Aortic stenosis and transcatheter aortic valve replacement

Degenerative valvular heart disease, including aortic stenosis (AS), is a major cause of morbidity and mortality in the aging population of industrialized countries (1,2). If left untreated, symptomatic, severe aortic stenosis is associated with poor prognosis (3,4). Therefore, surgical aortic valve replacement is indicated in these patients. However, many older patients with multiple co-morbidities and anticipated increased surgical risk are excluded from surgical intervention. For these patients, transcatheter aortic valve implantation (TAVI) is emerging as a viable treatment alternative. Transcatheter valvular heart procedures are characterized by lack of exposure and visualization of the operative field, therefore relying on image guidance, both for patient selection and preparation and the implantation procedure itself. This article describes the role of multi-detector row computed tomography (MDCT) for detailed assessment of the aortic valve, aortic root, and iliac arteries in the context of TAVI.

Key Words: Aortic stenosis; transcatheter aortic valve implantation; imaging; computed tomography

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In the decade following the first implantation in 2002, more than 30,000 TAVI procedures have been performed worldwide. Short and medium term results in patient populations with high surgical risk are encouraging (6-10). In patients not considered surgical candidates, the PARTNER trial cohort-B demonstrated a 20% absolute reduction of 1-year all-cause mortality in the TAVI cohort as compared with medical management (30.7% vs. 50.7%, respectively; P <0.001) (6). In the PARTNER trial cohort-A, which randomized high-risk patients with severe aortic stenosis to transcatheter or surgical procedures, both treatments were associated with similar rates of survival at 1 year, although there were important differences in periprocedural risks, specifically a higher risk of neurological events with TAVI (7).

The two most commonly used stent/valve systems are the balloon-expandable Edwards Sapien valve (Edwards Lifesciences, Irvine, California), and the self-expandable CoreValve ReValving system (Medtronic, Minneapolis, Minnesota). Both are approved for clinical use in the EU, but not yet in the US. Depending on the system, a retrograde transarterial technique (femoral or subclavian artery), or an ante-grade transapical implantation technique via the tip of the left ventricle are utilized. (11-13) Prior to positioning of the
Figure 1 Valvular and LV function: retrospective gated image acquisition allows 4-D image reconstruction of multiple adjacent reconstructions along the cardiac cycle displayed as cine-loops for evaluation of valvular and LV ventricular function. This figure shows images of LV analysis (left panel) and images focused on the mitral valve.

Movie 1 Retrospective gated image acquisition allows 4-D image reconstruction of multiple adjacent reconstructions along the cardiac cycle displayed as cine-loops for evaluation of valvular function. This movie shows the star-shaped opening of a tricuspid valve.

stent-valve within the aortic root, balloon aortic valvuloplasty is typically performed to prepare the landing zone. (14) The Edwards Sapien valve (Edwards Lifesciences) is then deployed and expanded by a balloon during rapid ventricular pacing to minimize cardiac output and prevent migration of the valve during deployment. The self-expanding CoreValve (Medtronic) is typically deployed without pacing.

The stent-valve is anchored within the annulus and, depending on stent design, additional stability is provided by the displaced calcified leaflets and/or the sinotubular junction. Optimal positioning of the transcatheter aortic prosthesis relative to the annulus is critical. If valve deployment is too high, increased risk of paravalvular regurgitation, aortic injury, or embolization into the aorta have been described. If positioning is too low, mitral valve dysfunction, heart block, paravalvular regurgitation, or embolization into the left ventricular cavity have been observed (15,16). The different valve systems have specific requirements regarding aortic annulus size, height and width of the aortic sinuses, as well as dimensions of the sinotubular junction. In addition, knowledge about the relationship between aortic valve leaflet height and distance between the insertion of the left/right coronary cusp and the coronary ostia is important to avoid coronary complications.

The relatively large delivery catheters currently required for insertion of the crimped valve are associated with the risk of vascular complications. It is therefore important to evaluate the size of the ilio-femoral or subclavian arteries, vessel calcification, and tortuosity. Recent developments of
lower profile delivery systems (≤18-F sheaths, outer diameter of approximately 7 mm) are expected to reduce vascular complications and expand patient eligibility for the procedure.

**Image Acquisition**

For dedicated cardiovascular imaging, CT systems with at least 64-detector technology are recommended. The spatial resolution of these systems is approximately 0.5 to 0.6 mm, which is sufficient for the anatomic evaluation of the aortic root and the iliofemoral arteries. However, the temporal resolution of current systems (about 75 ms for dual-source systems and >135 ms for single source scanners) is limited in comparison to echocardiography. CT imaging is associated with the administration of iodinated contrast and exposure to ionizing radiation exposure. Risk and benefit should be considered for individual patients (23,24).

The specific scan protocols used for TAVI assessment vary, but typically include imaging of the entire aorta including the aortic root, thoraco-abdominal aorta, and iliofemoral arteries. Depending on the desired detail of the valve/root images, this information is acquired with standard aortic protocols or separate standard aortic and high-resolution (4-D) root acquisitions. Because of the cyclic motion of the aortic valve, root, and ascending aorta, ECG-synchronized imaging of the root is critical to avoid image degradation secondary to motion artifacts. Two-options to achieve synchronization to the ECG are available, retrospective gating and prospective triggering. ECG-synchronization with retrospective gating acquires CT data throughout the entire cardiac cycle, and subsequently reconstructs images gated to a specific phase of the cardiac cycle. Reconstructions in specific phases are described in percent relative to the position in the R-R interval, e.g., a
systolic 30%-phase for valve area measurements and annular assessment corresponding to echocardiographic guidelines. Retrospective gating also allows 4-D image reconstruction of multiple adjacent reconstructions along the cardiac cycle displayed as cine-loops for evaluation of valvular and ventricular function (Figure 1, movie 1,2) (25,26).

Alternatively, ECG-synchronization can be achieved with prospective triggering, which is associated with significantly lower radiation exposure (27). With this acquisition mode, images are acquired only during a limited, pre-specified phase of the cardiac cycle (e.g. systole), with the tube turned off in-between. Therefore, reconstruction in other phases or 4-D cine-loops cannot be reconstructed. However, novel protocols with second-generation dual-source scanners allow prospective acquisition with a wider window and subsequent display of cine-loops (28).

While radiation exposure is important to consider with any CT acquisition, it is less a concern in the elderly patient populations currently considered for TAVI (24). Nonetheless, advanced scanner technology and imaging protocols, allowing lower dose imaging should be used. (29) Such advances include dual-source and volume scanners (256-320 slice), imaging with prospective triggering, high-pitch helical acquisition, and novel image reconstruction with iterative image reconstruction techniques (30,31).

In the frail patient populations currently considered for TAVI, it is critically important to weigh risk and benefit of contrast administration. Contrast is necessary for precise luminal diameter measurements of the annulus, aorta, and iliofemoral arteries. The amount of contrast depends on the specific protocol. A standard contrast-enhanced scan of the aorta uses commonly a volume of 80-120 mL. However, lower contrast volume protocols are being tested for reducing contrast-associated side effects. A scan of the pelvic arteries after intra-arterial contrast injection into the infrarenal abdominal aorta (left in place after coronary

Figure 2 Annulus Measurements: this figure demonstrates measurements at the annulus. For imaging, the annulus is defined at the level immediately below the lowest insertion point of the aortic leaflets. Measurements are taken from the phase with maximum valve opening. In this plane minimal, maximal, and mean diameter are described, with the mean diameter having the best correlation with echocardiography.
catheterization) can be performed with low-dose (15 mL) of contrast (32). If contrast administration is not feasible, a non-contrast scan still allows the assessment of overall vessel size, calcification, and tortuosity.

Pre-medication with beta-blocker or nitrates, which is standard for coronary CTA angiography, should be avoided in patients with severe AS at the time of scanning, in order to avoid haemodynamic complications. With modern scanner systems, reliable imaging of the aortic root, aorta, and iliofemoral arteries is possible even in patients with higher heart rates.

Imaging and tavi

Because of the lack of exposure and visualization of the operative field, transcatheter valvular procedures rely on image guidance for patient selection, pre-procedural planning, and intra-operative decision-making. In the context of TAVI, anatomic measurements of the aortic annulus, aortic root, aortic valve, coronary ostia, and vascular access site are of critical importance. Conventional angiography and echocardiography are essential prior and during the procedure. In addition, novel 3-dimensional imaging approaches, including computed tomography, magnetic resonance imaging, 3-D echocardiography, and C-arm CT are increasingly used (17-21).

The role and relative importance of these imaging modalities is evolving. Multidetector computed tomography (CT) has assumed an increasingly important, complementary role before and after TAVI. (22) It provides detailed anatomic assessment of the aortic root structures, the course of the descending aorta and the iliofemoral access, adding to the information obtained with echocardiography and...
angiography. Therefore routine screening with multidetector computed tomography (MDCT) is routinely performed in most large-volume centers for TAVI.

**Image reconstruction**

CT image acquisition results in (isotropic) 3-dimensional image data. The entire 3-D data can be reviewed in shaded, volume-rendered images giving depth perception (22). However, for measurement purposes the data is reconstructed along precisely defined 2D imaging planes, including those typically used in echocardiography or angiography. The strength of such 3-D-derived data analysis is the reconstruction along and perpendicular to the center axis of e.g., the aortic root or LV or the centerline of a vessel. These ‘double-oblique’ reconstructions allow precise diameter measurements, orthogonal to the axis of the structure. In particular in

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**Figure 4** Aortic Valve Area: detailed analysis of data in reconstructions along the cardiac cycle allows to identify the systolic phase with maximum valve opening (typically 20-30% RR-interval). In this reconstruction, a plane perpendicular to the valve plane is placed at the leaflet tips to measures the aortic valve opening area (AVA) by planimetry. This figure shows the star-shaped opening of a tricuspid valve.

**Figure 5** Bicuspid Aortic Valve: in contrast to Figure 4, this figure shows an example of a bicuspid valve with fusion of the non- and right coronary cusp and eccentric, slit-like opening. A systolic (left panel) and diastolic (right panel) reconstruction is shown.
non-circular structures, e.g. the oval-shaped annulus, such 3-D derived diameter measurements allow to identify minimal and maximal diameter, circumference, and area. Importantly, these diameter measurements are not limited by the acquisition plane, as 2-D derived images from angiography and echocardiography are. Although, the differences between 2-D and 3-D derived measurements appear to be small, such differences might impact device selection and outcome in a subgroup of patients. However, to date most current clinical recommendations are based on 2-D imaging with echocardiography and angiography and the incremental value of 3-D derived data is incompletely understood (17). Several groups increasingly rely on additional 3-D derived CT-measurements for device selection and sizing. A particular interesting area in imaging research is the finite element analysis of clinical imaging data, for device development and simulation of device implantation (33,34).

**Annulus**

Dedicated analysis and measurement of the annulus is important for correct selection of prosthesis size and type and to avoid damage of the annulus (15,16,35). Anatomically, the annulus is a crown-shaped 3-dimensional structure rather than a circular plane. The attachment of the aortic cusps is semi-lunar, extending throughout the aortic root from the left ventricle distally to the sinotubular junction. For clinical purposes, the annulus is defined as the plane at the lowest insertion point of the aortic valve leaflets, just above the left ventricular outflow tract (LVOT) (‘basal ring’) (Figure 2), (36,37).

Annulus measurements with 2 dimensional (2-D) transthoracic or transesophageal echocardiography, and aortic angiography provide a single diameter measurement, assuming a circular annular orifice. (38) In contrast, 3-dimensional CT reconstruction of the annulus orthogonal to the center-axis of the LVOT allows for the assessment

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**Figure 6** Aortic Valve Commisural Calcification: while CT is limited by blooming artifact from dense calcification, CT allows quantification and precise localization of calcification. The relationship between the calcification and postprocedural aortic regurgitation are unclear.
Mitral Annular Calcification: CT allows precise localization of mitral annular calcification. The impact of mitral annular calcification remains unclear of minimal and maximal diameter, circumference, and area measurements. The mean of the maximum and minimum diameter measurements is closest to 2-D measurement with echocardiography (17,36,37,39). In analogy to echocardiography, measurements are taken from systolic phase reconstructions at about 30% of the R-R interval, using the phase with maximum valve opening. (40) Recent papers describe the geometric change of the annulus during the cardiac cycle (41).

Despite the theoretical advantage of 3-D measurements as performed by CT (42), most experience and clinical recommendations regarding patient eligibility and sizing of the prosthesis is currently based upon measurements with TEE and angiography (17). Considering and comparing measurements from both TEE and MDCT may reduce the chance for error and provide a more comprehensive definition of the annular size and shape in patients considered for TAVI.

**Coronary ostia**

The pre-procedural evaluation for TAVI includes a complete assessment of coronary anatomy with conventional coronary angiography. Complete coronary assessment with CT is limited in the current population evaluated for TAVI because of the nearly universal presence of advanced calcified disease, which preclude precise assessment of luminal stenosis. However, CT allows to assess the relationship between leaflet height and distance between annulus and coronary ostia, which identifies patients at risk for coronary occlusion during the TAVI procedure. (Figure 3, Movie 3)
While no definite criteria exist to exclude patients, a distance <10 mm distance may identify increased risk (43). In these patients, peri-deployment placement of a guidewire in the left main should be considered to ensure access in case of complications.

**Aortic valve**

The aortic valve can be assessed in detail with CT. On single systolic reconstructions, and better on 4-D cineloops, the presence of a tricuspid valve can be confirmed. A tricuspid valve is characterized by symmetric leaflet height and star-shaped systolic opening versus a bicuspid valve with eccentric slit-like opening (Figures 4,5; Movies 1,2).

While CT is limited by blooming artifact from dense calcification, CT allows precise localization of calcification, e.g. extending from the commissure to the base of the anterior mitral leaflet and the mitral annulus (Figures 6,7). A non-contrast acquisition allows quantitative assessment of the aortic valve calcium score (44). Several investigators have started to investigate the potential relationship between aortic valve calcification and postprocedural aortic regurgitation and also potential impact or mitral annular calcification. However more data are needed before its impact on the TAVI procedure is fully understood (45,46).

For planimetry of the AVA, detailed analysis of data in reconstructions along the cardiac cycle allows to identify the systolic phase with maximum valve opening (typically

**Figure 8** Root Angulation: for the description of the aortic valve plane, double oblique transverse multiplanar reconstructions at the level of the aortic root are performed and rotated through a series of any angles. The right hand images show images perpendicular to the valve plane corresponding to angiographic images.
20-30% RR-interval). In this reconstruction, a plane perpendicular to the valve plane is placed at the leaflet tips to measures the aortic valve opening area (AVA) by planimetry. (42,47) Functional valvular assessment with CT is possible and demonstrates the relationship between valve calcification and leaflet motion.

**Aortic dimensions and plaque**

Measurement of the aortic sinus diameter, sinotubular junction, ascending and descending aorta are performed using centerline reconstructions. The presence of significant aneurysmal dilatation is considered a contraindication for TAVI. The extent of atherosclerotic plaque (48) is likely associated with complications including stroke (49).

**Root angulation**

Precise coaxial alignment of the stent-valve along the centerline of the aortic valve and aortic root is important during positioning. Inappropriate alignment of the device is associated with increased risk of procedural complications such as stent embolization (15,16,35). Root orientation is typically assessed using multiple repeat catheter aortograms in 1 or 2 orthogonal planes before starting the procedure or during the pre-procedural diagnostic angiogram. Typically a caudal angulation is chosen when in a right anterior oblique (RAO) projection and cranial angulation when in the left anterior oblique (LAO) projection. However, there are significant variations in patient anatomy. MDCT allows to assess the aortic root in relation to the body axis (50,51). Double oblique transverse multiplanar reconstructions are performed at the level of the root and then are rotated through a series of any angles (Figure 8).

Pre-procedural angle prediction with MDCT may decrease the number of aortograms required during the procedure, therefore shortening both procedure time and contrast usage, and potentially increases the likelihood of

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**Figure 9** Iliac Arteries: at least 1/3 of patients with critical aortic stenosis have unfavorable iliofemoral arteries. Centerline image processing of the iliac arteries (left panels) and cross-sectional images at the different levels allow assessment of dimensions, calcification, and angulation (right)
coaxial implantation by optimizing the orientation during device placement.

**Iliofemoral arteries**

Because of the relatively large diameter of the delivery sheaths (≥18F), appropriate vascular access is critical. Small luminal diameter, dense and circumferential and/or horseshoe calcification, and severe tortuosity are common in this patient population and increase the risk of access site complications (*Figure 9*) and central embolization.

Kurra *et al.* (52) reported that at least 1/3 of patients with critical aortic stenosis had unfavorable iliofemoral arteries, with the majority of those patients having minimal luminal diameters of 8 mm. In such patient alternative access approaches may include surgical sidegraft on the iliac arteries, transaxillary, or transapical access.

**Postprocedural imaging**

Postprocedural imaging relies primarily on echocardiography, which allows assessment of the valve and valvular dysfunction. CT allows to assess position of the stent-valve and its relationship to the annulus and coronary artery (47).

**Conclusion**

Because of the lack of exposure and visualization of the operative field, transcatheter valvular procedures rely on image guidance for patient selection, pre-procedural planning, and intra-operative decision-making. Anatomic measurements of the aortic annulus, aortic root, aortic valve, coronary ostia, and vascular access site are of critical importance. The role and relative importance of different imaging modalities is evolving. Multidetector computed tomography (CT) has assumed an increasingly important, complementary role before and after TAVI, and provides detailed anatomic assessment of the aortic root structures and iliofemoral access, adding to the information obtained with echocardiography and angiography. Therefore routine screening with multidetector computed tomography (MDCT) is utilized by several leading groups.

However, further evaluation of the clinical impact of MDCT and other 3-D modalities is necessary, anticipating future growth of transcatheter procedures. Similar to recent guidelines developed for the procedures itself (53), this evaluation of the role of imaging should be guided by international consensus.

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**References**


