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Assessment of paravalvular regurgitation after transcatheter aortic valve replacement using 2D multi-velocity encoding and 4D flow cardiac magnetic resonance

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Aims

To compare the novel 2D multi-velocity encoding (venc) and 4D flow acquisitions with the standard 2D flow acquisition for the assessment of paravalvular regurgitation (PVR) after transcatheter aortic valve replacement (TAVR) using cardiac magnetic resonance (CMR)-derived regurgitant fraction (RF).

Methods and results

In this prospective study, patients underwent CMR 1 month after TAVR for the assessment of PVR, for which 2D multi-venc and 4D flow were used, in addition to standard 2D flow. Scatterplots and Bland–Altman plots were used to assess correlation and visualize agreement between techniques. Reproducibility of measurements was assessed with intraclass correlation coefficients. The study included 21 patients (mean age \pm SD 80 ± 5 years, 9 men). The mean RF was $11.7 \pm 10.0\%$ when standard 2D flow was used, $10.6 \pm 7.0\%$ when 2D multi-venc flow was used, and $9.6 \pm 7.3\%$ when 4D flow was used. There was a very strong correlation between the RFs assessed with 2D multi-venc and standard 2D flow ($r = 0.88$, $P < 0.001$), and a strong correlation between the RFs assessed with 4D flow and standard 2D flow ($r = 0.74$, $P < 0.001$). Bland–Altman plots revealed no substantial bias between the RFs (2D multi-venc: 1.3%; 4D flow: 0.3%). Intra-observer and inter-observer reproducibility for 2D multi-venc flow were 0.98 and 0.97, respectively, and 0.92 and 0.90 for 4D flow, respectively.

Conclusion

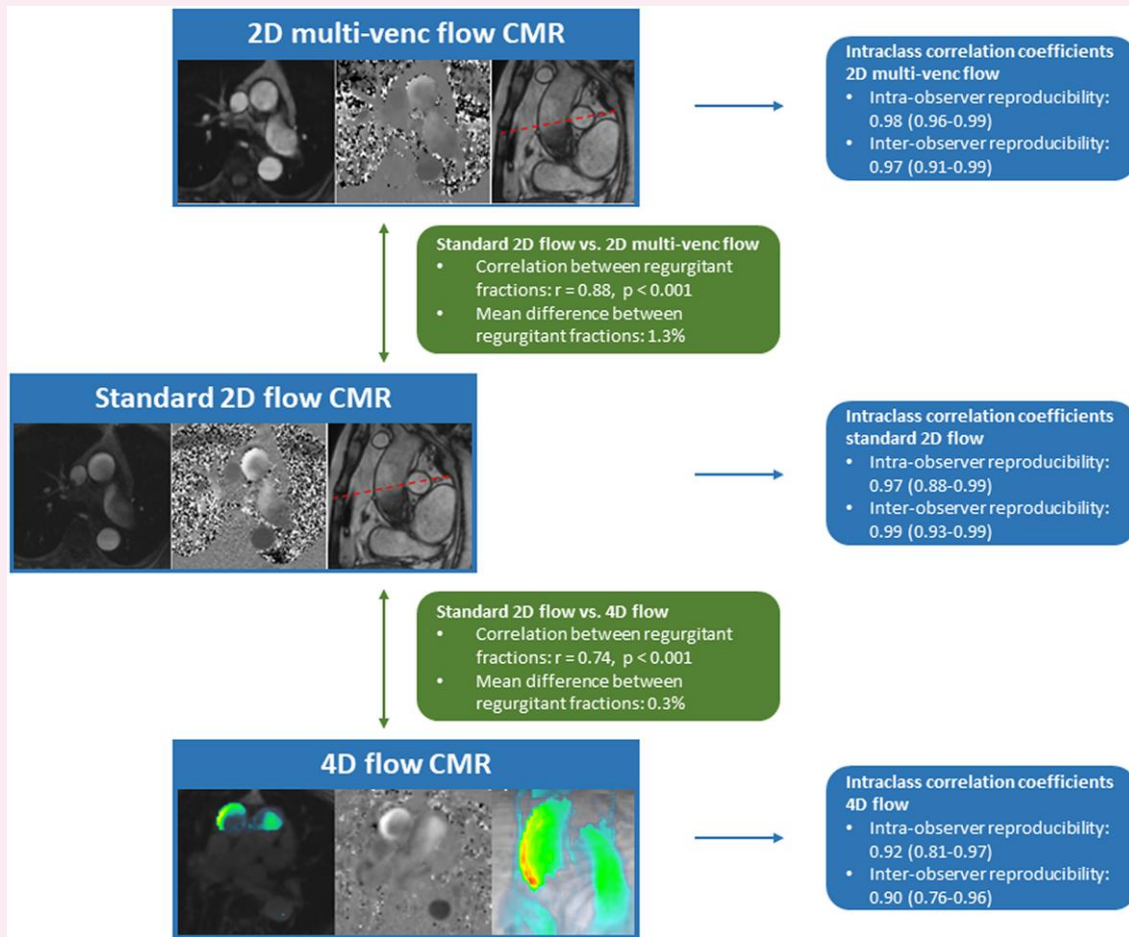
Two-dimensional multi-venc and 4D flow produce an accurate quantification of PVR after TAVR. The fast acquisition of the 2D multi-venc sequence and the free-breathing acquisition with retrospective plane selection of the 4D flow sequence provide useful advantages in clinical practice, especially in the frail TAVR population.

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Graphical Abstract



Assessment of paravalvular regurgitation after transcatheter aortic valve replacement using novel CMR flow assessment techniques. CMR, cardiac magnetic resonance; venv, velocity encoding.

Keywords aortic stenosis • cardiac magnetic resonance • paravalvular regurgitation • transcatheter aortic valve replacement

Introduction

Aortic stenosis (AS) is the most common valvular heart disease in the Western world. Randomized trials have demonstrated non-inferiority or superiority of transcatheter aortic valve replacement (TAVR) compared with surgical aortic valve replacement in patients with severe AS across the spectrum of surgical risk.¹⁻⁶ An important shortcoming of TAVR is the relatively high risk for the development of paravalvular regurgitation (PVR). The occurrence of PVR is common after TAVR, with incidences of mild PVR up to 40% and incidences of \geq moderate PVR up to 10% in contemporary TAVR studies.^{7,8} In patients with \geq moderate PVR, mortality is three times higher compared with patients with none to trace PVR.⁹⁻¹¹ Recently, increasing evidence suggests that mild-to-moderate or even only mild PVR could also have an impact on mortality.^{12,13} Given the strong implications for post-TAVR survival, early identification of relevant PVR is warranted to guide additional interventions (e.g. post-dilation) in order to reduce PVR and improve patient outcomes.

Although echocardiography remains the cornerstone of valvular heart disease assessment, visualization, and specifically quantification of aortic regurgitation after TAVR are challenging. Quantitative flow assessment using

2D through-plane phase-contrast (PC) cardiac magnetic resonance (CMR) has less inter-observer variability than echocardiography,¹⁴ is not limited by its acoustic windows secondary to patient characteristics (e.g. obesity and chronic obstructive pulmonary disease [COPD]), and allows unlimited imaging plane selection. Emerging techniques in the field of flow assessment after TAVR using CMR are 2D multi-velocity encoding (venv) flow mapping and 4D flow mapping. Two-dimensional multi-venv flow mapping facilitates the use of a single breath-hold to analyse two or three different venv values by combining these individual venv values into a single reconstruction that can be used for flow quantification.¹⁵ In 4D flow mapping, 3D blood flow patterns and haemodynamics can be assessed along all three spatial dimensions and over the complete cardiac cycle.¹⁶ Four-dimensional flow assessment is not dependent on breath-holds, thus allowing the patient to breathe freely during acquisition. As a result, no breathing-dependent variation is observed. Moreover, since the analysis plane can be set anywhere within the acquisition volume offline, 4D flow mapping is less operator dependent.¹⁷

The aim of this substudy of the Assessment of Paravalvular Regurgitation After Transcatheter Aortic Valve Replacement by Haemodynamic Measurements and Cardiac Magnetic Resonance

Table 1 Baseline characteristics

	Study population (N = 21)
Demographics	
Age, years	79.8 ± 4.9
Male sex, n (%)	9 (42.9)
Body mass index, kg/m ²	28.0 ± 3.7
Obesity, n (%)	5 (23.8)
Medical history	
EuroSCORE II	2.62 ± 1.32
NYHA Class III/IV, n (%)	12 (57.1)
Diabetes mellitus, n (%)	7 (33.3)
Coronary artery disease, n (%)	13 (61.9)
COPD, n (%)	3 (14.3)
Atrial fibrillation, n (%)	4 (19.0)
Pre-procedural echocardiographic parameters	
Aortic valve area, cm ²	0.81 ± 0.19
Aortic valve mean gradient, mmHg	45.7 ± 6.4
Aortic valve maximum velocity, m/s	4.3 ± 0.3
LVEF, %	53.3 ± 9.3
Moderate or severe aortic regurgitation, n (%)	2 (9.5)

Data are presented as mean ± standard deviation or as number (%). COPD, chronic obstructive pulmonary disease; EuroSCORE, European System for Cardiac Operative Risk Evaluation; LVEF, left ventricular ejection fraction; NYHA, New York Heart Association.

Results

Baseline characteristics

Baseline characteristics of the 21 patients are presented in Table 1. The mean age was 79.8 ± 4.9 years, and 42.9% of patients were men. The mean European System for Cardiac Operative Risk Evaluation (EuroSCORE) II was 2.62 ± 1.32, with 57.1% of patients being in New York Heart Association (NYHA) function Class III or IV. Baseline echocardiographic parameters were as follows: the mean LVEF was 53.3 ± 9.3%, with a mean aortic valve area and aortic valve mean gradient of 0.81 ± 0.19 cm² and 45.7 ± 6.4 mmHg, respectively.

Due to the extensiveness of the CMR scan protocol, in which the 2D multi-venic and 4D flow mapping sequences were performed at the end of the scanning procedure, four patients failed to undergo the full scan protocol. Hence, there were two patients with missing 2D multi-venic data and two patients with missing 4D flow mapping data. In the remaining 17 patients, all 3 different flow sequences were successfully acquired.

Echocardiographic assessment of PVR

The mean duration between TAVR and TTE was 38 ± 11 days. TTE assessment showed none/trace PVR in 14 (66.7%) patients, mild PVR in 7 (33.3%) patients, and no patients with ≥ moderate PVR.

CMR quantification of RF

The CMR assessment of PVR is provided in Table 2.

Table 2 CMR and echocardiographic assessment of PVR

	Study population
Days after TAVR	38 ± 11
2D flow measurements (N = 21)	
FV, mL	83.5 ± 19.6
RV, mL	9.8 ± 7.9
RF, %	11.7 ± 10.0
Classification of PVR	
Mild or less than mild (RF ≤ 20%), n (%)	18 (85.7)
Moderate (RF 21–39%), n (%)	2 (9.5)
Severe (RF ≥ 40%), n (%)	1 (4.8)
2D multi-venic flow measurements (N = 19)	
FV, mL	73.6 ± 17.2
RV, mL	7.7 ± 4.9
RF, %	10.6 ± 7.0
Classification of PVR	
Mild or less than mild (RF ≤ 20%), n (%)	17 (89.5)
Moderate (RF 21–39%), n (%)	2 (10.5)
Severe (RF ≥ 40%), n (%)	0
4D flow measurements (N = 19)	
FV, mL	71.7 ± 14.6
RV, mL	7.0 ± 5.7
RF, %	9.6 ± 7.3
Classification of PVR	
Mild or less than mild (RF ≤ 20%), n (%)	16 (84.2)
Moderate (RF 21–39%), n (%)	3 (15.8)
Severe (RF ≥ 40%), n (%)	0
TTE classification of PVR^a	
None/trace, n (%)	14 (66.7)
Mild, n (%)	7 (33.3)
Moderate, n (%)	0
Severe, n (%)	0

Data are presented as mean ± standard deviation or as number (%).

^aTTE was performed on the same day as CMR.

FV, forward volume; PVR, paravalvular regurgitation; RF, regurgitant fraction; RV, regurgitant volume; TAVR, transcatheter aortic valve replacement; TTE, transthoracic echocardiography; venc, velocity encoding.

Standard 2D flow mapping

Standard 2D flow assessment was done in 21 patients. The mean FV measured with a high venc (≥ 180 cm/s) was 83.5 ± 19.6 mL. The mean RV measured with a low venc (75 cm/s) was 9.8 ± 7.9 mL, resulting in a mean RF of 11.7 ± 10.0%. The mean acquisition times of the high and low venc sequences were 15.0 ± 2.2 and 15.0 ± 2.0 s, respectively. Compared with standard 2D flow mapping assessment, TTE underestimated the degree of PVR in 3 of 21 patients.

Two-dimensional multi-venic flow mapping

In 19 patients, 2D multi-venic flow mapping measurements were performed. The mean FV and RV were 73.6 ± 17.2 and 7.7 ±

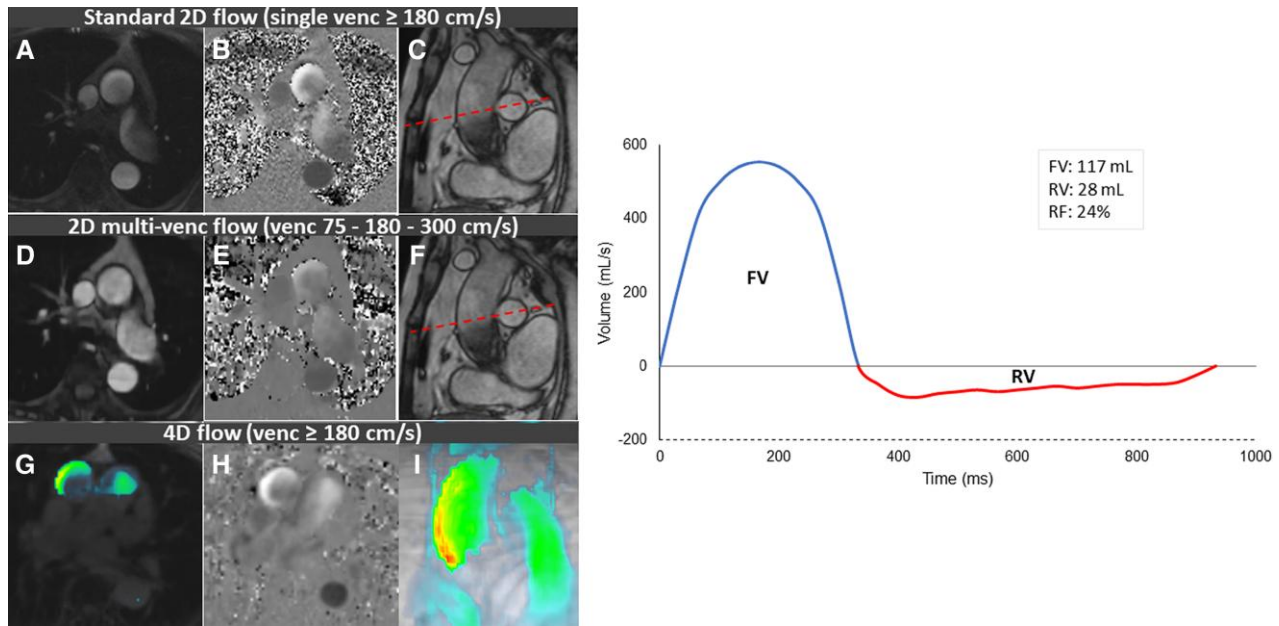


Figure 3 Example of the three different CMR flow assessment techniques. Left: upper row: standard 2D flow mapping with (A) magnitude image; (B) phase image; (C) the level of acquisition of PC CMR indicated with the dashed line. Middle row: 2D multi-venic flow mapping with (D) magnitude image; (E) phase image; (F) the level of acquisition of PC CMR indicated with the dashed line. Lower row: 4D flow mapping with (G) magnitude image; (H) phase image; and (I) 4D flow image reconstruction. Right: the quantification of the RF was done by dividing the RV by the FV multiplied by 100. FV, forward volume; RF, regurgitant fraction; RV, regurgitant volume; venc, velocity encoding.

cardiac anatomy knowledge or specific imaging planes for acquisition, since it can cover the whole heart.³⁶ The continuous presence of a CMR laboratory technician could therefore be reduced. Third, the analysis plane can be set anywhere within the acquisition area retrospectively, which reduces planning effort, allows standardization of acquisition, and facilitates post-processing. Last, 4D flow mapping acquisitions do not depend on breath-holds, allowing the patient to breathe freely. Especially in the TAVR population, this can be of great benefit. Besides the numerous advantages of the 4D flow mapping acquisition, important disadvantages are the longer acquisition time of ~3 min and the risk of breathing artefacts. In addition, adequate 4D flow planning and acquisition has a learning curve and requires strong collaboration between cardiac imagers and their technicians during the implementation phase.

Future perspectives

The annual number of patients treated by TAVR is still rising,²⁶ with an inherent increase in cases in which the degree of PVR as assessed with TTE can be debatable. In these cases, CMR can be the designated imaging modality to provide clarity regarding PVR severity. The advantages of the novel CMR techniques addressed in this study can be considered in the decision of which acquisition technique to use. Given the high correlation with the standard 2D flow mapping technique and the acceptable inter-observer variability, 4D flow mapping appears to be a valuable alternative to standard 2D flow mapping, limiting the risk of inadequate plane selection and reducing the need for the continuous presence of a CMR laboratory technician.

Limitations

This study has some limitations. First, the number of patients in this study is relatively low, and not all patients underwent all three different flow assessment techniques due to the extensive scan protocol.

However, we do believe that the results of this study would not change even if a larger sample size is used, given the strong correlation between the CMR sequences and the high reproducibility of these sequences. Second, the 4D flow imaging plane was visually set at the same level as for the standard 2D and 2D multi-venic sequences. Therefore, it is unknown whether differences in flow volumes can be attributed to differences in plane selection. Third, since the number of patients with ≥ moderate PVR is limited, conclusions on the reclassification between the CMR modalities for this patient category can not be drawn. Fourth, this study was performed solely with the Abbott Portico valve, precluding the direct translation of these results to other types of TAVR devices. However, since the focus of this study was to compare the three different CMR flow mapping acquisitions for PVR assessment, rather than describing the CMR-RF of different valve types, the aforementioned point is of less importance. Fifth, the 4D flow mapping sequence uses only a single (high) venc value, as opposed to the standard 2D flow and 2D multi-venic flow sequences, thereby potentially limiting the accuracy of the RVs.

Conclusion

Two-dimensional multi-venic and 4D flow mapping are two novel techniques that have a high correlation with standard 2D flow mapping in the quantification of PVR after TAVR. The high acquisition speed of the 2D multi-venic sequence and the free-breathing acquisition with retrospective plane selection with 4D flow mapping provide useful advantages, especially in the frail TAVR population. Given these practical advantages, the use of these techniques should be considered.

Conflict of interest: M.H.v.W. has been a proctor and consultant for Abbott Vascular. D.G. has been an employee of Siemens Healthcare GmbH. N.v.R. has received research funding from Abbott, Philips, Medtronic, and Biotronik; has served as a consultant for RainMed,

Castor, and Medtronic; and has received speaker fees from Abbott and Bayer. R.N. has received research funding from Philips Volcano and Biotronik; has served as a consultant for BMS and Sanofi; and has received speaker fees from BMS, Canon, Pfizer, and Sanofi. The other authors do not have potential conflicts of interest or disclosures to report.

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Data availability

The data underlying this article are available from the corresponding author on reasonable request.

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