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#### STATE-OF-THE-ART REVIEW

# Multimodality Imaging in Evaluating and Guiding Percutaneous Left Atrial Appendage Occlusion

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**Tweet:** State of the Art Review on multiple modalities that can be used for all steps in LAAO procedure #echofirst @yesCT @renujain19 @EGarciaSayan @PWesslyMD @nishath\_quader

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#### **Abstract**

Left atrial appendage occlusion (LAAO) has emerged as an important intervention for stroke prevention in patients with non-valvular atrial fibrillation (AF) who are unable to tolerate longterm anticoagulation. The development of advanced imaging technologies and techniques, such as 3D echocardiography with multiplanar reconstruction (MPR), multidetector computed tomography (MDCT), 3D intracardiac echocardiography (ICE), 3D printing, and simulation, has revolutionized pre-procedural planning, intra-procedural guidance, and post-procedural surveillance, ensuring improved precision and outcomes. Transesophageal echocardiography (TEE) remains a foundational imaging modality for assessing LAA morphology, excluding thrombi, and obtaining accurate measurements for device sizing. Recent advances in 3D TEE and MPR techniques enable enhanced visualization of complex LAA anatomies, improving device selection and procedural planning. Multidetector computed tomography has a growing role, offering high-resolution 3D reconstructions for detailed anatomical assessment. Additionally, its applications in 3D printing and virtual device simulation provide patientspecific insights, facilitating optimal device sizing and improving procedural efficiency. Intraprocedurally, 3D ICE has gained traction as a valuable alternative to TEE. With its real-time imaging capabilities and high spatial resolution, 3D ICE allows precise guidance during transseptal puncture and device deployment while reducing the need for general anesthesia. Postprocedurally, both TEE and MDCT play critical roles in assessing device stability and identifying complications such as device-related thrombus and peri-device leak. This review highlights the evolving role of multimodality imaging in LAAO, including innovations such as 3D ICE, 3D printing, and simulation. It also reviews recent literature to establish state-of-the-art

imaging practices, providing a comprehensive discussion of imaging applications across pre-, intra-, and post-procedural phases to optimize outcomes and minimize complications in LAAO.

**Keywords:** echocardiography, left atrial appendage, left atrial appendage occlusion, multimodality imaging, transesophageal echocardiography

#### **Abbreviations**

3DP = 3D printing

AF = atrial fibrillation

CT = computed tomography

**DRT** = device-related thrombus

**FDA** = US Food and Drug Administration

**HAT** = hypoattenuated thickening

ICE = intracardiac echocardiography

**LAA** = Left atrial appendage

**LAAO** = Left atrial appendage occlude/occlusion

**LUPV** = left upper pulmonary vein

**MDCT** = multidetector CT

**MPR** = multiplanar reconstruction

**PDL** = peri-device leak

**TEE** = transesophageal echocardiogram/echocardiography

#### INTRODUCTION

Atrial fibrillation (AF) affects more than 33 million people worldwide and is the second-leading cause of stroke, resulting in significant clinical morbidity. Lechocardiographic studies show that more than 90% of thrombi in non-valvular AF form in the left atrial appendage (LAA). Anticoagulation therapy significantly reduces the risk of ischemic stroke and all-cause mortality. However, more than 50% of eligible patients either discontinue or avoid anticoagulants because of bleeding concerns, side effects, or noncompliance. Alternative strategies for stroke prevention are therefore needed. The concept of LAA exclusion dates to 1949 when John L. Madden performed surgical resection to prevent recurrent arterial thrombi. Surgical LAA exclusion strategies offered a viable option for patients unable to tolerate long-term anticoagulation, but evidence of efficacy was limited.

The PROTECT AF (WATCHMAN Left Atrial Appendage System for Embolic Protection in Patients With Atrial Fibrillation)<sup>6</sup> and PREVAIL (WATCHMAN LAA Closure Device in Patients With Atrial Fibrillation Versus Long Term Warfarin Therapy)<sup>7,8</sup> trials demonstrated that percutaneous LAA occlusion (LAAO) device implantation was non-inferior to warfarin for stroke prevention in non-valvular AF. The approval of the first plug-like transcatheter LAAO device (with a single occlusive mechanism) in 2015 marked a milestone in LAAO therapy. In 2021, the US Food and Drug Administration (FDA) approved a second device, a disk-and-lobe system (a dual occlusive mechanism) that was shown in the Amulet IDE (Amplatzer Amulet LAA Occluder) trial<sup>9</sup> to have non-inferior efficacy, further expanding the available stroke prevention options (Figure 1).

LAA size, shape, and relationship with surrounding structures vary significantly between patients, and these anatomical differences impact the success of LAAO procedures.

Transesophageal echocardiography (TEE) has traditionally been used to evaluate LAA morphology, guide LAAO procedures, and monitor post-implantation outcomes. Recently, real-time 3D intracardiac echocardiography (ICE) has gained popularity for LAAO guidance, offering imaging with high spatial and temporal resolution. Multidetector cardiac computed tomography (MDCT) has emerged as a key tool in pre-procedural planning, providing detailed 3D reconstructions that allow for accurate device sizing and visualization of complex LAA anatomy.

This review aims to explore the role of multimodality imaging in pre-procedural evaluation, intra-procedural guidance, and post-procedural follow-up in percutaneous LAAO (Central Illustration). It also addresses the use of imaging in assessing device-related complications during follow-up and highlights the latest advances in and future directions of this evolving field.

#### PRE-PROCEDURAL EVALUATION

#### Transesophageal Echocardiography

Transesophageal echocardiography served as the primary imaging modality in pivotal clinical trials. <sup>7,9,10</sup> Accurate image acquisition is operator-dependent, and adherence to a comprehensive protocol is crucial for reliable measurements and conclusions. <sup>11</sup> The goals of TEE in the context of screening and planning for percutaneous LAAO procedures are to determine LAA morphology, exclude LAA thrombus, and perform measurements of the landing zone, thereby facilitating optimal device selection and sizing. A thorough TEE examination is recommended to evaluate left ventricular size and function, aortic plaque, aortic or mitral valve pathology (especially mitral stenosis), pericardial effusion, intracardiac masses, and the atrial septum.

Baseline TEE should particularly evaluate for mitral stenosis, a marker for valvular AF, to ensure

eligibility by identifying patients excluded from trials like PROTECT AF and PREVAIL. The role of percutaneous LAAO for valvular AF is currently unknown, with clinical equipoise persisting regarding its efficacy. <sup>6-8</sup> Atrial septal defects, atrial septal aneurysms, and existing atrial occluder devices may all present challenges or contraindications to the procedure.

On 2D TEE, the LAA is typically imaged at 4 omniplane angles in 45-degree intervals (0°, 45°, 90°, 135°) with and without color Doppler. Color-flow Doppler imaging can reveal areas with decreased or absent color flow within the appendage, which is highly suggestive of thrombi. Dedicated and zoomed images are recommended for improved accuracy. The shape of the LAA is determined by the number and location of appendage lobes. A "windsock" morphology is characterized by a prominent single lobe, a "cactus" morphology by 2 roughly equal lobes, a "chicken wing" morphology by a lobe that arises at an acute angle from the LAA ostium (anterior or posterior to the ostium), and a "cauliflower/broccoli" morphology by multiple lobes and variable depth (Figure 2). These morphologies influence device selection and success. Although 2D imaging can be used to determine LAA morphology, 3D echocardiography with a zoom volume acquisition of the LAA provides a more accurate depiction of the shape, number, and location of lobes. New 3D rendering algorithms, such as light-source manipulation and transillumination technologies, can enhance visualization of the LAA morphology and allow orientation of the images to resemble fluoroscopic views (Figure 2). 12,13 Both the 3D LAA multiview (LAAM) and tilt up and turn left (TUPLE) maneuvers are algorithms developed in step-wise fashion to assess LAA morphology in 3D (Figures 3 and 4).

For the plug-like LAAO device, key measurements include the landing zone, with the ostium width measured as the distance from the left circumflex coronary artery extending radially to the opposing limbus, often approximating 2 cm from the tip of the limbus (Figure

5A). The maximum width of the anticipated landing zone is determined and used for device sizing according to the manufacturer's recommendations. The measured landing zone must be between 14 mm and 36 mm to accommodate the WATCHMAN FLX/PRO device (Boston Scientific, Marlborough, Mass). The LAA depth is measured from the center of the anticipated landing zone to the LAA apex and should be more than 50% of the labeled device diameter.

For the disk-and-lobe LAAO device, the LAA orifice should be measured to ensure it is smaller than the anticipated diameter of the device disk. The anticipated landing zone of the device lobe is measured 10-12 mm distal to the LAA orifice, defined as the plane extending from the tip of the limbus to a location just proximal to the left circumflex coronary artery. The measured landing zone must be between 11 mm and 31 mm and is used for device sizing, as determined by the device manufacturer. The LAA depth is measured from the center of the LAA orifice ostium and then perpendicular to the length of the LAA orifice. Figure 5B demonstrates the 2D LAA sizing measurements for the Amplatzer Amulet device (Abbott Cardiovascular, Abbott Park, IL).

For both device styles, the manufacturers' recommendations and instructions for use suggest 2D measurements at the 4 omniplane angles. However, 2D measurements of a 3D orifice can lead to uncertainty about the exact plane of measurement and potential underestimation if not centered in the orifice. Three-dimensional echocardiography with multiplanar reconstruction (MPR) can provide precise measurements of the orifice and landing zone, ensuring all measurements are performed in the same plane. This approach also allows visualization of the deepest lobe and has been shown to lead to more accurate device sizing (Figure 6). 14,15

#### **Multidetector Cardiac Computed Tomography**

Multidetector cardiac computed tomography has a growing role in pre-procedural planning for LAAO procedures owing to its excellent spatial resolution and ability to generate precise, high-quality 3D measurements. It can replicate the key aspects of TEE in evaluating LAA morphology, maximum ostium diameter, and depth for device selection and eligibility (Central Illustration). Increasingly, MDCT is recognized as a primary imaging modality for LAAO planning, often reducing the need for pre-procedural TEE. In many centers, patients may undergo TEE only at the time of device implantation, whereas in others, LAAO may be performed using ICE-guided deployment. Filby et al. demonstrated that pre-procedural planning guided by cardiac computed tomography (CT), in combination with ICE-guided device deployment, achieved a 97.2% accuracy in device sizing and 100% procedural success in 71 patients, supporting the feasibility of non-TEE workflows with comparable outcomes. <sup>16</sup> This shift underscores MDCT's ability to provide comprehensive anatomical and functional data, facilitating efficient procedural planning and enabling same-day discharge strategies in select cases.

Standardized MDCT acquisition protocols for evaluating the LAA have been established using a 2-phase scan approach and may vary based on the scanner system used.<sup>17</sup> The first phase consists of a prospective or retrospective electrocardiographically gated scan acquisition during systole (typically 30-60% of the R-R interval) that provides high-resolution images for a detailed assessment of LAA morphology and size. This is followed by a delayed acquisition, performed 30-180 seconds later, to reliably exclude the presence of an LAA thrombus in patients who may have delayed opacification of the LAA. A meta-analysis of 19 studies showed that MDCT has a high negative predictive value (96-100%) for detecting LAA thrombi. Delaying imaging by at

least 30 seconds after the contrast bolus is administered is mandatory and improves the mean positive predictive value from 41% to 92% or greater. 18

Several image processing workstations (Vitrea Workstation, Aquarius Workstation, Brilliance Workstation, and 3mensio software among others) use MPR (a volume-rendering process used to produce 2D planes from 3D datasets) to reassemble and manually manipulate images of the LAA and surrounding structures. By rotating the axial view 30° right anterior and the sagittal view 10° cranial, an oblique angle is reached to view the LAA orifice en face. Here, the maximum LAA depth, ostium diameter, and landing zone diameter are measured (Figure 7A and 7B).

The 3D data provided by MDCT images enable advanced applications in procedural planning. For example, advanced software applications can assist with planning the optimal transseptal puncture location and C-arm position (Figure 7C) and display a virtual device in a 3D rendering (Figure 7D). Multidetector cardiac computed tomography datasets can also be used for 3D printing (3DP), generating models to visualize the LAAO implantation and transseptal puncture *in vitro*. Studies have demonstrated that 3DP improves device size accuracy compared to CT and TEE alone and can identify optimal transseptal puncture locations. <sup>19-21</sup> Additionally, artificial intelligence-enabled MDCT anatomical analyses and computer simulations can improve outcomes. De Backer *et al.* demonstrated that using MDCT with FEops HEARTguide simulation technology (FEops NV, Ghent, Belgium) significantly improved procedural efficiency over standard practices, reducing total procedure times (55.2±24.7 vs 45.1±18.3 min, p=0.01) and radiation times (17.6±11.4 vs 12.5±6.8 min, p<0.001). <sup>22</sup>

#### **Comparison of Imaging Modalities**

The first studies comparing imaging modalities for LAAO emerged in 2016. Wang et al. (n=53) concluded that measurement by both 2D and 3D TEE significantly undersized the LAA landing zone (p≤0.0001).<sup>23</sup> Saw et al. (n=50) highlighted that CT and TEE imaging modalities were not interchangeable, with MDCT measurements for both plug-like and disk-and-lobe devices being larger on CT than TEE.<sup>24</sup> In one small study, the use of MDCT-derived LAA measurements significantly improved device size accuracy, achieving 92% accuracy (11/12) compared to 27% accuracy (3/11) with TEE (p=0.01). This accuracy translated into enhanced procedural efficiency with reduced procedural time (55±17 vs 73±24 min, p=0.05), fewer devices required per procedure  $(1.3\pm0.7 \text{ vs } 2.5\pm1.2 \text{ devices}, p=0.01)$ , and fewer delivery sheaths (1 per case vs 1.7±0.7 sheaths, p=0.01) used.<sup>25</sup> A meta-analysis by Sattar et al. found that despite LAA orifice measurements that were significantly larger on MDCT imaging than on TEE imaging, there was no significant difference in the number of devices used or in the odds of correct device sizing when only MDCT and TEE measurements were compared. However, when studies utilizing MDCT-based 3D modeling were included, MDCT demonstrated significantly higher odds of correct device sizing compared to TEE (OR 1.64; 95% CI 1.05-2.56, p=0.03). Additionally, MDCT imaging resulted in significantly reduced fluoroscopy time compared to TEE.<sup>26</sup> These findings underscore the utility of pre-procedural MDCT imaging in improving procedural efficiency and outcomes in LAAO procedures.

#### Favorable and Unfavorable Anatomies for LAAO

The success of LAAO procedures depends on the anatomical characteristics of the LAA, which must be evaluated to select the appropriate device and optimize outcomes. Favorable and unfavorable morphologies can be described for plug-like devices and disk-and-lobe devices (Table 1).

#### Plug-like devices

Favorable Anatomies: Windsock- and cactus-shaped LAAs that have an orifice measuring 14-36 mm and a minimum depth of approximately 50% of the device width allow higher compression and better success rates (Figure 8A).

Unfavorable Anatomies: The chicken wing and cauliflower/broccoli morphologies, or orifices <14 mm or >36 mm, pose challenges with lower compression and success rates. Additionally, extremely elliptical LAA orifice or high eccentricity index (maximum diameter/minimum diameter  $\ge 1.5$ ) pose significant challenges, increasing the risk of residual leaks even with oversized devices. Insufficient depth (<50% of device length) or large mid-cavitary pectinates may lead to a large shoulder with incomplete sealing, residual leaks, or device embolization (Figure 8B).<sup>27</sup>

#### Disk-and-Lobe Devices

Favorable Anatomies: These devices perform well in LAAs with an orifice diameter of 11-31 mm and depth of ≥10-12 mm. They are ideal for anatomies with oval-shaped landing zones, large mid-cavitary pectinates, and secondary lobes (<1 cm depth). Disk-and-lobe devices are positioned more proximally than plug-like devices, allowing for adequate LAA closure in these anatomies. For LAAs that have a very short neck and sharp acute bends, such as in the chicken wing morphology, the implantation and complete closure rates are very high (98-99%) owing to the dual-closure design and ability to anchor on the proximal part of the LAA.²8

Unfavorable Anatomies: Very small (<11 mm) or very large (>31 mm) landing zone diameters and insufficient depth (<10 mm) compromise device stability and sealing.

Accurate pre-procedural imaging and detailed anatomical assessment are essential for choosing the most suitable device, enhancing procedural success, and optimizing long-term patient outcomes.

#### INTRA-PROCEDURAL EVALUATION

#### **Transesophageal Echocardiography**

Intra-procedural TEE is the primary imaging modality for LAAO and was used in all the LAAO clinical trials. Intra-procedural TEE is performed to confirm LAA measurements and to rule out LAA thrombi. When spontaneous echocardiographic contrast (SEC) or sludge is present in the left atrium or LAA, differentiating a thrombus from the sludge/SEC becomes challenging, necessitating additional TEE enhancements. <sup>29</sup> Guidelines from the American Society of Echocardiography recommend the consideration of ultrasound enhancing agents during TEE for the assessment of LAA thrombus when there is SEC or poor visualization (COR IIa, LOE B-NR). <sup>30</sup> Ultrasound enhancing agents improve thrombus identification and differentiation of SEC degrees, with studies demonstrating increased interpretative confidence, reader agreement, and prevention of procedure cancellations. <sup>31</sup> Dobutamine and isoproterenol, both positive inotropic agents, have been shown to resolve LAA SEC and significantly increase peak LAA velocity. <sup>32,33</sup> There were no strokes, systemic embolisms, or device-related thrombi reported on follow-up with the use of dobutamine or isoprotenerol. <sup>34</sup>

Disk-and-lobe and plug-like devices both require transfemoral venous access and have several common steps (Figure 9, Videos 1-4). Once femoral access is obtained, TEE imaging is utilized to guide the transseptal puncture. A delivery catheter and needle are advanced to the interatrial septum. Both 2D or 3D TEE views can be used to visualize the inferior, superior, anterior, and posterior portions of the interatrial septum. The choice of transseptal puncture site

is guided by the LAA's anatomy, with an inferoposterior puncture recommended in 75-80% of cases to ensure coaxial alignment between the delivery sheath and the proximal LAA central axis. However, an inferior but more central or anterior transseptal puncture is preferred in 20-25% of cases, particularly in those with reverse chicken wing LAA morphology or posterior bending of the proximal LAA. 35,36 As most LAAs are anteriorly directed, a posterior puncture is desirable in most cases. The 90° TEE view aids in gauging LAA direction, complemented by pre-procedural cardiac CT analysis. Once the delivery sheath has crossed the interatrial septum, it is advanced into the left atrium and the distal end of the sheath is positioned near the LAA ostium.

The plug-like device is initially unsheathed to form a ball, which can be manipulated in the LAA to achieve an ideal trajectory and then be fully deployed. Figure 10 and Videos 5-8 outline the steps for deployment of a plug-like device. The location of the circumflex artery in relation to the ostium should be noted, with the plane of maximal device diameter positioned at or just distal to the ostium, ensuring that any protrusion or "shoulder" (if present) does not exceed 40-50% of the device depth. For plug-like devices, the anatomical orifice is defined by a line connecting the circumflex artery and a point 10-20 mm inside the LAA from the left upper pulmonary vein ridge, described as the beginning of the trabeculated LAA, and is intended to be covered by the device. Compression is measured from shoulder to shoulder with the central metallic "threaded insert" visible on the left atrial side and is assessed at 0°, 45°, 90°, and 135°. A "tug test" is performed under TEE and/or fluoroscopic visualization to confirm stability, ensuring that the device returns to its original position. Once stability is confirmed, peri-device leaks (PDLs) should be carefully assessed using multiplanar imaging at 0°, 45°, 90°, and 135°

with color Doppler and the Nyquist limit set at 20-30 cm/sec to detect low-velocity flows. Any significant PDL should be avoided, and the device recaptured and redeployed if needed.

For disk-and-lobe devices, the steps for deployment are outlined in Figures 11 and 12 (Videos 9-17). The device lobe should be two-thirds deeper than the circumflex artery to ensure proper sealing and compression. The lobe is initially unsheathed to form a ball. This ball can be manipulated into the LAA. Once the position and trajectory are optimized, this lobe is fully deployed, initially appearing as a triangle shape before full deployment. The disk is deployed subsequently, ensuring that it covers the anatomic ostium of the LAA and does not impinge upon the left superior pulmonary vein, which is separated from the LAA by a ridge.

Once the device is deployed, each type of device has a protocol for the assessment of position, stability, compression, and seal (Figure 13 for plug-like device, Figure 14 for disk-and-lobe device). A color Doppler "twinkling" artifact might be noted over the proximal surface of nitinol-based PTFE-covered devices. This artifact arises from ultrasound interactions with the strongly reflective surface and appears as a red-blue shimmer, mimicking flow, and should not be misinterpreted as PDL. This can be distinguished by its diffuse pattern, a lack of continuity across the device, and the absence of a corresponding signal on spectral Doppler. If the release criteria are met, the device is released and the guide catheter is withdrawn from the left atrium; color Doppler and 2D TEE are then used to assess the size of the atrial septal defect and flow direction. A 3D view of the LAAO device is obtained to confirm its proximity to the mitral annulus and surrounding structures. A final assessment of pericardial effusion is performed prior to removing the TEE transducer.

#### **Micro- and Mini-TEE**

Micro- and mini-TEE probes, originally designed for pediatric diagnostic imaging, are emerging as alternatives to standard TEE probes for LAAO guidance, particularly in frail patients or those for whom general anesthesia is unsuitable. These smaller-diameter probes (with 32 and 48 imaging elements, respectively, vs. >2,500 in standard TEE) enable the use of conscious sedation, thereby reducing anesthesia-related risks and logistical challenges. In a recent study, micro- and mini-TEE provided adequate imaging in 99.3% of LAAO cases, with only 0.7% of cases requiring conversion to standard TEE, supporting the feasibility of these probes for this procedure. However, limitations include reduced image quality as a result of fewer imaging elements, challenges in maintaining esophageal wall contact because of their smaller flexible design, and diminished far-field resolution in patients with large atria owing to higher probe frequencies. Despite these challenges, micro- and mini-TEE are particularly appealing to operators who seek minimalist LAAO approaches, offering a balance between TEE's familiarity and ICE's reduced anesthesia requirements, with the added benefit of lower gastrointestinal complication risk compared to standard TEE.

### **Intracardiac Echocardiography**

Intracardiac echocardiography has gained popularity in guiding LAAO procedures, particularly with the advent of 3D real-time ICE imaging. The procedure follows the same steps as with TEE. The ICE transducer is inserted into the right atrium and utilized to cross the interatrial septum. Imaging in 3D is essential to evaluate anterior-posterior and superior-inferior portions of the septum. Once the septum has been crossed, the ICE transducer is advanced through the puncture site into the left atrium. With the ICE transducer in the left atrium, the absence of any LAA thrombi is confirmed and measurements of the LAA ostium and depth are obtained, preferably from the 3D dataset. Intracardiac echocardiographic views should be optimized to create 3

"TEE-like" views (Figure 15) to optimize the position and trajectory of the device. The mid left atrial view resembles the 45° TEE view and is used during LAA device deployment. The left superior pulmonary vein view resembles the 90° TEE view and is used to assess maximum LAA depth. The supramitral view is akin to the 135° (high angle) TEE view, which is critical to assess device position, seal, and compression (Figure 15).

At this stage, it is unclear whether ICE-guided LAAO is a more cost-efficient method for LAA procedures and whether it is associated with better outcomes and shorter procedure times. In some cases, ICE could potentially eliminate the need for general anesthesia.<sup>38</sup> In a study by Berti et al., ICE guided LAAO with a procedural success rate of 93% and there was a significant correlation in measurements for device sizing between ICE and angiography.<sup>39</sup> A recent metaanalysis demonstrated that ICE provides comparable efficacy and safety, with reduced use of contrast and shortened fluoroscopy and procedure times. 40 However, a multicenter study showed that, although ICE-guided LAAO achieved high procedural success rates, TEE-guided procedures were associated with shorter procedure and fluoroscopy times.<sup>41</sup> The potential for ICE-guided LAAO without fluoroscopy is a significant advancement, with studies demonstrating feasibility for sizing, deployment, and PDL assessment without contrast. Successful cases with plug-like devices achieved zero fluoroscopy, with comparable procedural times to the conventional method. 42,43 This approach reduces radiation risks and eliminates the need for general anesthesia, enhancing patient comfort, although it involves a larger transseptal puncture and a learning curve. Further, each modality comes with unique complications: whereas TEE has been associated with higher gastrointestinal complications, ICE has been associated with increased peripheral vascular and renal complications. 44 Additional considerations include the cost of the ICE catheter, the learning curve for operators, and current reimbursement challenges.

#### **Complications of LAAO Procedures**

Intra-procedural complication rates are low with both types of devices. Pericardial effusion/tamponade can be a complication of transseptal puncture. Other complications include device-related embolization and procedure-related ischemic stroke. <sup>10</sup> The disk-and-lobe device has been found to have a slightly higher risk of pericardial tamponade or major bleeding. <sup>45</sup> Table 2 details the complications of LAAO procedures. <sup>7,10,46-48</sup> Importantly, the rate of complications decreases with operator experience. Intraprocedural imaging with TEE or ICE is key for both device placement and the monitoring of these complications.

#### POST-PROCEDURAL EVALUATION

#### **Post-Procedural Imaging Protocols**

The 2023 consensus statement by the Society for Cardiovascular Angiography and Interventions and the Heart Rhythm Society recommends follow-up imaging with TEE or MDCT between 45 and 90 days post-LAAO to assess device sealing. A second follow-up at 1 year should be considered if there is concern about device-related thrombus (DRT) or PDL. If DRT was detected, repeat imaging using TEE or MDCT every 45 to 90 days is recommended.<sup>36</sup>

Nestelberger *et al.* recommended similar surveillance strategies: TEE or MDCT 6-12 weeks post-LAAO and another routine TEE or MDCT at 12 months if the patient is at high risk for DRT. A key difference from the 2023 consensus statement is their recommendation for routine transthoracic echocardiography at 12 months to assess atrial sizes, ventricular and valvular function, and potential late device embolization. They also favor MDCT for surveillance at 6-12 weeks owing to its higher spatial resolution, sensitivity in the detection of PDL, and non-invasive and non-fasting nature. However, they acknowledged the radiation and subsequent risk

associated with CT utilization and emphasized that TEE remains a valuable modality for LAAO follow-up imaging.<sup>49</sup>

# **Imaging of DRT**

DRT is one of the more troublesome findings on follow-up imaging, with the reported incidence ranging from 1.7% to 7%.<sup>50,51</sup> Device-related thrombi vary widely in size,<sup>52</sup> and their clinical impact is worrisome (Table 3). Few trials were reassuring,<sup>9,56</sup> and the majority of data point to a 3.5-4 times higher risk of ischemic events in patients with DRT than in those without DRT.<sup>53,54,57</sup> The timing of the occurrence of DRT is somewhat unpredictable: 58% of DRT cases were diagnosed after 3 months in a meta-analysis and 18-36% of DRT were seen after 6 months in large registry studies, well after the presumed device endothelialization period.<sup>52,53,57</sup>

Predicting the occurrence of DRT requires consideration of several clinical, imaging, and procedural risk factors. The most commonly cited clinical risk factors include advanced age, <sup>50,58</sup> prior embolic events, <sup>54,58</sup> higher CHA<sub>2</sub>DS<sub>2</sub>-VASc score, <sup>59</sup> and permanent AF. <sup>50,52</sup> Pre-implant imaging findings that are associated with subsequent DRT include larger LAA orifice width, larger left atrial size, lower left ventricular ejection fraction, lower peak emptying velocity, and SEC. <sup>54,57,60-62</sup> Interestingly, neither LAA morphology nor anticoagulation regimen predicted DRT occurrence. <sup>53,57</sup>

Deep implantation has emerged as the strongest procedural risk factor for DRT.<sup>57,59,63</sup> Similarly, one study correlated the uncovered area—defined as the triangular area between the tip of the ridge, the surface of the device, and the mitral annulus—with the risk of DRT.<sup>63</sup> This cul-de-sac between the exposed ridge and the face of the device has been termed a "neo-appendage"<sup>63,64</sup> and has been shown to increase turbulence. Additionally, off-axis deployment

and residual PDL have also been associated with DRT risk.<sup>57,65</sup> Finally, the literature suggests that DRT can also occur centrally, adherent to the central screw of the device (Figure 16).<sup>52,57</sup>

TEE has historically been the standard post-deployment surveillance imaging modality.<sup>66</sup> The expert consensus definition of DRT by TEE is an echo density that is not explained by artifact, not typical of healing, visible in multiple planes, in contact with the device, and independently mobile.<sup>67,68</sup> The rise in MDCT imaging has yielded markedly high rates of hypoattenuated thickening (HAT), which resembles thrombus but has an uncertain clinical significance. Recent studies show HAT in a significant majority of cases, leading to classification of HAT into low-grade (smooth, covering the entire device, and continuous with the left atrial wall) and high-grade (not continuous with the left atrial wall, having irregular borders or being pedunculated).<sup>69,70</sup> High-grade HAT, which occurs far less commonly at 2.8-5%, is associated with an increased risk of stroke compared to low-grade HAT.<sup>69,71</sup>

### **Imaging of PDL**

Peri-device leak following LAAO presents a notable risk for thromboembolic complications (Table 4). The characteristics of these leaks can vary significantly based on the occlusion device used and the anatomical features of the LAA. With plug-like devices, PDL typically manifests as flow between the left atrium and the distal part of the LAA. With disk-and-lobe devices, PDL may occur between the left atrium and the space between the disk and the lobe or beyond the lobe itself.

The mechanisms of leaks for plug-like devices include edge leak, uncovered lobe or proximal LAA, and fabric leak (Figure 17). For disk-and-lobe devices, leaks can occur between the disk and the lobe, at the distal LAA owing to uncovered proximal tissue, or from fabric

leak.<sup>77</sup> The clinical implications of the different types of leaks with disk-and-lobe devices have not been adequately studied, highlighting the need for further research.

Size of PDLs and Thromboembolic Risk

Traditionally a vena contracta of PDL edge leak of <5 mm has been considered acceptable for the plug-like device; however, emerging data challenge this threshold. 9,72,73,75 Table 3 summarizes key studies showing the PDL incidence and its association with ischemic events. Alkhouli *et al.* reported a 10-15% increase in the 1-year risk-adjusted rates of systemic thromboembolic events for patients with small leaks, with a hazard ratio of 1.152 (95% CI: 1.025-1.294), compared to patients without leaks. Though rare, have been associated with a significantly higher risk of thromboembolism when compared to patients with no PDL. This underscores the importance of accurate detection and ongoing surveillance of these leaks to manage and mitigate associated risks effectively.

Imaging Modalities for Detecting PDL

Detecting and characterizing PDL is critical yet challenging owing to varied detection methods and the lack of a standardized definition. Transesophageal echocardiography and MDCT are the primary imaging modalities used to assess PDL. In a study involving 346 patients who underwent both TEE and MDCT at 8 weeks post-LAAO with a disk-and-lobe device, PDL was detected in 110 patients (32%) via TEE whereas MDCT identified PDL in 210 patients (61%). Another study also demonstrated that the incidence of PDL detected by MDCT was higher (52%) than observed with TEE (34.3%). These studies indicate a substantially higher occurrence of PDL detection with MDCT compared to TEE, revealing a significant discrepancy in leak quantification between these modalities. However, no standardized sizing thresholds currently

exist for cardiac CT, since initial trials were performed with TEE. This highlights an area where further research is needed.

#### CONCLUSIONS

Multimodality imaging is essential for optimizing outcomes in LAAO. Pre-procedural evaluation using TEE and MDCT is essential for assessing LAA morphology and determining the appropriate device size. Multidetector cardiac computed tomography offers superior multiplanar and 3D imaging, whereas TEE provides functional insights that are vital for determining patient eligibility and selecting the optimal device for the procedure. Intra-procedural imaging with TEE or ICE ensures real-time guidance for accurate and efficient device placement. Post-procedural follow-up with TEE or MDCT is essential for monitoring device stability and assessing PDL and DRT (Central Illustration).

The selection of appropriate imaging modalities for percutaneous LAAO is crucial for ensuring optimal outcomes. The integration of multimodality imaging—leveraging the strengths of TEE, MDCT, and ICE—is essential for accurate device sizing, successful implantation, and timely detection of complications. As imaging technologies advance, these strategies will continue to evolve, further improving long-term outcomes for patients undergoing LAAO.

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#### **Declarations of Interest**

MS is a member of the speaker bureau for Abbott, Boston Scientific, Medtronic, and Philips and the advisory board of Siemens. RJ is a speaker and consultant for Medtronic, Philips, Edwards and GE Healthcare and a member of the advisory board for Medtronic and GE Healthcare.

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#### Legends

Figure 1 Commercially approved percutaneous LAAO devices. (A) Plug-like device: This is a self-expanding nitinol device equipped with dual-row precision fixation anchors for enhanced stability. It features a low-profile threaded insert covered by polyethylene terephthalate fabric and is available in 6 sizes, 20-40 mm. (B) Disk-and-lobe device: Composed of a self-expanding nitinol mesh, this device forms a lobe and a disk, connected by a central waist. It is available in 8 sizes, 11-31 mm. Device images are used with permission from Boston Scientific and Abbott Cardiovascular.

**Figure 2** LAA morphologies illustrated in 2D and 3D TEE. (A) Windsock, (B) chicken wing, (C) cactus, and (D) cauliflower/broccoli LAA morphologies are demonstrated in 2D (left side of each panel) and 3D. 3D TEE utilizes light-source manipulation and transillumination technology to provide high-definition, photorealistic, 3D volume-rendered images of the LAA.

Figure 3 LAAM maneuver to characterize LAA morphology by 3D TEE. (A) Obtain a 3D midesophageal LAA en face view at a 2D 45° angle (green line) with the transillumination feature activated. Establish an orthogonal plane (red line) using Multiview. (B) Select the red reference plane and display the cropping plane (dotted lines). (C) Adjust the cropping plane leftward to reveal the full 3D anatomy of the LAA. (D) Rotate the 3D image 90° counterclockwise to simulate the standard fluoroscopic view used during percutaneous LAAO procedures. MV, mitral valve; PA, pulmonary artery.

**Figure 4** TUPLE maneuvers to characterize LAA morphology by 3D TEE. (A) Display the standard en face view of the LAA orifice from the left atrial perspective. (B) Apply light-source manipulation to enhance visual details. (C) Utilize transillumination rendering of the LAA, increasing transparency to clearly demonstrate the orifice and outline the walls of the LAA body.

- (D) Rotate the image leftward along the vertical axis of the LAA to reveal the full body.
- (E) Selectively crop tissue to focus on specific areas. (F) Finalize the fully rendered LAA volume after decreasing transparency and increasing image gain. This orientation corresponds to the 2D TEE image at 135°. AV, aortic valve; MV, mitral valve; PA, pulmonary artery.

**Figure 5** LAA sizing for LAAO devices. (A) 2D TEE sizing for a plug-like occlusion device.

The LAA landing zone diameters and depths are measured at 4 angles: 0°, 45°, 90°, and 135°. The LAA landing zone, which can be up to 2 cm below the left atrial ridge, is depicted by a dotted white line measured at the level of the circumflex artery (red arrow). (B) 2D TEE sizing for the disk-and-lobe device. The LAA landing zone diameters and depths are measured at 4 angles: 0°,

orifice (yellow arrow), and the depth is measured from the ostium to the LAA wall in a plane

45°, 90°, and 135°. The LAA landing zone diameter is measured 10-12 mm from the anatomic

perpendicular to the ostium. The red arrow indicates the circumflex artery.

Figure 6 Sizing of the LAA orifice using 3D TEE MPR. (A) 3D view of the LAA from the left atrium; the degrees shown correspond to 2D omniplane TEE views to assess the LAA. (B) A single-beat zoom capture acquires the entire LAA and surrounding structures. MPR is obtained. The red and green planes are locked and oriented toward the LAA apex, leaving the blue plane free for adjustment. The blue plane is oriented toward the plane of the LAA orifice, typically at the level of the left circumflex coronary artery (red arrow). The 3D Auto LAA feature (Philips) is used to obtain the maximum diameter of the LAA orifice.

**Figure 7** Cardiac computed tomography for pre-procedural planning for LAAO. In addition to excluding LAA thrombi, MDCT with 3D MPR can be utilized to perform precise measurements of the expected landing zone (A), LAA orifice (B), LAA depth, 3D morphology, and relations with the surrounding structures. Advanced software applications can also be of assistance in

planning the optimal transseptal puncture location and C-arm position (C) and display a virtual device in the 3D rendering (D).

Figure 8 Anatomical considerations for LAAO device selection. (A) Favorable morphologies for plug-like LAAO devices include windsock- and cactus-shaped LAAs, typically with orifice diameters ranging from 14-36 mm and depths approximately 50% of the intended device width. Favorable anatomies for disk-and-lobe devices include LAAs with orifice diameters of 11-31 mm and depths of at least 10-12 mm. (B) Unfavorable morphologies for plug-like devices include the chicken wing and cauliflower/broccoli shapes with shallow depths and orifice diameters <14 mm or >36 mm. Unfavorable conditions for disk-and-lobe devices include very small (<11 mm) or very large (>31 mm) landing zone diameters with insufficient depth (<10 mm).

Figure 9 Common steps for LAA closure procedure. (A) Transeptal access. Guided by TEE and fluoroscopy, transseptal access is usually achieved through an inferior and posterior approach. Yellow arrows demonstrate the sheath (Video 1). (B) Guidewire placement. The crossing sheath is exchanged with a guidewire (yellow arrow), which is placed in the LUPV (Video 2). (C) Delivery system advancement. A delivery system with a pigtail catheter (yellow arrow) is advanced into the left atrium. Contrast angiography is performed to fluoroscopically evaluate the LAA (Video 3). (D) Catheter navigation. The guide catheter/pigtail combination is navigated to align the corresponding radiopaque marker band for the device size with the LAA ostium. Once the guide catheter (yellow arrow) is properly positioned, the pigtail is removed. Yellow arrows demonstrate the device and sheath (Video 4).

**Figure 10** Plug-like device procedure with TEE. (A) The delivery system (yellow arrow, left panel) is advanced until the distal marker bands of the delivery system and access sheath are

aligned. The device is then unsheathed slowly until a ball shape (yellow arrow, right panel) is visualized (Video 5). (B) The device (yellow arrows) is fully unsheathed, and its position is evaluated using fluoroscopy and TEE (Video 6). (C-E) A pre-release evaluation is conducted using the "PASS" criteria (Position: Ensure that the plane of maximum diameter of the device is at or distal to the LAA ostium. Anchor: Test stability by retracting the deployment knob and letting go, assessing whether the device returns to its original position. Size: Confirm that the device shoulder is compressed to 10-30% of its original size. Seal: Use TEE to assess for any residual flow, which must be <5 mm before release (Video 7). Once all "PASS" criteria are met, the device may be released to deploy (Video 8).

Figure 11 Disk-and-lobe procedure steps with TEE. (A) The cable is advanced to push the device to the tip of the delivery sheath. Subsequently, the sheath is backed out until the device forms the "ball" (yellow arrow). It is essential to verify that the ball is coaxial to the neck of the LAA (Videos 9, 10). (B) The cable is advanced slowly to form the "triangle" shape (yellow arrow) (Video 11). (C) The delivery cable is advanced further to complete the deployment of the device lobe (yellow arrow) (Video 12). (D) Deployment of the device disk (yellow arrow) is completed by advancing the delivery cable while simultaneously unsheathing the device disk (Video 13).

Figure 12 Disk-and-lobe deployment steps with TEE. (A) Once the disk is deployed, a tension test is conducted to ensure the lobe position remains unchanged (Video 14). The yellow arrows point to the football shape of the disk. (B) Residual flow around the device is assessed using TEE (Video 15). (C) After checking for any leaks and verifying a suitable position for release, the "CLOSE" criteria are re-evaluated before the device is released (Video 16). (D) Following deployment, the device is visualized on conventional 3D TEE (Video 17). The red arrow

indicates the proximal end screw. (E) The disk-and-lobe occluder device using light source manipulation, showcasing a figure of 8. Yellow arrows demonstrate the device and sheath.

**Figure 13** Assessment of device release PASS criteria for plug-like device. The yellow arrows indicate the threaded insert, the red arrow indicates the left circumflex artery, and the blue double-headed arrow indicates compression of 15%.

Figure 14 Assessment of device release CLOSE criteria for disk-and-lobe device.

**Figure 15** Standard ICE views for LAAO. (A) The ICE catheter is visualized in the left atrium on fluoroscopy. The three key ICE probe positions commonly utilized to visualize the LAA during LAAO are the (B) mid left atrial view (equivalent to TEE 45° view), the (C) left superior pulmonary vein view (equivalent to TEE 90° view), and the (D) supramitral view (equivalent to TEE 135° view).

**Figure 16** DRT. (A) A thrombus (arrow) on a deeply implanted device is seen on 2D TEE. (B) A thrombus (arrow) on the threaded insert of a device is seen on 2D TEE. (C) A device-related thrombus (arrow) in the neo-appendage created between the device face and the left atrial pulmonary vein/LAA ridge is seen on 3D TEE.

**Figure 17** Mechanisms of residual leaks after LAAO with a plug-like occluder. (A) Edge leak seen on TEE. (B) Uncovered lobe seen on CT. (C) Fabric leak seen on TEE.

**Central Illustration** Utility of multimodality imaging for LAAO.

Table 1 Favorable and unfavorable anatomies for left atrial appendage occlusion

| Parameter                              | Favorable characteristics   | Unfavorable characteristics              |
|--|---|--|
| Plug-like device                       |   |  |
| Landing zone                           | 14-36 mm  | >36 mm                                   |
| Depth                                  | ~50% of device width  | <50% of device width                     |
| Morphology                             | Windsock<br>Cactus  | Chicken wing,<br>Cauliflower/broccoli    |
| Landing zone shape/ eccentricity index | Round shape   | Elliptical shape ≥1.5 eccentricity index |
| Pectinates                             |   | Large mid cavitary pectinates            |
| Disk-and-lobe device                   |   |  |
| Landing zone                           | 11-31 mm  | <11 mm or >31 mm                         |
| Depth                                  | ≥10-12 mm   | <10 mm                                   |
| Morphology                             | Secondary lobes <1 cm from landing zone Chicken wing with short neck, acute bends |  |
| Landing zone shape/ eccentricity index | Oval landing zone   |  |
| Pectinates                             | Large mid cavitary pectinates   |  |

Table 2 Complications of left atrial appendage procedures 7,10,46-48

| Complications                | Percentage |
|------------------------------|------------|
| Serious pericardial effusion | 1.2%-4.8%  |
| Major bleeding               | 0.6%-1.3%  |
| Ischemic stroke              | 0.2-0.7%   |
| Device embolization          | 0.1%-0.6%  |
| Procedure-related death      | <0.5%      |
| Total major safety events    | 3.2%-8.7%  |

Table 3 Summary of studies reporting DRT incidence, imaging modalities, and association with ischemic events

| Study   | DRT incidence   | Patient factors   | Procedural factors  | Outcomes   |
|---|---|---|---|--|
| Meta-analysis<br>Alkhouli et al. <sup>53</sup>  | <ul> <li>3.8% (pooled incidence)</li> <li>Following LAAO</li> <li>90 days: 42%</li> <li>90 to 365: 57%</li> <li>&gt;365 days: 1%</li> <li>No difference in DRT incidence between devices</li> </ul> | No specific consistent<br>predictor was<br>identified   | No specific consistent<br>predictor was<br>identified   | Increase in ischemic events (13.5% vs. 4.4%; OR 4.15; 95% CI 2.77-6.22; p<0.001; I <sup>2</sup> =0)  |
| Protect AF,<br>PREVAIL, CAP,<br>and CAP2 ad<br>hoc analysis<br>Dukkipati et al. <sup>54</sup> | <ul> <li>3.74%</li> <li>51% of DRT detected by scheduled TEE at 1 year</li> <li>62% detected during unscheduled TEE at 1 year</li> </ul>  | <ul> <li>Permanent AF (OR 2.24; 95% CI 1.19-4.2, p=0.012)</li> <li>History of TIA/stroke (OR 2.31; 95% CI 1.26-4.25; p=0.007)</li> <li>Vascular disease (OR 2.06; 95% CI 1.08-3.91; p=0.028)</li> </ul> | <ul> <li>LAA diameter (OR 1.06; 95% CI 1.01-1.12; p=0.019</li> <li>Lower LVEF (OR 0.96 per 1% increase; 95% CI 0.94-0.99; p=0.009)</li> </ul> | Higher rates of ischemic stroke or systemic embolization (adjusted HR 3.9; 95% CI 2.3-6.8; p<0.001)  |
| Amulet IDE<br>Schmidt et al. <sup>55</sup>  | <ul> <li>3.9% through 18 months</li> <li>Amulet: 82% DRT ≤45 days</li> <li>Watchman: 73.8% DRT &gt;45 days</li> </ul>   | • AF (HR 2.44; 95% CI 1.42-4.22; p<0.01) • Female sex (HR 1.65; 95% CI 1.01-2.71; p=0.04) • Older age (HR 1.04; 95% CI 1.01-1.08; p=0.02)   |   | No statistically significant associations between DRT and ischemic stroke or systemic embolization (3.1% vs 2.6% with vs without DRT), although numerically there were more strokes and systemic embolization after DRT vs no-DRT in |

| the Watchman arm        |
|-------------------------|
| (5.5% vs 2.5%; HR       |
| 2.15; 95% CI 0.50-9.19) |

EUROC-DRT registry Sedaghat et al.<sup>52</sup>

- Median of 93 days (IQR 54-161)
- 82% detected <6 months
- 20% detected >6 months

• Deep implantation

Higher stroke rates in patients with residual DRT vs DRT resolution after 1 year (7.6% vs. 6.5%; p=0.09; mortality 15.0% vs. 1.4%; p=0.01)

EWOLUTION registry Sedaghat et al. 56

- 4.1% by TEE or CT at a median 54 days (IQR 42-111)
- 91.2% of DRT detected <3 months or at first TEE

• Permanent AF (82.4 vs. 64.9%; p<0.01)

• Dense SEC (26.5 vs. 11.9%; p=0.03)

• LAA diameter (22.8±3.5 vs. 21.1±3.5 mm; p<0.01)

No difference in rate of ischemic stroke/TIA (DRT 1.7 vs. no-DRT 2.2% per year; p=0.8).

Global DRT registry Simard et al.<sup>57</sup>

- 2.8%
- ≤180 days: 64%
- >180 days: 36%
- Hypercoagulopathy (OR 17.50; 95% CI 3.39-90.45)
- Permanent AF, (OR 1.90; 95% CI 1.22-2.97)
- Renal insufficiency (OR 4.02; 95% CI 1.22-13.25)
- Pericardial effusion (OR 13.45; 95% CI 1.46-123.52)
- Deep implantation >10 mm from limbus (OR 2.41; 95% CI 1.57-3.69)

Increased rates of ischemic stroke in DRT vs no-DRT patients (16.9% vs. 3.6%; p=0.0001)

| Multicenter registry c et al. <sup>58</sup>                              | <ul> <li>7.2% per year</li> <li>8.3% with TEE and 5.3% with CT</li> <li>Mean time: 3.1±2.6 months</li> </ul>                                       | <ul> <li>Older age (HR 1.07; 95% CI 1.01-1.14; p=0.02)</li> <li>Prior TIA/stroke (HR 3.68; 95% CI 1.17-11.62; p=0.03)</li> </ul>  | <ul> <li>No OAC (HR 0.26;<br/>95% CI 0.09-0.77;<br/>p=0.02)</li> <li>APT post-LAAO (HR 0.10; 95% CI 0.01-<br/>0.76; p=0.03)</li> </ul> | Independent predictor of stroke and TIA (HR 4.39; 95% CI 1.05-18.43; p=0.04)  |
|--|--|---|--|---|
| Single center<br>Kaneko et al. <sup>59</sup>                             | <ul><li>5% DRT</li><li>5.1% at 45-day follow-up</li></ul>  | • Higher CHA <sub>2</sub> DS <sub>2</sub> -<br>VASc score (OR 2.8;<br>95% CI 1.2-7; p=0.02)   | • Deep device implantation (OR 24.7; 95% CI 1.3-458.2; p=0.03)   | Not assessed  |
| Prospective<br>global Amulet<br>registry<br>Aminiam et al. <sup>60</sup> | <ul> <li>1.7% per year</li> <li>DRT scheduled follow-<br/>up TEE in 10 cases,<br/>unscheduled TEE 7<br/>cases, and CT 1 patient</li> </ul>         | • Larger LAA orifice width (HR 1.09, 95% CI 1-1.19; p=0.04)   |  | Greater risk for ischemic stroke/TIA (HR 5.27; 95% CI 1.58-17.55; p=0.007)  |
| EUROC-DRT<br>registry<br>Vij et al. <sup>61</sup>                        | <ul> <li>20% patients with DRT</li> <li>TEE: 88.4% of cases detected</li> <li>CT: 11.6% of cases detected</li> <li>Mean of 147±219 days</li> </ul> | <ul> <li>Age (OR 1.16; 95% CI 1.07-1.25; p&lt;0.01)</li> <li>Prior stroke/TIA (OR 3.71; 95% CI 1.19-11.56; p=0.02)</li> <li>SEC (OR 5.08; 95% CI 1.37-18.84; p=0.02)</li> </ul> | Deep implantation<br>predictive in univariate<br>analysis after PSM  | Significant increased risk of stroke with DRT 13.55, compared to no DRT 3.8% within 2 years (HR 4.21; 95% CI 1.88-9.49; p<0.01) |

AF, atrial fibrillation; APT, antiplatelet; CI, confidence interval; CT, computed tomography; DRT, device-related thrombus; HR, hazard ratio; IQR, interquartile range; LAA, left atrial appendage; LAAO, LAA occlusion; LVEF, left ventricular ejection fraction; OAC, oral anticoagulation; OR, odds ratio; PSM, propensity score matching; SEC, spontaneous echocardiographic contrast; TEE, transesophageal echocardiography; TIA, transient ischemic attack.

Table 4 Summary of studies reporting PDL incidence, imaging modalities, and association with ischemic events

| Study                                    | Device(s)   | Study<br>populat<br>ion | Imaging<br>modality       | PDL incidence   | Association of PDL with ischemic events   |
|--|---|-------------------------|---------------------------|---|---|
| NCDR LAAO<br>registry <sup>72</sup>      | Watchman 2.5  | 51,333                  | TEE (45 days)             | <5 mm: 25.8%<br>>5 mm: 0.7%   | Small PDLs (1-5 mm): ↑ stroke/TIA/systemic embolization (aHR 1.15; 95% CI 1.02-1.29) compared to no PDL   |
| PROTECT AF & PREVAIL+CAP2                | Watchman 2.5  | 1,054                   | TEE (45 days, 12 months)  | ≤3 mm: 24.2%<br>3-5 mm: 14.1%<br>>5 mm: 1.5%  | PDL ≤5 mm at 1 year was a significant predictor of ischemic stroke/systemic embolism (HR 1.75; 95% CI 1.06-2.89; p=0.03)  |
| AMULET IDE <sup>9</sup>                  | Amulet and<br>Watchman 2.5                            | 1,878                   | TEE (45 days, 12 months)  | At 45 days  Amulet  0-3 mm: 27%  >3-5 mm: 9%  >5 mm: 1%  Watchman  0-3 mm: 29%,  3-5 mm: 22%  >5 mm: 3% | Not assessed  |
| AMULET IDE<br>Sub-analysis <sup>74</sup> | Amulet (801 patients) and Watchman 2.5 (792 patients) | 1,593                   | TEE at 45 days, 12 months | At 45 days Watchman 0-3 mm: 74.1% ≥3-5 mm: 25.9% >5 mm: 3.2% Amulet 0-<3mm: 88.8%                       | PDL ≥3 mm: higher 1-month rates of ischemic stroke or systemic embolization compared with those with PDL <3 mm (3.6% vs 1.8%; unadjusted HR 2.03; 95% CI 0.96-4.29; p=0.06) |

|   |                        |              |                                     | ≥3-5mm: 1.2%<br>>5 mm: 1.1%  |   |
|---|------------------------|--------------|-------------------------------------|--|---|
|   |                        |              |                                     | p<0.01 for all   |   |
|   |                        |              |                                     | At 12 months  Watchman – residual PDL  ≤5 mm: 97.2%  <5 mm: 2.8%  Watchman – new PDL  ≥3 mm: 8.3%  Amulet – residual PDL  ≤5 mm: 99.4%  >5 mm: 0.6%  Amulet – new PDL  ≥ 3mm: 4.2% |   |
| Meta-analysis of<br>residual leaks<br>following<br>LAAO <sup>75</sup> | Amulet and<br>Watchman | 61,666       | TEE, CT                             | TEE  Any PDL: 26.5%  >1 mm: 15%  >3 mm: 9.6%  >5 mm: 0.9%  CT  PDL >0 mm: 57.3%  | Any TEE-reported PDL was significantly associated with 2-fold increase odds of thromboembolism (pOR 2.04, 95% CI 1.52-2.74; I <sup>2</sup> =28%) compared to no PDL |
| PINNACLE<br>FLX <sup>51</sup>   | Watchman FLX           | 400 patients | TEE (45 days, 6 months, and 1 year) | 17.2% at 45 days 10.5% at 6 months (>0 and ≤5 mm) No PDL >5 mm   | Not assessed  |
| EWOLUTION <sup>56</sup>   | Watchman               | 835          | TEE (45 days)                       | 8.4% (any PDL)   | Not assessed  |

Global Amulet 1,088 TEE (45 days) 9.9% (any PDL) Not assessed Study<sup>76</sup>

aHR, adjusted hazard ratio; CI, confidence interval; CT, computed tomography; HR, hazard ratio; LAAO, left atrial appendage occlusion; OR, odds ratio; pOR, pooled odds ratio; PDL, peri-device leak; TEE, transesophageal echocardiography; TIA, transient ischemic attack.

### **Video Legends**

Video 1 Common steps – transseptal wire on 2D transesophageal echocardiography

**Video 2** Common steps – wire in the left upper pulmonary vein on 2D transesophageal echocardiography

**Video 3** Common steps – pigtail placement in the left atrial appendage on 2D transesophageal echocardiography

**Video 4** Common steps – sheath guided into the left atrial appendage on 3D biplane transesophageal echocardiography

**Video 5** Plug-like device – "ball" shape visualized on 3D biplane transesophageal echocardiography

**Video 6** Plug-like device – deployed device on 3D biplane on 2D transesophageal echocardiography

**Video 7** Plug-like device – deployed device assessed with color Doppler to evaluate for peridevice leak on 2D transesophageal echocardiography

**Video 8** Plug-like device – released device in the left atrial appendage on 3D transesophageal echocardiography

**Video 9** Disk-and-lobe device – "ball" shape visualized on 3D biplane transesophageal echocardiography

**Video 10** Disk-and-lobe device – "ball" shape visualized on 3D transesophageal echocardiography

**Video 11** Disk-and-lobe device – "triangle" shape visualized on 3D biplane transesophageal echocardiography

**Video 12** Disk-and-lobe device – lobe deployment visualized on 3D biplane transesophageal echocardiography

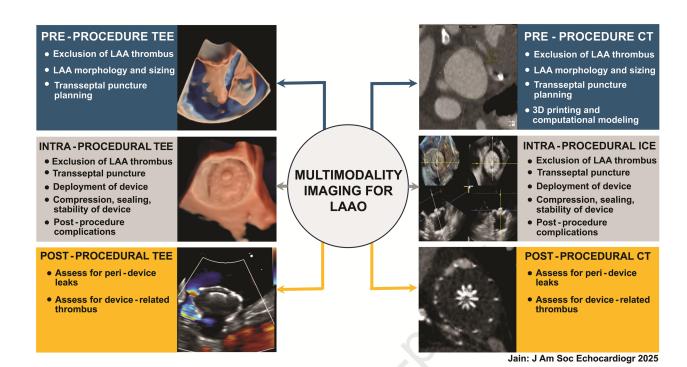
**Video 13** Disk-and-lobe device – disk release is visualized on 3D biplane transesophageal echocardiography

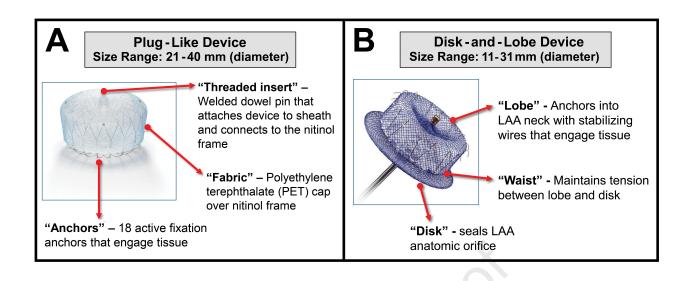
**Video 14** Disk-and-lobe device – tension test performed on 3D biplane transesophageal echocardiography

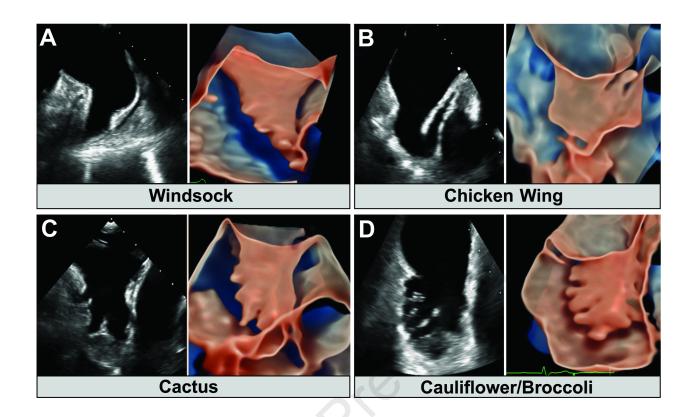
**Video 15** Disk-and-lobe device – deployed device assessed with color Doppler to evaluate for peri-device leak on 2D transesophageal echocardiography

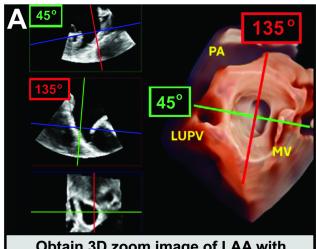
**Video 16** Disk-and-lobe device – device release visualized on 3D biplane transesophageal echocardiography

**Video 17** Disk-and-lobe device – disk released and seen on 3D transesophageal echocardiography

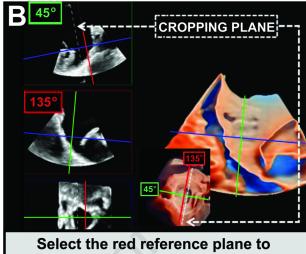




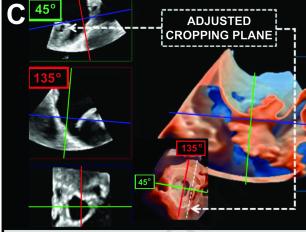




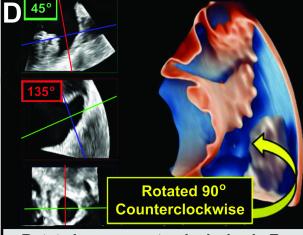
Obtain 3D zoom image of LAA with transillumination in multiplane mode



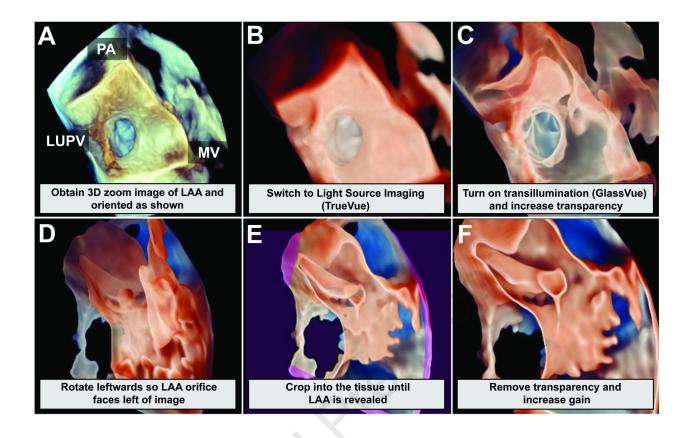
reveal cropped LAA image



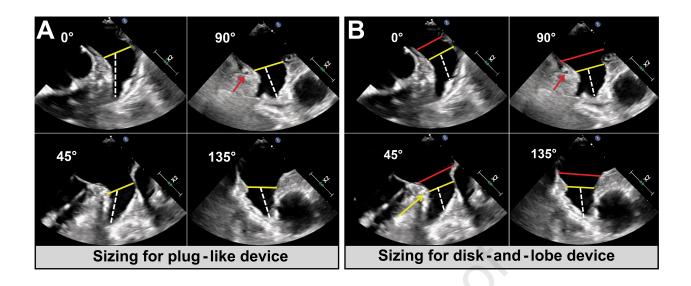
Move cropping plane to uncrop and reveal the entire LAA shape

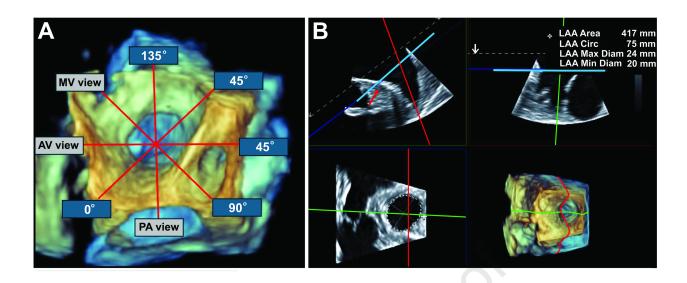


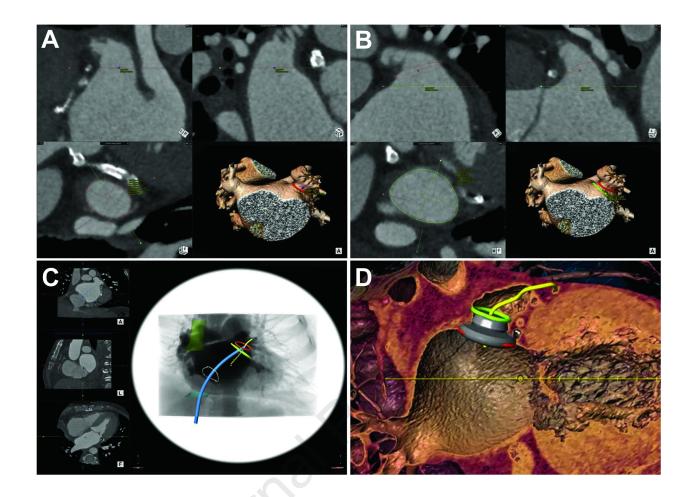
Rotate image counterclockwise in Z plane to emulated RAO CAUDAL view



#### Journal Pre-proof

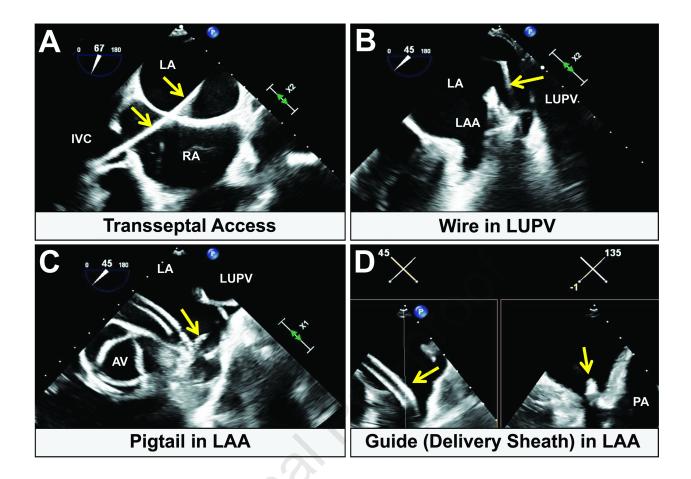


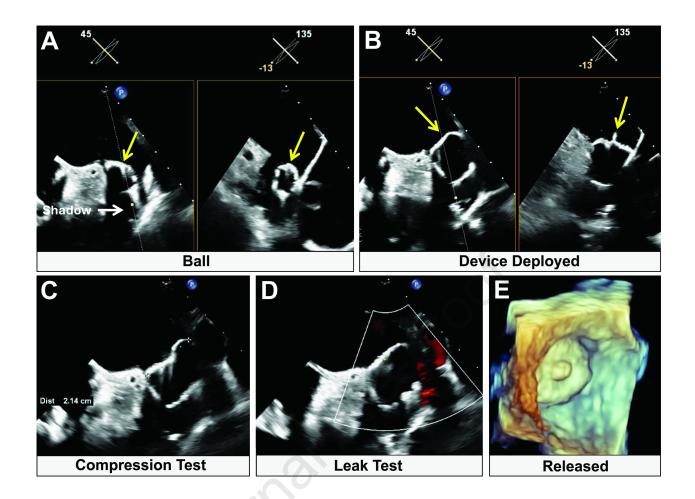


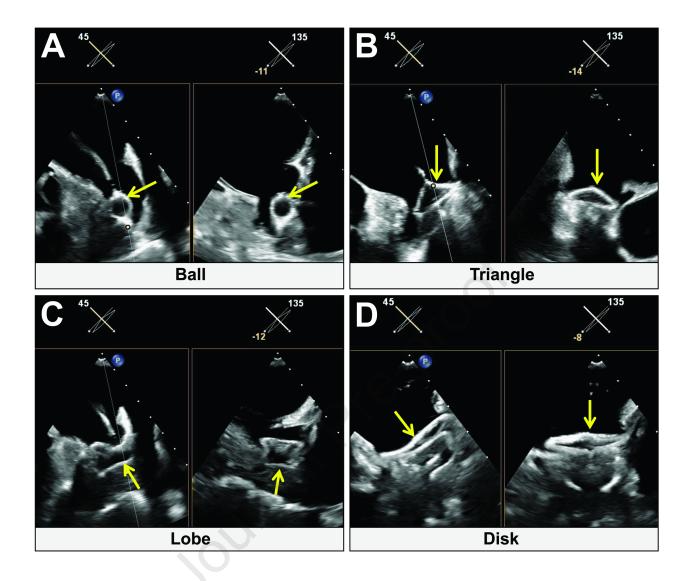


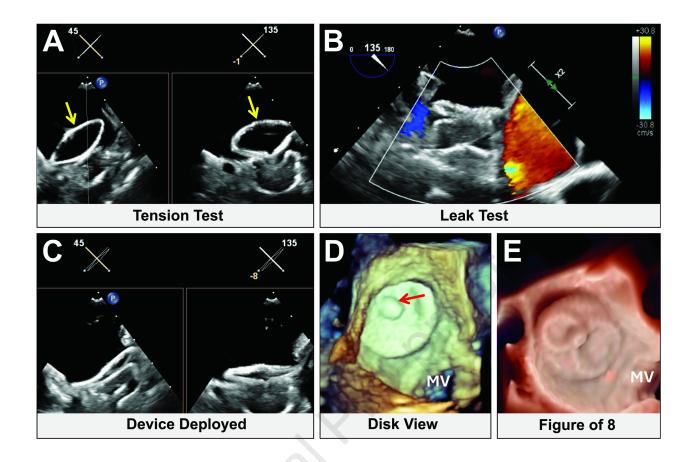






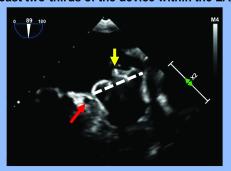






### Position (P)

Device shoulder (white dotted line) at the LAA ostium, or at least two-thirds of the device within the LAA



### Anchor/Stability (A)

Tug test for stability performed by retracting the deployment knob and letting go, to assess return to original position

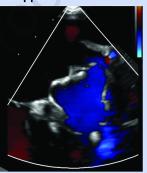
### Size/Compression (S)

10-30% of the original size compression for appropriate stability



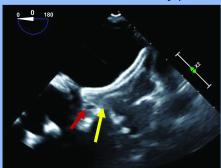
### Seal (S)

Ensure all lobes are distal to device and sealed with ≤5 mm PDL on Doppler



## Circumflex (C)

At least two-thirds of the device lobe (yellow arrow) should be distal to the circumflex artery (red arrow)



# Orientation (O) Lobe should be coaxial with the landing zone

## Separation (S)

The disk (green arrow) must be separated from the lobe (yellow arrow)



### Lobe position (L)

Device lobe should be slightly compressed (tireshaped) and have good apposition to the LAA wall

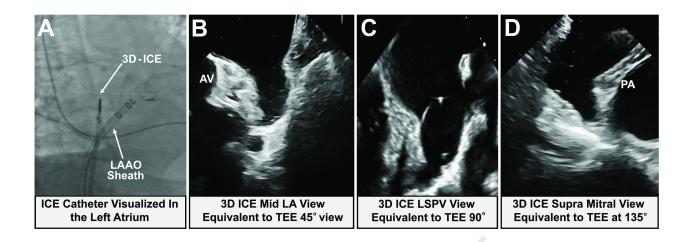


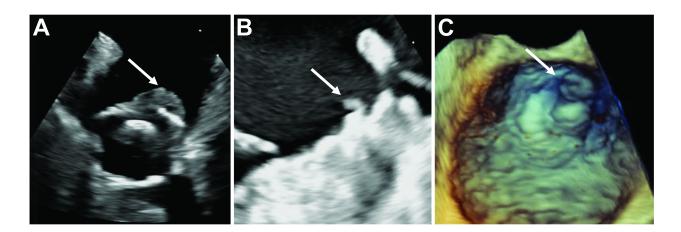
### Elliptical (E)

The disk (green arrow) should be a concave shape with respect to the left atrium



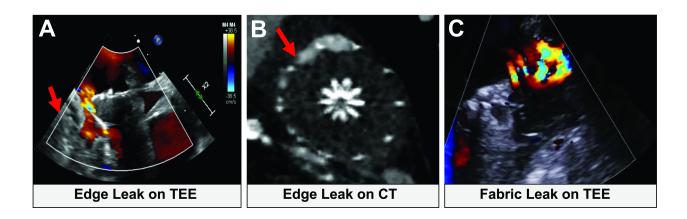
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### **Highlights**

- Advanced imaging technologies and techniques have revolutionized LAA occlusion.
- 3D TEE and multiplanar reconstruction aid device sizing and procedural planning.
- MDCT allows 3D printing and virtual device simulation useful during planning.
- 3D ICE can guide transseptal puncture and device deployment.
- Post-procedure, TEE and MDCT can assess device stability and complications.