Importance of dynamic aortic evaluation in planning TEVAR

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Dynamic aortic evaluation in planning thoracic endovascular aortic repair (TEVAR) is important to provide optimal stent graft sizing. Static imaging protocols do not consider normal aortic dynamics and may lead to stent graft to aorta mismatch, causing stent graft related complications, such as type I endoleak and stent graft migration. Dynamic imaging can assist in accurate stent graft selection and sizing preoperatively, and evaluate stent graft performance during follow-up. To create new imaging technologies, integration of knowledge between diverse scientific fields is essential (i.e., engineering, informatics and medicine). Different dynamic imaging modalities, such as electrocardiographic-gated computed tomography angiography (ECG-gated CTA) and four-dimensional phase-contrast MRI (4D PC-MRI), are progressively investigated and implemented into clinical practice as important instruments in preoperative planning for TEVAR. In time, further application of dynamic imaging tools for preoperative screening and follow-up after TEVAR might lead to better outcomes for patients. The advances in dynamic imaging for evaluation of the thoracic aorta using new imaging modalities and their future perspectives are addressed in this manuscript.

Keywords: Dynamic imaging; thoracic endovascular aortic repair (TEVAR); type B aortic dissection; thoracic aortic disease; computational fluid dynamics (CFD)



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Introduction

The biomechanics of the thoracic aorta play an essential role in the endovascular treatment of the descending thoracic aorta and its side branches. The pulsatility of the aorta and its side branches during the cardiac cycle, the heartbeat-dependent out-of-plane movement of the aorta, and other morphologic factors might all be of influence on the endovascular treatment for thoracic aortic pathology (1). When patients affected by thoracic aortic diseases are selected for thoracic endovascular aortic repair (TEVAR), preoperative assessment of aortic morphology is essential. In particular, the aortic arch angulation and the proximal and distal landing zones are important in order to achieve an optimal result. Adequate sizing in patients with acute type B aortic dissection (ABAD) is challenging, because the actual size of the affected true lumen is unknown and the aortic wall is expected to be more fragile than in patients

with thoracic aortic aneurysm (TAA). Moreover, a stent graft in a patient with ABAD will most likely have sealing over the entire length, while in an aneurysm the stent graft has only proximal and distal sealing and fixation zones. Therefore, moderate stent graft oversizing, less radial force of the stent graft and the absence of anchoring pins should be considered when planning TEVAR for ABAD.

Computed tomography angiography (CTA) is most commonly used to analyze the aorta and the surrounding structures, while magnetic resonance angiography (MRA) is usually adopted as second choice. Generally, using CTA, static images are acquired, and these images have been obtained at any arbitrary point during the cardiac cycle (during diastole, during systole or in between). Dynamic electrocardiographic-gated (ECG-gated) CTA, MRA and ultrasonography imaging have demonstrated significant changes in aortic dimensions during the cardiac cycle

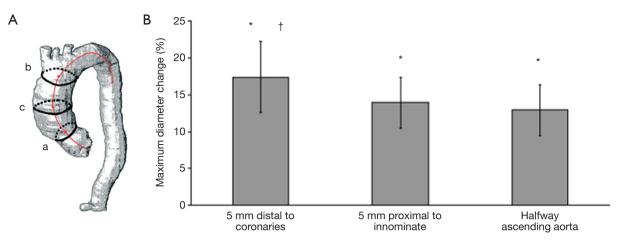


Figure 1 (A) Diagram showing the 3 measured aortic levels with the central lumen line. Level a: 5 mm distal to the coronary arteries, b: 5 mm proximal to the innominate artery, and c: halfway up the ascending aorta; (B) The mean percentage of maximum diameter change is shown at each of the 3 measured levels. Maximum diameter change at all levels is significant (*P<0.001). Level a differs significantly from b and c (+P=0.02). Adapted from van Prehn *et al.* with permission of the Journal of Endovascular Therapy (6).

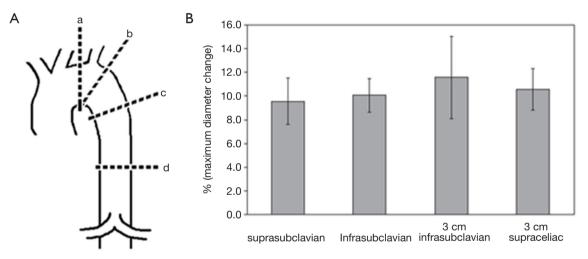


Figure 2 (A) Diagram showing the four measured thoracic aortic levels. These levels included 1 cm proximal to the left subclavian artery, 1 cm distal to the left subclavian artery, 3 cm distal to the left subclavian artery, and 3 cm proximal to the celiac artery; (B) The maximum percentage diameter change is shown at each of the four measured levels. Mean maximum diameter pulsatility is approximately 10% at all four levels (P<0.05). There is no difference in pulsatility between any of the measured levels. Adapted from Muhs *et al.* with permission of Elsevier (5).

(2,3). Irrespective of the imaging modalities, static imaging protocols do not take normal aortic dynamics into account and may subsequently lead to incorrect measurements. Guidelines state that a stent graft should be 10-20% oversized in comparison to the aortic diameter (4). However, a maximum aortic difference in pulsatility has been reported around 18% in the ascending and descending thoracic aorta (5,6) (*Figures 1,2*), and therefore the use of static imaging for TEVAR planning can lead to relative undersizing of the

stent graft to the aortic diameter. This potential mismatch might be the reason for common stent graft related complications, such as migration and endoleak type I (1,3). Distensibility of the thoracic aorta is maintained after TEVAR, which makes adequate sizing even more crucial (7). The integration of dynamic imaging into clinical practice, as a standard approach, can lead to better stent graft sizing and subsequently an improved outcome (8). Additionally, it can provide insight into the causative mechanisms for

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stent graft related complications and improve stent graft performance.

Advances in medical imaging

Over the past two decades, vascular medical imaging has progressed extensively, with the introduction of new imaging modalities considering aortic hemodynamics. ECG-gated CT and ECG-gated MRI both permit studying the pulsatility of the aorta and surrounding tissue during the cardiac cycle, and allow insight into aortic morphologic changes.

ECG-gated CT and Dual-energy CT

ECG-gated CT allows studying the pulsatility of the aorta and stent graft, by obtaining images at different time points during the cardiac cycle. ECG synchronization with a 16 to 256 slices CT scanner is mandatory for such scans, with more slices resulting in lower temporal solution, and congruently in less radiation exposure and contrast administration (9). A minimum temporal resolution of 8 phases per cardiac cycle can offer an adequate perspective of the aortic dynamics. Moreover, all major vendors have introduced iterative reconstruction algorithms for CT, which are noise-reducing methods that aim to either enhance image quality using a constant CT radiation dose, or to reduce the CT radiation dose with similar image quality (10). Dual-energy CT (DECT) imaging is a relatively new technique using two different X-ray tubes in a single CT unit, which discriminates materials based on diverse interactions with photons between the different X-ray energies. In the near future, this technique may be further implemented into clinical practice for the evaluation of aortic conditions. In particular, DECT can be used to remove bones from the datasets and virtual images allow differentiation between the contrast agent, and calcifying thrombus in the thoracic aorta of endovascularlytreated patients (11).

ECG-gated MRI

ECG-gated MRI, similar to ECG-gated CT, enables study of the pulsatility of the aorta and stent graft, by obtaining a sequence of images throughout the cardiac cycle and combining the MR images with an ECG. These images contain three spatial dimensions and one temporal solution, which is the time between the subsequent images. Compared to CTA, MR imaging is considered safer, as no radiation exposure is necessary for this high-resolution imaging modality. However, the processing time is relatively long and it is vital that patients do not move during the scan for optimal results. ECG-gated MRI is frequently used for cardiac examination and can also be used to assess the aortic motion in the preoperative setting for optimal stent graft sizing in patients with thoracic aortic pathology.

Besides stent graft sizing, dynamic MRA may also be useful for other purposes in the preoperative examination including preservation of the artery that provides the main blood supply of the anterior spinal artery (i.e., artery of Adamkiewicz), which is important to reduce the risk of spinal cord ischemia after TEVAR (12). Dynamic MRA has shown high sensitivity in detecting this artery, and subsequently the endovascular procedure can be adjusted to provide maximal spinal cord protection. Dynamic MRA is also adopted for postoperative evaluation, and, compared to conventional angiography, it has been shown to be an excellent imaging modality for the classification of endoleaks after endovascular aortic repair (13,14).

An additional dynamic application of MRI is the flow study with phase contrast sequence to assess the peak velocity, forward and reverse flow, and visualization of flow. The raw MR signal consists of two components: a magnitude and a phase. The magnitude holds anatomic information, and the phase holds velocity information, in addition to various background sources (15).

4D PC-MRI

An imaging modality with the potential to be widely implemented into clinical practice is four-dimensional phasecontrast magnetic resonance imaging (4D PC-MRI) (16,17). This imaging technique can be used for *in vivo* imaging of the blood flow within the cardiovascular system with several potential clinical applications. Volumetric flow measurements, such as cardiac output and valvular regurgitation, are already used in clinical practice, and other applications will undoubtedly follow. Coronary flow assessment to evaluate coronary artery disease, as well as aortic flow evaluation in patients with dissection and other applications, such as blood flow imaging in peripheral arteries and neurovascular systems, are of potential clinical use (18-20).

In patients with aortic dissection, 4D PC-MRI can accurately visualize and quantify flow characteristics and provide valuable information about stroke volume, velocity, dominant proximal and distal entry tears and helical flow, which seem all related to aortic expansion (*Figure 3*) (16).

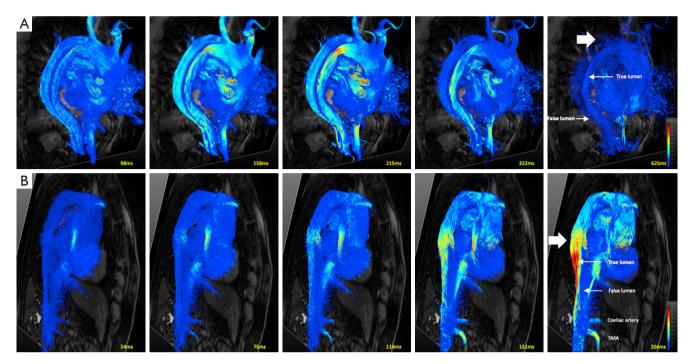


Figure 3 Visualization of entry tears. Communications between the true and false lumen (entry tears) were identified using path line analysis. These images can be viewed in 3D over time in any chosen orientation. A single plane has been selected for illustration and flow shown at five time points in the cardiac cycle. (A) One entry tear is seen at the origin of the left subclavian artery. In systole (158 and 215 ms), the velocity in the true and false lumen is approximately equal; (B) Blood flows from the true to the false lumen through an entry tear in the proximal descending thoracic aorta. An area of high velocity is seen just distal to the entry tear as the false lumen expands (due to the additional flow) and true lumen narrows. Flow is seen in the celiac and superior mesenteric arteries (SMA) distally. Figure adapted from Clough *et al.* with permission of Elsevier (16).

The velocity is measured in three directions encoded in 3D imaging over time, therefore enabling extensive analysis of physiologic hemodynamics *in vivo*. Four-dimensional PC-MRI is currently only available in some specialized centers, but has the potential to be widely adopted, because no ionizing radiation is used. With MRI scanners becoming more common in the clinical setting and continuously improving technology (i.e., accelerated acquisition, imaging hardware), the application of velocity imaging with MRI will also continue to grow. Additionally, imaging technologists and clinicians are collaborating and also developing visualization and quantitative tools to streamline the large amount of information obtained in the complex velocity fields.

Computational fluid dynamics (CFD) and TEVAR

In recent years, an increasing number of studies focused on aortic hemodynamics after TEVAR using CFD, which is a numerical technique for evaluating the hemodynamic environment of a vessel segment (21-23). Phase contrast and MRA have been extensively used in the CFD field, providing flow boundary conditions in which numerical simulations can be performed and represent a method of validating of calculated flow fields (24,25).

A patient-specific approach to both stent graft design and procedural approach is required to distinguish between the pathologic and the healthy aortic condition. To successfully accomplish this goal, integration of knowledge between different scientific fields is required (i.e., engineering, informatics and medicine). The technological development in medical imaging and parallel computing has contributed to the ability to perform complex simulations for patientspecific modeling. These simulations might be used and further implemented into clinical decision-making in the near future (26).

A patient-specific computational model can be created, using information from medical imaging, to investigate the different forces acting on the aortic wall and stent graft (21).

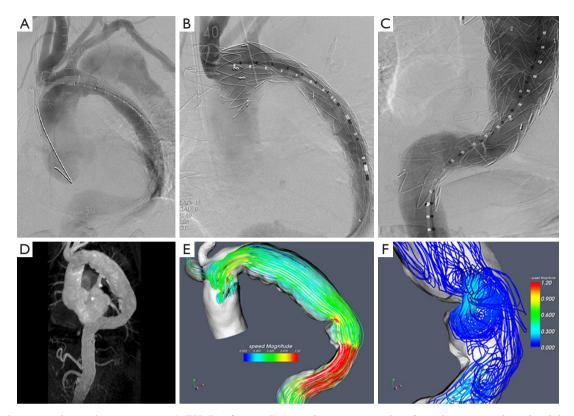


Figure 4 Thoracic endovascular aortic repair (TEVAR) of a type B aortic dissection extending from the aortic arch to the abdominal aorta. (A) Pre- and (B,C) post-implantation angiograms showing two stent grafts placed from the arch, covering the left subclavian artery, to the distal descending aorta in order to treat different entry tears and the true lumen compression; (D) MRA and (E) velocity computational fluid dynamics (CFD) study at 1 year from the implantation: the true lumen is well expanded at the proximal and mid descending aorta, while a smaller diameter is observed at the distal part of the vessel. Acceleration of blood flow is highlighted by red streamlines corresponding to >1.20 m/s velocity profiles, as indicated by the color-coded scale (speed in meters per second); (F) Patterns of turbulent flow are depicted during different phases of the cardiac cycle just above the stenotic segment. Figure adapted from Midulla *et al.* with permission of Springer (28).

In addition, CFD can help in understanding the magnitude and orientation of the loads experienced by thoracic aortic stent grafts *in vivo*, with the aim of improving stent graft design and performance (27). Furthermore, the different hemodynamic and biomechanical forces acting on the intermodular junctions of a multi-component stent graft can be investigated through computational modeling, focusing on the development of endoleak due to disconnection of the stent graft segments (22). Moreover, CFD can depict the flow conditions and the magnitude of flow by streamlines throughout the thoracic aorta (*Figure 4*) (28).

Other applications for TEVAR

TEVAR has proven to be beneficial over open surgical repair, both in TAA and ABAD, and is currently the

preferred therapy in these patients. Previous studies have shown that there is significant distension of the thoracic (descending) aorta, both pre- and post TEVAR, during the cardiac cycle in patients with TAA (7). Most interestingly, distension was preserved after stent graft placement. Based on these conditions, it has to be concluded that aortic dynamics have a profound impact on correct stent graft sizing, design and durability (7).

The primary goal of TEVAR for ABAD is to cover the primary entry tear. However, due to possible uncovered re-entry tears, communication between the true and false lumen can still be present postoperatively, inhibiting the false lumen thrombosis and aortic remodeling process. By using preoperative computational flow analyses, it is possible to detect and quantify the importance of these reentry tears/multiple entry tears, potentially assisting the physician in the choice of endovascular procedure and its extent (29).

Current status of dynamic aortic evaluation and guidelines for future research

Currently, numerous relative new dynamic imaging applications (i.e., ECG-gated CT and MRI, 4D PC-MRI, CFD analysis) have been reported in the literature. This development has expanded the multidisciplinary integrated knowledge and evaluation of the diseased thoracic aorta. However, several issues concerning the aortic arch biomechanics and their clinical consequences remain undefined and should be investigated in future research.

Larger prospective studies are required to measure flow characteristics and clinical outcomes in preoperative planning for TEVAR and in the postoperative phase. The large load of quantitative information needs to be further streamlined to be able to widely implement different new dynamic imaging tools and modalities into daily clinical practice. This will allow physicians to further determine the predictive value of the dynamic measurements. More specifically, in stratifying patients with ABAD, a new imaging modality such as 4D PC-MRI may have predictive value.

Conclusions

Technological development and interdisciplinary collaboration provide the opportunity to progressively implement dynamic imaging for aortic evaluation in planning TEVAR into clinical practice. Dynamic imaging can help preoperatively in accurate stent graft selection and sizing, and postoperatively to evaluate stent graft performance. Important imaging modalities, such as 4D PC-MRI, can give valuable information about adequate sizing and the presence and potential coverage of entry tears. Eventually, further implementation of (new) dynamic imaging tools will lead to a better outcome for the patient, which is the bottom line of research in this field.

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