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SCIENTIFIC EDITORIAL

Four-dimensional flow cardiovascular magnetic resonance: Towards accurate flow quantification?

IRM cardiovasculaire 4D des flux : vers une quantification précise des flux ?

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Received 12 February 2019; received in revised form 4 March 2019; accepted 4 March 2019
Available online 21 March 2019

KEYWORDS

Cardiovascular magnetic resonance;
Flow study;
4D flow

MOTS CLÉS

IRM cardiovasculaire ;
Étude des flux ;
Flux

Cardiovascular magnetic resonance (CMR) has become an important tool for the clinical assessment of patients with cardiovascular disease. Besides morphological information, CMR provides functional information on cardiac function, perfusion, myocardial viability and blood flow. Phase contrast CMR (PC-CMR) relies on the direct relationship between

blood flow velocity and the phase of the magnetic resonance signal. Two-dimensional (2D) cine PC-CMR yields a series of anatomical (magnitude) and flow velocity (phase difference) images that represent the temporal changes in morphology and blood flow over the cardiac cycle. The 2D location is positioned normal to the direction of the vessel, with typical imaging parameters: spatial resolution 1.5–2.5 mm; temporal resolution 30–40 ms; slice thickness 5–8 mm. 2D PC-CMR is a routine part of standard CMR for the assessment of regional blood flow in the heart and great vessels [1,2]. The velocity encoding sensitivity (Venc), which represents the maximum flow velocity that can be acquired, is set by the operator to avoid velocity aliasing, but also minimize velocity noise.

More recently, time-resolved PC-CMR, with velocity encoding along all three flow directions and three-dimensional (3D) anatomical coverage (four-dimensional [4D] flow CMR), has been applied for the evaluation of cardiovascular haemodynamics. Whole heart 4D flow CMR is acquired using electrocardiography and (usually) respiratory motion compensation. 3D velocity encoding is used to obtain velocity-sensitive phase images, which are subtracted from reference images to calculate blood flow velocities along all three spatial dimensions (V_x, V_y and V_z).

Abbreviations: 2D, two-dimensional; 3D, three-dimensional; 4D, four-dimensional; CMR, cardiovascular magnetic resonance; PC-CMR, phase contrast cardiovascular magnetic resonance.

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<https://doi.org/10.1016/j.acvd.2019.03.001>

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The acquisition generates a 3D anatomical representation of the underlying cardiovascular geometry needed for 3D flow visualization and retrospective flow quantification. 4D flow CMR can provide information on the temporal and spatial evolution of 3D blood flow, with full volumetric coverage of any cardiac or vascular region of interest (at least 40 ms temporal resolution). A flexible retrospective quantification may be assessed at any location within the 3D data volume on an off-line workstation, and provides variables such as total flow, flow-time curves, peak velocities, pressure gradients or regurgitant fraction [3,4]. The combination of 3D blood flow visualization with flow quantification enables unprecedented comprehensive evaluation of the impact of cardiovascular diseases on global and local changes in cardiac or vascular haemodynamics [5,6].

Recent technical developments, including the utilization of advanced parallel imaging techniques, have resulted in reasonable overall scan times of <10 minutes for 4D flow CMR of the aorta or whole heart coverage, depending on the breathing patterns and heart rate of the patients. Continued developments in sparse sampling techniques (compressed sensing, radial acquisition and k-t GRAPPA) or multidimensional parallel imaging might further improve scan times. The clinical application of 4D flow CMR has become more feasible, as illustrated by the large number of recent reports on the clinical utility of the technique. Some 4D flow CMR studies have compared blood flow variables with markers determined by Doppler ultrasound and echocardiography, such as peak pressure gradient ($r=0.96$; $P<0.05$) [7] or peak and mean velocities ($r=0.83$ and $r=0.76$, respectively) [8]. In addition, 4D flow CMR can provide improved clinical haemodynamic assessment, by deriving additional metrics of cardiovascular haemodynamics, such as wall shear stress [9], pressure difference [7,10], pulse wave velocity [11] or turbulent kinetic energy [12]. Data have shown excellent agreement with 2D cine-PC-MR for flow quantification [13]. Furthermore, 4D flow CMR has good scan-rescan reproducibility and low interobserver and intraobserver variability of flow quantification for intracranial, cervical, thoracic and abdominal applications.

4D flow CMR has been applied and validated in many clinical applications. We will provide a concise and non-exhaustive overview. 4D flow CMR may be very useful for congenital heart disease, and whole heart techniques allow for a noninvasive comprehensive assessment of cardiovascular haemodynamics in the heart and surrounding great vessels, with flexible off-line quantitative analysis in any plane. 4D flow CMR is also very useful for diseases of the thoracic aorta. In patients with bicuspid aortic valve and aortic aneurysm, 4D flow CMR has shown high velocity flow jets impinging on the ascending aortic wall and resulting in increased wall shear stress, as well as increased helical flow in the ascending aorta [14]. Patients with aortic coarctation have flow jet eccentricity following the coarctation, resulting in jet impingement along the descending aorta. 4D flow CMR facilitates a comprehensive velocity assessment throughout the entire affected region near the coarctation [15]; it allows for direct accurate quantification of intracardiac shunts, and may show potential for aortic and pulmonary valve stenosis and regurgitation. 4D flow data may provide more advanced metrics of haemodynamics associated with complex blood flow patterns, such as

highly helical flow in aortic valve disease. These variables include the quantification of vorticity, helicity, flow angle, wall shear stress and turbulent and viscous energy loss. Alterations in these haemodynamic variables may affect signalling and organization of endothelial cells. 4D flow may bring new insights in the assessment of hepatic and portal venous flow of patients with advanced liver cirrhosis.

Errors and limitations

There are various sources of phase offset errors in 4D flow CMR that can alter image quality and impair measurements. The most common include phase offset errors due to Eddy currents, Maxwell terms and gradient field non-linearity. Appropriate correction strategies must be applied to compensate for these potential sources of error before processing of the data. The corrections for Maxwell terms and gradient field non-linearity are performed during image reconstruction, without user interaction. Correction for Eddy currents has to be integrated into the data analysis workflow. The approach is based on thresholding, to identify regions with static tissue. These regions are then used to estimate Eddy current-induced linearly varying phase offset errors, which are subsequently subtracted from the entire image. Postprocessing is time consuming, and is performed off-line on specific workstations. Limited spatial resolution is a drawback for analysis in small vessels. An additional barrier to the widespread clinical implementation of 4D flow CMR is the lack of validation against clinical gold standards, e.g. invasive measurements of blood flow variables. A common effort is needed to work toward increased availability and clinical adoption of 4D flow CMR, by refining the clinical 4D flow workflow, determining the most useful ways to present the data and enabling quick data postprocessing in the archive systems.

Disclosure of interest

The author declares that he has no competing interest.

References

- [1] Kilner PJ, Yang GZ, Mohiaddin RH, Firmin DN, Longmore DB. Helical and retrograde secondary flow patterns in the aortic arch studied by three-directional magnetic resonance velocity mapping. *Circulation* 1993;88:2235–47.
- [2] Pelc NJ, Herfkens RJ, Shimakawa A, Enzmann DR. Phase contrast cine magnetic resonance imaging. *Magn Reson Q* 1991;7:229–54.
- [3] Bogren HG, Mohiaddin RH, Yang GZ, Kilner PJ, Firmin DN. Magnetic resonance velocity vector mapping of blood flow in thoracic aortic aneurysms and grafts. *J Thorac Cardiovasc Surg* 1995;110:704–14.
- [4] Wigstrom L, Sjoqvist L, Wranne B. Temporally resolved 3D phase contrast imaging. *Magn Reson Med* 1996;36:800–3.
- [5] Bogren HG, Buonocore MH, Valente RJ. Four-dimensional magnetic resonance velocity mapping of blood flow patterns in the aorta in patients with atherosclerotic coronary artery disease compared to age-matched normal subjects. *J Magn Reson Imaging* 2004;19:417–27.

- [6] Kozerke S, Hasenkam JM, Pedersen EM, Boesiger P. Visualization of flow patterns distal to aortic valve prostheses in humans using a fast approach for cine 3D velocity mapping. *J Magn Reson Imaging* 2001;13:690–8.
- [7] Bock J, Frydrychowicz A, Lorenz R, et al. In vivo noninvasive 4D pressure difference mapping in the human aorta: phantom comparison and application in healthy volunteers and patients. *Magn Reson Med* 2011;66:1079–88.
- [8] Jiang J, Strother C, Johnson K, et al. Comparison of blood velocity measurements between ultrasound Doppler and accelerated phase contrast MR angiography in small arteries with disturbed flow. *Phys Med Biol* 2011;56:1755–73.
- [9] Markl M, Wallis W, Harloff A. Reproducibility of flow and wall shear stress analysis using flow-sensitive four-dimensional MRI. *J Magn Reson Imaging* 2011;33:988–94.
- [10] Ebbers T, Wigstrom L, Bolger AF, Engvall J, Karlsson M. Estimation of relative cardiovascular pressures using time-resolved three-dimensional phase contrast MRI. *Magn Reson Med* 2001;45:872–9.
- [11] Markl M, Wallis W, Bredecke S, Simon J, Frydrychowicz A, Harloff A. Estimation of global aortic pulse wave velocity by flow-sensitive 4D MRI. *Magn Reson Med* 2010;63:1575–82.
- [12] Dyverfeldt P, Sigfridsson A, Kvitting JP, Ebbers T. Quantification of intravoxel velocity standard deviation and turbulence intensity by generalizing phase contrast MRI. *Magn Reson Med* 2006;56:850–8.
- [13] Wentland AL, Grist TM, Wieben O. Repeatability and internal consistency of abdominal 2D and 4D phase contrast MR flow measurements. *Acad Radiol* 2013;20:699–704.
- [14] Barker AJ, Markl M, Burk J, et al. Bicuspid aortic valve is associated with altered wall shear stress in the ascending aorta. *Circ Cardiovasc Imaging* 2012;5:457–66.
- [15] Allen BD, Barker AJ, Carr JC, Silverberg RA, Markl M. Time-resolved three-dimensional phase contrast MRI evaluation of bicuspid aortic valve and coarctation of the aorta. *Eur Heart J Cardiovasc Imaging* 2013;14:399.