthy nonaneurysmal aorta), no endoleak or sac expansion occurred (shaded in green). In the 14 distal stents with neither adequate oversize (distal stent diameter <10% of aortic diameter) nor sufficiently long distal seal (<30 mm of stent apposition with healthy nonaneurysmal aorta), 7 developed endoleak (3 of which required thoracic endovascular aneurysm repair), and 3 developed sac expansion. Only 4 did not develop a complication. This demonstrates that selection of an appropriately oversized FET distal stent of sufficient length to appose healthy nonaneurysmal aorta is likely to prevent the complications of endoleak and sac expansion, potentially reducing further endovascular reintervention.
were 2 of 40 type 2 endoleaks), suggesting again that preoperative planning can reduce endoleak. Conversely, in Jakob’s case series of FET procedures, clinicians selected a $<10\%$ distal aortic diameter device oversize for the 39 patients with aneurysmal disease, which resulted in a 26\% rate of endovascular reintervention to the thoracic aorta.\textsuperscript{15}

In patients requiring TEVAR alone, device manufacturers recommend 10\% to 20\% device oversize compared with aortic diameter and 20-mm distal sealing length in healthy aorta. Tolenaar and colleagues\textsuperscript{16} found that for TEVAR, a 10\% to 20\% oversize resulted in fewer long term type 1 endoleaks compared with $<10\%$ or $>20\%$. Ranney and colleagues\textsuperscript{17} showed that TEVAR performed according to the instructions for use resulted in a 7.3\% reintervention rate over 11 years, lower than that reported for FET procedures over a longer follow-up period. Our thresholds of 10\% oversize and 30-mm distal landing zone are conservative compared with the recommended instructions for use for FET devices, but we have demonstrated that adhering to these criteria resulted in no endoleaks and no reinterventions. Indeed, Damberg and colleagues\textsuperscript{18} recommended a 10\% to 20\% oversize of the distal stent for FET procedures performed for aneurysmal disease. In cases where the thoracic aorta was aneurysmal where the distal stent landed, it might not have been possible to oversize the stent owing to the limited range of stent graft diameters. In these instances, identification of a more distal, nonaneurysmal landing zone and planned extension into this with second-stage TEVAR would have prevented endoleak, with the distal stent of the FET acting as the proximal landing zone for the TEVAR. Greenberg and colleagues\textsuperscript{19} have shown that completion TEVAR in this context is feasible and can be accomplished with minimal mortality. This option needs to be considered before selection of the FET device, given that a significant size discrepancy between the proximal and distal thoracic stent graft diameters is difficult to mitigate using off-the-shelf TEVAR devices.

Moving the landing zone more distally raises concern about the increased risk of spinal cord ischemia due to more extensive coverage of intercostal arteries supplying the thoracic spinal cord,\textsuperscript{20} and in our series this might explain the reluctance to extend aortic coverage with second-stage TEVAR. None of the 36 patients in this series experienced permanent paraplegia or temporary spinal cord

Figure 5: Example of late endoleak after intended single-stage frozen elephant trunk (FET) requiring unplanned thoracic endovascular aneurysm repair (TEVAR) to extend the seal into healthy, nonaneurysmal aorta more distally. A and B, Axial and sagittal computed tomography angiogram (CTA) of distal stent 17 days after a planned single-stage FET procedure, showing small type 1b endoleak at the distal aspect of the distal stent (arrows). C and D, Axial and sagittal CTA showing increase in endoleak (arrows) and aneurysm sac expansion 2 years later. E, Post-TEVAR CTA showing extension of distal seal into healthy nonaneurysmal aorta using a further stent-graft and no endoleak. This patient’s original FET distal stent was undersized relative to the aortic diameter and had an insufficient sealing length into healthy nonaneurysmal aorta. Preoperative planning of the distal stent length and diameter might have avoided the endoleak and need for reintervention.
Although some have reported that staging these procedures is a protective factor against spinal cord ischemia, a recent meta-analysis concluded that this was not the case. The risk of causing paraplegia with TEVAR extension must be balanced against the risks of endoleak, sac expansion, and subsequent rupture.

Involvement of endovascular surgeons in the preoperative planning of these procedures might enable the identification of an appropriate landing zone in the thoracic aorta and facilitate preoperative planning of the distal stent, with or without the need for TEVAR extension. Because the majority of cases are elective, inclusion of endovascular surgeons in multidisciplinary decision making could improve stent graft planning. In emergency cases, the development of aortic emergency rotas could facilitate access to a team member with appropriate expertise in stent graft sizing and planning. The introduction of a multidisciplinary approach has already been shown to contribute to a reduction in operative mortality in the context of acute type A aortic dissection.

This study is limited by the small sample size, which precluded any identification of risk factors for the development of sac expansion or endoleak during CTA follow-up. Because of the small number of cases and events, our statistical analysis can only be interpreted as descriptive, and inference that these findings apply to larger datasets cannot be computed. The retrospective analysis of data and the short follow-up period limits comparisons with other case series. Stent graft use and planning during the long data collection period might have changed as surgeons learned from previous experience with endoleak.

CONCLUSIONS

Our results demonstrate the importance of appropriate preoperative sizing of the distal stent in FET procedures. We recommend >10% aortic diameter oversize and >30-mm sealing length in healthy aorta, with planning of distal extension with TEVAR if this cannot be achieved with the FET device alone. We encourage an increase in multidisciplinary team collaboration involving endovascular surgeons at an early stage, along with the introduction of all-hours access to an endovascular specialist to facilitate optimal FET planning (Figure 6).

Conflict of Interest Statement
The authors reported no conflicts of interest.
The Journal policy requires editors and reviewers to disclose conflicts of interest and to decline handling or reviewing manuscripts for which they may have a conflict of interest. The editors and reviewers of this article have no conflicts of interest.

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Key Words: computed tomography angiography, descending thoracic aorta, frozen elephant trunk, multidisciplinary team, thoracic endovascular aneurysm repair.