



# Comparison of the effective orifice area of prosthetic mitral valves using two-dimensional versus three-dimensional transesophageal echocardiography

Lei Zhou<sup>1</sup>, Hai-yan Wei<sup>2</sup>, Ya-li Ge<sup>2</sup>,  
Zheng-nian Ding<sup>3,\*</sup>  and Hong-wei Shi<sup>4,\*</sup>

## Abstract

**Objective:** This study compared the continuity equation-based effective orifice area (EOA) of prosthetic mitral valves between two-dimensional (2D) and 3D transesophageal echocardiography (TEE).

**Methods:** Thirty-four patients without major aortic valve abnormalities underwent mitral valve replacement surgery. The EOAs of prosthetic mitral valves were calculated using the continuity equation with 2D and 3D TEE. For 18/34 patients using a biological valve prosthesis, the EOA of the prosthesis was obtained from commercial records.

**Results:** The EOA of prosthetic mitral valves significantly varied between the 2D and 3D methods ( $2.22 \pm 0.71$  vs  $2.35 \pm 0.70$  cm<sup>2</sup>,  $n = 34$ ). The area of the diameter of the left ventricular outflow tract as determined by the 3D method was significantly higher than that by the 2D method (mean difference:  $-0.14 \pm 0.20$  cm<sup>2</sup>), with 95% coherence boundaries of  $-0.53$  and  $0.25$  cm<sup>2</sup>. The regression equation for the EOA by 3D and 2D TEE was  $y = 0.27 + 0.94x$ , with a good correlation.

<sup>4</sup>Department of Anesthesiology, Nanjing First Hospital & Nanjing Cardiovascular Disease Hospital, Nanjing Medical University, Nanjing, Jiangsu, China

\*These authors contributed equally to this work.

## Corresponding author:

Hong-wei Shi, Department of Anesthesiology, Nanjing First Hospital & Nanjing Cardiovascular Disease Hospital, Nanjing Medical University, 68 Changle Road, Qinhuai District, Nanjing, Jiangsu 210006, China.  
Email: mdshw@163.com

<sup>1</sup>Department of Anesthesiology, Changzhou Maternity and Child Health Care Hospital Affiliated to Nanjing Medical University, Changzhou, Jiangsu, China

<sup>2</sup>Department of Anesthesiology, Nanjing First Hospital, Nanjing Medical University, Nanjing, Jiangsu, China

<sup>3</sup>Department of Anesthesiology, Jiangsu People's Hospital, the First Affiliated Hospital of Nanjing Medical University, Nanjing, Jiangsu, China



**Conclusions:** The EOA of prosthetic mitral valves is underestimated using the 2D TEE method compared with the 3D TEE method. The 3D-TEE method has the advantage of higher precision over the 2D TEE method, and it may be helpful for better assessment of prosthetic mitral valves intraoperatively.

### Keywords

Prosthetic mitral valve, transesophageal echocardiography, three-dimensional, ventricular outflow tract, effective orifice area, stroke volume

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## Introduction

Estimation of the effective orifice area (EOA) of a prosthesis is challenging, especially when the prosthesis is damaged. The continuity equation is frequently used to assist such judgment. In the clinical situation and experimental studies, calculations of the continuity equation valve area have been conclusively validated.<sup>1,2</sup> Furthermore, calculating the continuity equation valve area is considered to be a credible parameter when making predictions of clinical outcome and clinical decisions.

According to the principle of continuous equations, in the absence of valvular reflux and intra-cardiac shunts, the blood flow rate passing through the normal aortic valve and that through the mitral valve should be equal.<sup>2</sup> Measurement of the diameter of the left ventricular outflow tract (LVOT) needs to be accurate because it is essential for calculation of the continuity equation. This equation is calculated according to the assumption that the shape of the LVOT is circular. Therefore, a single diameter can be used to obtain a precise assessment of the LVOT area. However, the LVOT is currently recognized as not circular, but elliptical, with major and minor axes in a large percentage of

patients.<sup>3-5</sup> The cross-sectional area of the left ventricular outflow tract ( $CSA_{LVOT}$ ) originating from a two-dimensional (2D)-based diameter of the LVOT may result in different outcomes of the EOA of a prosthetic mitral valve with calculation of the continuity equation compared with the 3D-planimetered LVOT area with real-time 3D transesophageal echocardiography (TEE).

We conducted this study to measure and compare the EOA of prosthetic mitral valves between two techniques. This area was calculated with the  $CSA_{LVOT}$  originating from a 2D-based diameter of the LVOT and with a 3D-planimetered LVOT area using real-time 3D TEE in patients undergoing a mitral valve replacement operation. This study aimed to determine a more accurate and reliable method for determining the EOA of a prosthesis.

## Materials and methods

The study was part of a current protocol for collection of intraoperative echocardiographic data. This study was approved by the ethics committee of Nanjing First Hospital, Nanjing Medical University (approval number: KY20170811-03). All patients were informed that they would participate in the study and provided written

consent before the operation. They were also informed before the operation that they would receive a TEE examination, and informed of the relevant risks and precautions. We studied 2D and 3D data that were routinely collected from patients who underwent elective mitral valve replacement surgery in Nanjing First Hospital, Nanjing Medical University.

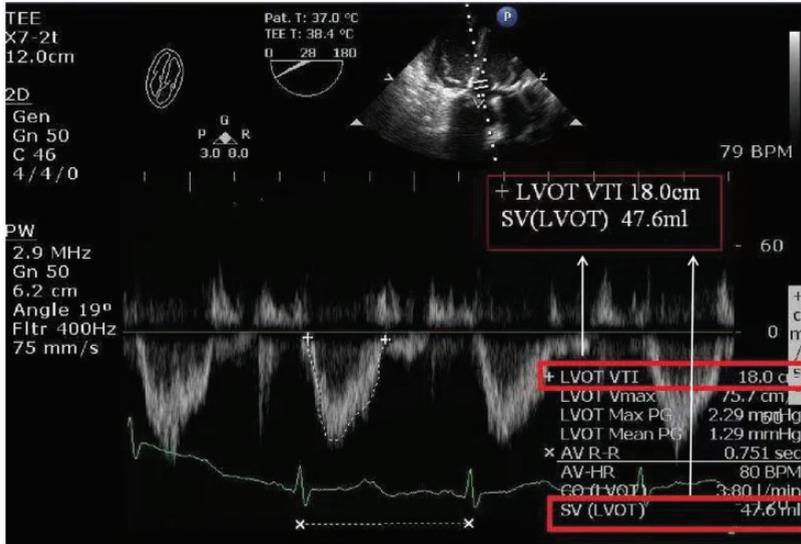
We collected echocardiographic data from patients who received mitral valve replacement surgery with intraoperative 3D TEE during March 2019 to August 2019. We excluded patients who had undergone emergency procedures, those with previous valvular heart surgery, those who had more than moderate regurgitation of the aortic valve, those with moderate regurgitation of the mitral valve after surgery, those with combined procedures (e.g., ascending aortic surgery or aortic valve replacement), and those with suboptimal 3D images from this study. Two more patients were excluded because of suboptimal 3D images.

All operations were performed by two senior attending physicians with more than 5 years of experience in TEE examinations and a TEE advanced qualification certification. The examination methods and programs were consistent. The 3D images were transferred to a computer from the ultrasonic equipment after the operation and an investigator analyzed acquired 3D data. The intraoperative values were blinded for the investigator. For the archived 3D data, the investigator measured the results again 1 week later, and the results were averaged with the first measurement.

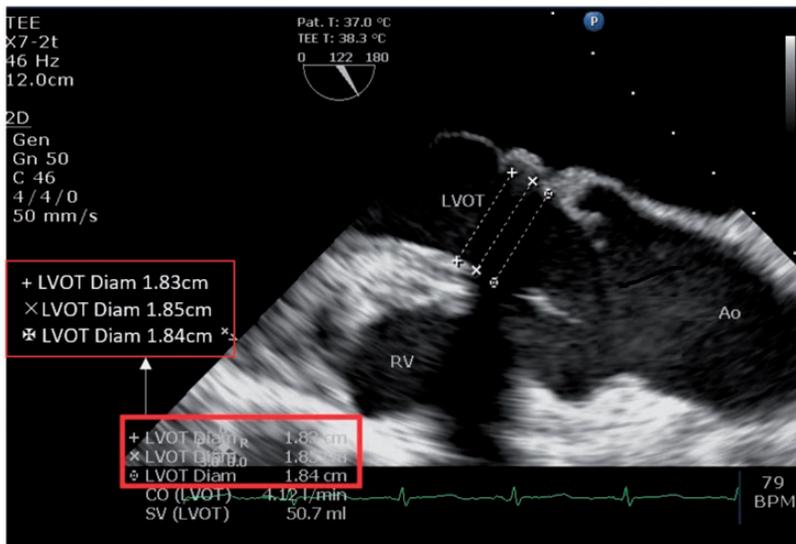
Using a Philips CX50 ultrasound system (Philips Healthcare, Andover, MA, USA) and its X7-2t probe, we conducted intraoperative TEE examinations after closing the chest and before transferring the patient. According to the European and American Echocardiographic Societies Recommendations for Image Acquisition

and Display Using Three-dimensional Echocardiography, we selected the 2D mode at a frame rate above 45 HZ and the 3D mode at a frame rate of 15 to 25 HZ. Comprehensive 2D exams were performed according to these guidelines. Stroke volume (SV) was calculated as  $CSA_{LVOT} \times \text{velocity time integral (VTI)}$  through the LVOT ( $VTI_{LVOT}$ ).  $CSA_{LVOT}$  was calculated using 2D measurement results and data derived from 3D images when using the 3D method. The EOA of the prosthetic mitral valve was calculated with the continuity equation as follows:  $EOA = SV / VTI_{pmv}$ , where  $VTI_{pmv}$  is velocity-time through the mitral valve prosthesis. Therefore,  $EOA = CSA_{LVOT} \times VTI_{LVOT} / VTI_{pmv}$ . For calculation of the EOA, we obtained  $VTI_{pmv}$ , as measured by continuous-wave Doppler below the mitral valve prosthesis. We also collected the  $VTI_{LVOT}$  using pulse-wave Doppler in the deep transgastric window (Figure 1). The same values of  $VTI_{LVOT}$  and  $VTI_{pmv}$  were used in the 2D and 3D methods.

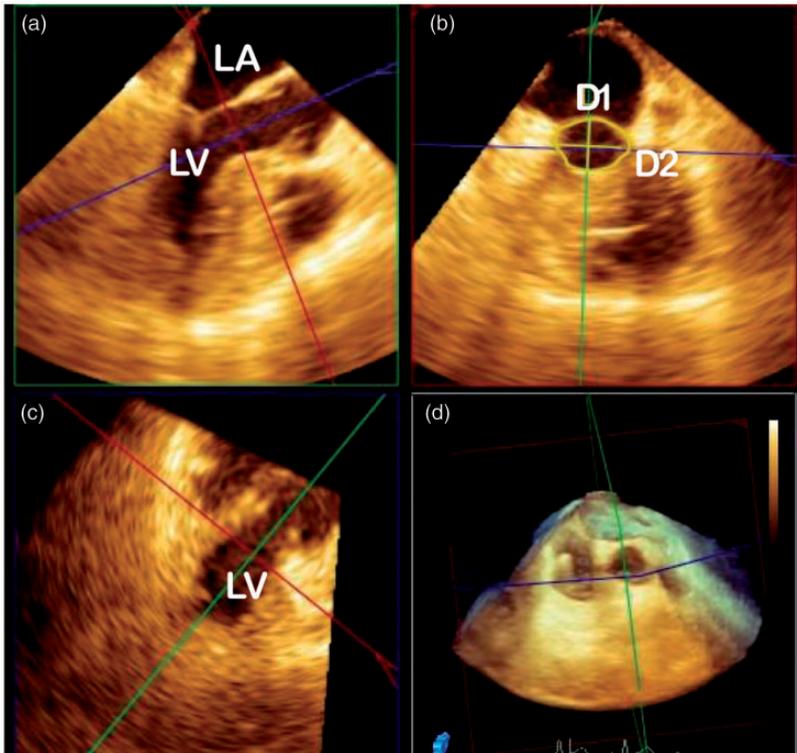
In the 2D method, from the mid-esophageal position, we obtained LVOT images in the long-axis view ( $120^\circ$ – $150^\circ$ ). Using the zoom function, we measured the LVOT diameter approximately 5 mm from insertion of the aortic leaflets in the mid-systolic phase (Figure 2). The  $CSA_{LVOT}$  was calculated as  $\pi \times (LVOT \text{ diameter} / 2)^2$  and was automatically derived by the instrument's software. In the 3D method, measurement and 2D TEE were performed simultaneously with the same system. We acquired 3D data of the LVOT and aortic valve in the center of the screen. When we aligned the multi-planar reformatting planes, including the sagittal, coronal, and transverse planes, in the mid-systolic position (Figure 3), we displayed three geometrically orthogonal views of the LVOT and the aortic valve. To achieve the highest spatial and temporal resolution, we avoided absence of motion or electrical interference.



**Figure 1.** The + LVOT VTI in the deep transgastric window was obtained. SV was calculated as + LVOT VTI $\times$ CSA<sub>LVOT</sub>. + LVOT VTI, velocity–time integral through the left ventricular outflow tract; SV, stroke volume; LVOT, left ventricular outflow tract; CSA<sub>LVOT</sub>, cross-sectional area through the left ventricular outflow tract.



**Figure 2.** The LVOT diameter was obtained from LVOT images in the long-axis view. In this case, we took measurements three times with +LVOT Diam, xLVOT Diam, and ■ LVOT Diam, and used the average value of these results. LVOT, left ventricular outflow tract; CO, cardiac output; SV, stroke volume; RV, right ventricle; Ao, aorta.



**Figure 3.** Alignment of orthogonal multi-planar reformatting planes to each other. An accurate en face view of the left ventricular outflow tract was obtained from three-dimensional images. An elliptical shape of the cross-sectional area through the left ventricular outflow tract was displayed. LA, left atrium; LV, left ventricle; D1, minor axis diameter; D2, major axis diameter.

After surgery, the 3D data were transferred to a computer with Q-Lab Version 8.1.2 Advanced Ultrasound Quantification Software (Philips Healthcare). Using this software, an investigator performed geometrical reestablishment and analyzed the LVOT. The shape of the 3D-planimetered LVOT area ( $CSA_{\text{planimetered}}$ ) was traced and the result was obtained from the software mentioned above. SV was calculated as  $CSA_{\text{planimetered}} \times VTI_{\text{LVOT}}$ .

The EOAs of 18 patients using a biological valve prosthesis were obtained from commercial records (R-EOAs). We did not find EOA data from commercial records for 16 patients with mechanical valves. The indexed EOA was defined as the EOA of

a prosthetic mitral valve divided by the patients' body surface area. We defined prosthesis-patient mismatch (PPM) as an indexed EOA  $\leq 1.20 \text{ cm}^2/\text{m}^2$ , with severe PPM defined as an indexed EOA  $< 0.9 \text{ cm}^2/\text{m}^2$ .<sup>6</sup>

Continuous data are expressed as mean  $\pm$  standard deviation. Categorical data are expressed as the number and percentage. For comparison of two continuous variables, the two-sided paired t test or Mann-Whitney U test was used, with or without a normal distribution. The relationship of EOAs was assessed by applying Pearson's correlation coefficient analysis and simple regression analysis. We also used Bland-Altman analysis to define the biases and

limits of agreement between the two methods. All of the data were analyzed using IBM SPSS version 20.0 (IBM Corp, Armonk, NY, USA).

## Results

We examined 34 patients and their baseline clinicopathological characteristics are shown in Table 1. The mean age was  $57.26 \pm 9.87$  years, and 34% ( $n=12$ ) were men and 66% ( $n=22$ ) were women.

The mean EOA of the prosthetic mitral valve as estimated by the 3D method was significantly larger than that estimated by the 2D method ( $2.35 \pm 0.70$  vs  $2.22 \pm 0.71$  cm<sup>2</sup>,  $P < 0.001$ ). The  $CSA_{LVOT}$  as estimated by the 3D method was also significantly larger than that obtained by the 2D method ( $3.12 \pm 0.83$  vs  $2.95 \pm 0.87$  cm<sup>2</sup>,  $P < 0.001$ ). SV as estimated by the 3D method was significantly higher than that estimated by the 2D method ( $52.39 \pm 3.42$  vs  $49.34 \pm 3.36$  cm<sup>2</sup>,  $P < 0.01$ ).

Bland–Altman analysis (Figure 4) was used to test the coherence of the EOA calculated by the two techniques. We found

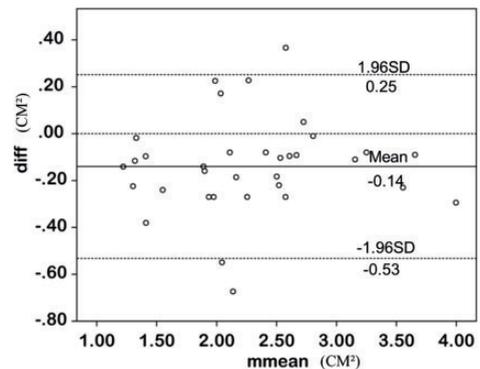
**Table 1.** Characteristics of the patients ( $n=34$ ).

|   |                   |
|---|-------------------|
| Age (years)                               | $57.26 \pm 9.87$  |
| Sex                                       |                   |
| Men                                       | 12 (34)           |
| Women                                     | 22 (66)           |
| Body surface area (m <sup>2</sup> )       | $1.77 \pm 0.15$   |
| Body weight (kg)                          | $64.06 \pm 10.16$ |
| Mean arterial pressure (mmHg)             | $80.42 \pm 9.49$  |
| Central venous pressure (mmHg)            | $8.72 \pm 2.19$   |
| Aortic occlusion time (minutes)           | $69.00 \pm 13.34$ |
| Extracorporeal circulation time (minutes) | $93.42 \pm 17.97$ |
| Temporary pacemaker                       | 20 (59)           |
| Mechanical prosthesis                     | 16 (47)           |
| Biological prosthesis                     | 18 (53)           |

Values are mean  $\pm$  standard deviation or n (%).

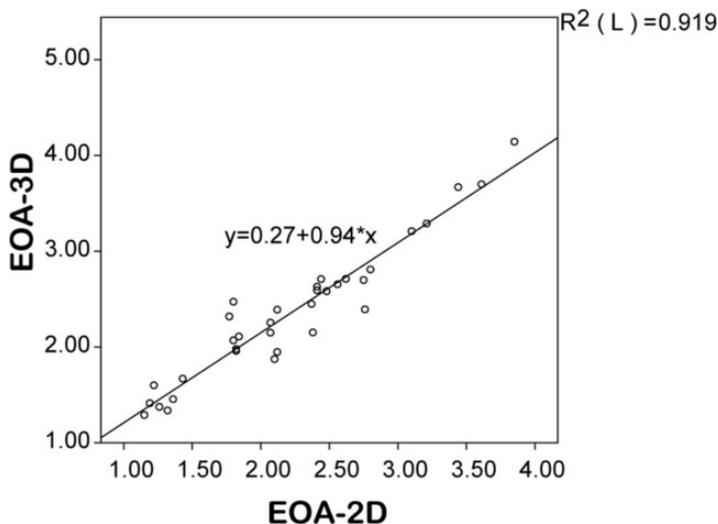
good agreement of the EOA calculated by the two methods, which showed a fixed bias (mean difference was  $0.14 \pm 0.20$  cm<sup>2</sup>,  $P < 0.001$ ), and the 95% coherence boundaries were  $-0.53$  and  $0.25$  cm<sup>2</sup>. We also found a good correlation of the EOA of the mitral valve between the 2D and 3D methods ( $r=0.94$ ,  $P < 0.001$ ) (Figure 5). The observed mean difference in the EOA between the 2D method and R-EOA was  $0.37$  cm<sup>2</sup> ( $2.22 \pm 0.71$  vs  $2.59 \pm 0.37$  cm<sup>2</sup>). The observed mean difference in the EOA between the 3D method and R-EOA was  $0.24$  cm<sup>2</sup> ( $2.35 \pm 0.70$  vs  $2.59 \pm 0.37$  cm<sup>2</sup>). These mean differences between the 2D and 3D methods and R-EOA were not significant.

In the 2D method, mitral PPM was found in 32.3% (11/34) of the patients. Four of the 11 patients with mitral PPM had severe PPM. Using the 3D method, mitral PPM was found in 26.4% (9/34) of patients. Three of nine patients with mitral PPM had severe PPM. The probability of occurrence of mitral PPM was significantly



**Figure 4.** Comparison of the effective orifice areas of the two methods with Bland–Altman analysis.

SD, standard deviation; diff, difference; mmean, mean effective orifice area by two-dimensional transesophageal echocardiography and effective orifice area by three-dimensional transesophageal echocardiography (for the same patient measured at the same time).



**Figure 5.** Correlation of the EOA of the mitral valve between 2D and 3D measurements ( $R^2 = 0.919$ ,  $P < 0.001$ ).

EOA, effective orifice area; 2D, two-dimensional; 3D, three-dimensional.

higher using the 2D method than the 3D method ( $P < 0.05$ ).

## Discussion

The current study suggested that use of the continuity equation to calculate the EOA of prosthetic mitral valves may have been affected by estimation of the  $CSA_{LVOT}$  with 2D echocardiography. The 2D method is based on the diameter of a single axis. However, the LVOT is not circular, but elliptical instead, with major and minor axes in a large percentage of patients.<sup>3–5</sup> Therefore, the  $CSA_{LVOT}$  with 2D echocardiography may be either under- or overestimated depending on which single diameter (i.e., minor or major axis<sup>7</sup>) is used. With calculation of the continuity equation, a difference in SV leads to a difference in calculation of the EOA of the mitral valve.<sup>8,9</sup>

In our study, calculation using the 2D-based  $CSA_{LVOT}$  of SV led to underestimation of SV by approximately 6% compared with the 3D-planimetered LVOT area. This

finding is in agreement with previous reports on the effect of the elliptical shape of the LVOT by observation and calculation of the aortic valve area.<sup>10–14</sup> Our study showed a slightly smaller underestimation of SV than that reported previously (10%).<sup>15</sup> The reason for this difference between studies may be due to differences in different types of surgery and different time points of observation. Two-dimensional-derived calculations may have limitations compared with 3D imaging.<sup>16,17</sup> In the mid-esophageal long-axis view of 2D, there may be a good opportunity to visualize the minor axis of the LVOT. Therefore, the area may be underestimated using this measurement. However, in the 3D method, acquisition of an en face view of the LVOT might be more precise because it may exclude the assumption of circularity.<sup>18</sup>

We recorded the R-EOA of 18 patients using bioprosthetic valves from commercial records. We found that the difference between EOA values measured by the 3D method and R-EOA was smaller than that between EOA values measured by the 2D

method and R-EOA. Bioprosthetic valves are biologically closer to human valves compared with mechanical valves, and the valid valve area recorded by commercial products is easier to obtain. However, after implantation of a valve, the EOA of the valve is also affected by other factors. The R-EOA can only be used as a reference.<sup>9</sup>

The shape of the LVOT may be slightly distorted after mitral valve replacement because of the presence of a prosthesis. Additionally, continuing to use the 2D method to measure the  $CSA_{LVOT}$  after implantation of a prosthesis may lead to inaccurate results. However, when the  $CSA_{LVOT}$  is measured by the 3D method, the resulting error caused by distortion can be corrected because the shape can be adjusted by tracing according to the actual situation. This situation also requires attention especially after mitral valve replacement surgery.

Mitral PPM may adversely affect long-term survival of patients with regurgitation of the mitral valve.<sup>18</sup> We found that the incidence rate of mitral PPM using the 3D method was lower than that using the 2D method. Therefore, the 3D method may be useful in early identification of PPM perioperatively.

A limitation of our study is that a relatively small number of patients were included. A further study on continuity equation-based estimation of the EOA of prosthetic mitral valves using 2D and 3D TEE with a larger population is required in the future. Additionally, SV as measured by pulmonary artery catheterization may need to be added for providing vital information in future studies. We performed analysis of data using software in a computer and routine analysis of 3D data of geometrically orthogonal views (sagittal, coronal, and transverse planes). However, because of resource constraints, this analysis cannot be achieved on a routine basis.

Calcification of the elliptical shape of the LVOT also needs to be further investigated. However, the possibility that the LVOT area as measured with the 2D method may be more reliable in certain situations cannot be excluded.<sup>19,20</sup> Additionally, because of the low frame rate in early studies of 3D ultrasound, frame loss may have severely affected the effectiveness of 3D results. However, in our study, the 3D mode at a frame rate of 15 to 25 HZ may have overcome this weakness of frame loss. The 3D mode also has a limitation that if 3D data in multiple cardiac cycles need to be collected, the present frame rate is insufficient.

## Conclusions

When the continuity equation is used, the EOA of a prosthetic mitral valve may be underestimated using the 2D TEE method based on the diameter of a single axis compared with the 3D-planimetered area. The advantage of the 3D method versus the 2D method is higher precision, which leads to better assessment of a prosthetic mitral valve intraoperatively.

## Declaration of conflicting interest

The authors declare that there is no conflict of interest.

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## ORCID iD

Zheng-nian Ding  <https://orcid.org/0000-0002-9813-6664>

## References

1. Alame AJ, Karatasakis A, Karacsonyi J, et al. Comparison of the American College of Cardiology/American Heart Association

- and the European Society of Cardiology Guidelines for the Management of Patients With Valvular Heart Disease. *J Invasive Cardiol* 2017; 29: 320–326.
2. Nishimura RA, Otto CM, Bonow RO, et al. 2017 AHA/ACC Focused Update of the 2014 AHA/ACC Guideline for the Management of Patients With Valvular Heart Disease: A Report of the American College of Cardiology/American Heart Association Task Force on Clinical Practice Guidelines. *J Am Coll Cardiol* 2017; 70: 252–289.
  3. Husser O, Rauch S, Endemann DH, et al. Impact of three-dimensional transesophageal echocardiography on prosthesis sizing for transcatheter aortic valve implantation. *Catheter Cardiovasc Interv* 2012; 80: 956–963.
  4. Doddamani S, Bello R, Friedman MA, et al. Demonstration of left ventricular outflow tract eccentricity by real time 3D echocardiography: implications for the determination of aortic valve area. *Echocardiography* 2007; 24: 860–866.
  5. Doddamani S, Grushko MJ, Makaryus AN, et al. Demonstration of left ventricular outflow tract eccentricity by 64-slice multi-detector CT. *Int J Cardiovasc Imaging* 2009; 25: 175–181.
  6. Hwang HY, Kim YH, Kim KH, et al. Patient-prosthesis mismatch after mitral valve replacement: a propensity score analysis. *Ann Thorac Surg* 2016; 101: 1796–1802.
  7. Wu VC, Kaku K, Takeuchi M, et al. Aortic root geometry in patients with aortic stenosis assessed by real-time three-dimensional transesophageal echocardiography. *J Am Soc Echocardiogr* 2014; 27: 32–41.
  8. Singh M, Sporn ZA, Schaff HV, et al. ACC/AHA Versus ESC Guidelines on Prosthetic Heart Valve Management: JACC Guideline Comparison. *J Am Coll Cardiol* 2019; 73: 1707–1718.
  9. Zamorano JL, Badano LP, Bruce C, et al. EAE/ASE recommendations for the use of echocardiography in new transcatheter interventions for valvular heart disease. *Eur Heart J* 2011; 32: 2189–2214.
  10. Tidholm A, Bodegård-Westling A, Höglund K, et al. Real-Time 3-Dimensional Echocardiographic Assessment of Effective Regurgitant Orifice Area in Dogs With Myxomatous Mitral Valve Disease. *J Vet Intern Med* 2017; 31: 303–310.
  11. Tan TC, Zeng X, Jiao Y, et al. Three-Dimensional Field Optimization Method: Clinical Validation of a Novel Color Doppler Method for Quantifying Mitral Regurgitation. *J Am Soc Echocardiogr* 2016; 29: 926–934.
  12. Bayramoğlu A, Taşolar H, Otlu YÖ, et al. Assessment of left atrial volume and mechanical functions using real-time three-dimensional echocardiography in patients with mitral annular calcification. *Anatol J Cardiol* 2016; 16: 42–47.
  13. Elsayed M, Thind M and Nanda NC. Two- and Three-dimensional Transthoracic Echocardiographic Assessment of Tricuspid Valve Prolapse with Mid-to-Late Systolic Tricuspid Regurgitation. *Echocardiography* 2015; 32: 1022–1025.
  14. Koto D, Izumo M, Machida T, et al. Geometry of the left ventricular outflow tract assessed by 3D TEE in patients with aortic stenosis: impact of upper septal hypertrophy on measurements of Doppler-derived left ventricular stroke volume. *J Echocardiogr* 2018; 16: 162–172.
  15. Montealegre-Gallegos M, Mahmood F, Owais K, et al. Cardiac output calculation and three-dimensional echocardiography. *J Cardiothorac Vasc Anesth* 2014; 28: 547–550.
  16. Santosa CM, Rose DD and Fleming NW. The Utility of 3D Left Atrial Volume and Mitral Flow Velocities as Guides for Acute Volume Resuscitation. *Biomed Res Int* 2015; 2015: 697327.
  17. Wunderlich NC, Beigel R and Siegel RJ. Management of mitral stenosis using 2D and 3D echo-Doppler imaging. *JACC Cardiovasc Imaging* 2013; 6: 1191–1205.
  18. Tsubota H, Sakaguchi G, Arakaki R, et al. Impact of prosthesis-patient mismatch after mitral valve replacement: A propensity score analysis. *Semin Thorac Cardiovasc Surg* 2020; S1043-0679(20)30299-9. DOI: 10.1053/j.semtcvs.2020.09.026
  19. Kwon SH and Gopal AS. 3D and 4D Ultrasound: Current Progress and Future

- Perspectives. *Curr Cardiovasc Imaging Rep* 2017; 10: 43.
20. Papachristidis A, Papitsas M, Roper D, et al. Three-Dimensional Measurement of Aortic Annulus Dimensions Using Area or Circumference for Transcatheter Aortic Valve Replacement Valve Sizing: Does It Make a Difference. *J Am Soc Echocardiogr* 2017; 30: 871–878.