Received: 04 July 2011,

Revised: 21 October 2011,

Accepted: 27 October 2011,

Published online in Wiley Online Library: 19 December 2011

(wileyonlinelibrary.com) DOI: 10.1002/nbm.1812

Real-time phase-contrast MRI of cardiovascular blood flow using undersampled radial fast low-angle shot and nonlinear inverse reconstruction

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Velocity-encoded phase-contrast MRI of cardiovascular blood flow commonly relies on electrocardiogram-synchronized cine acquisitions of multiple heartbeats to quantitatively determine the flow of an averaged cardiac cycle. Here, we present a new method for real-time phase-contrast MRI that combines flow-encoding gradients with highly undersampled radial fast low-angle shot acquisitions and phase-sensitive image reconstructions by regularized nonlinear inversion. Apart from calibration studies using steady and pulsatile flow, preliminary *in vivo* applications focused on through-plane flow in the ascending aorta of healthy subjects. With bipolar velocity-encoding gradients of alternating polarity that overlap the slice-refocusing gradient, the method yields flow-encoded images with an in-plane resolution of 1.8 mm, section thickness of 6 mm and measuring time at 3 T of 24 ms (TR/TE = 3.44/2.76 ms; flip angle, 10°; seven radial spokes per image). Accordingly, phase-contrast maps and corresponding velocity profiles achieve a temporal resolution of 48 ms. The evaluated peak velocities, stroke volumes, flow rates and respective variances over at least 20 consecutive heartbeats are in general agreement with literature data. Copyright © 2011 John Wiley & Sons, Ltd. Supporting information may be found in the online version of this paper

Keywords: cardiovascular MRI; flow imaging; heart function; nonlinear inverse reconstruction; phase-contrast imaging; real-time imaging

INTRODUCTION

Velocity-encoded phase-contrast MRI is a well-established method for quantitative mapping of flow velocities. It is based on Erwin Hahn's 1960 observation of a phase shift for spins moving in the presence of a magnetic field gradient (1), and has been developed during the early stages of MRI [see, for example, refs. (2–5)]. In its simplest form, quantitative velocities of a constant flow may be derived from the phase difference between two cross-sectional images with different velocity-encoding gradients (e.g. two bipolar gradients with opposite polarity). The resulting three-dimensional dataset comprises two spatial dimensions and one velocity (phase-difference) dimension, so that each image pixel presents with a single mean velocity value. Applications to clinical studies of the human heart and great vessels have been reviewed several times [see, for example, refs. (6–8)].

In order to obtain sufficient temporal resolution during the cardiac cycle, MRI studies commonly rely on electrocardiogram (ECG)-synchronized cine acquisitions of multiple heartbeats, either during free breathing or in suspended respiration. The approach may be affected by an inappropriate temporal and/or spatial averaging of phase information as a result of the calculation of a phase-difference value from the complex mean of signals of multiple spins (9), and therefore bears the risk of underestimating maximum flow velocities compared with, for example, Doppler sonography. Moreover, it renders the technique susceptible to flow inconsistencies caused by cardiac arrhythmias. Indeed, the direct assessment of physiological variability or pathological

irregularities, such as arrhythmia, is not feasible with acquisitions extending over several heartbeats.

Many ideas have been proposed to ameliorate the detrimental effects of motion in MRI. In order to allow for free breathing in cardiovascular applications, the most successful attempts range from navigator-type corrections based on concentric rectangular strips or blades rotated about the center of k space (10) to the use of respiratory self-gating (11). A similar wide range of methods has been proposed for real-time MRI since the early work by Chapman *et al.* (12) using echo-planar imaging. The approaches employ low-flip-angle radiofrequency (RF) excitations of gradient echoes and cover k space with the use of an echo-planar (13), Cartesian (14–16), radial (17) or spiral (18,19) trajectory. In recent

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Abbreviations used: ECG, electrocardiogram; FLASH, fast low-angle shot; FOV, field of view; GRAPPA, generalized autocalibrating partially parallel acquisition; RF, radiofrequency; SD, standard deviation; SENSE, sensitivity encoding.



years, further acceleration has been achieved by combinations with parallel imaging based on sensitivity encoding (SENSE) (20–23) or generalized autocalibrating partially parallel acquisition (GRAPPA) (24), or by further exploiting temporal continuities (25). Nevertheless, the hitherto still limited spatiotemporal resolution and/or overall image quality of many proposed techniques must be considered responsible for a lack of general clinical acceptance.

Most recently, we have made significant advances in high-quality real-time MRI using highly undersampled radial fast low-angle shot (FLASH) acquisitions with image reconstruction by regularized nonlinear inversion (26–28). Preliminary applications to cardiovascular MRI allowed for serial real-time images (i.e. movies) with an in-plane resolution of 1.5–2.0 mm and an acquisition time of 20–30 ms (28,29). Thus, the purpose of this study was to combine the principles of velocity-encoded phase-contrast MRI with the acquisition of two highly undersampled radial FLASH images to achieve a temporal resolution for real-time flow MRI of less than 50 ms.

METHODS

SUBJECTS

Apart from pilot studies that were performed to optimize the experimental conditions and define the measurement protocol, a total of six healthy male subjects without prior cardiovascular symptoms and no known illness participated in the real-time assessment of through-plane flow in the ascending aorta. The deliberately chosen heterogeneous group of subjects presented with different ages (range, 16–57 years), body weights and physical constitutions. All subjects gave written informed consent before each MRI examination.

Velocity-encoded phase-contrast MRI in real time

All studies were conducted at 3 T using a commercial MRI system (TIM Trio, Siemens Healthcare, Erlangen, Germany) and a body coil for RF excitation. Subjects were examined in a supine position and MRI signals were acquired with the use of a 32-channel cardiac coil comprising an anterior and posterior 16-element array.

Real-time phase-contrast MRI employed a highly undersampled RF-spoiled radial FLASH sequence (30) with a motioncompensated slice-selective gradient and pairs of flow-encoding gradients in two interleaved acquisitions (every other TR). Velocity encoding to measure through-plane flow was implemented along the slice-selective gradient axis in two different ways. As indicated in Fig. 1, the first scheme employed bipolar velocity-encoding gradients with opposite polarity (Min-Max version), and the second scheme applied no flow-encoding gradient during one acquisition and a correspondingly stronger velocity-encoding gradient during the other (Zero-Max version, not shown). In either case, the field of view (FOV) in the phase-contrast or velocity dimension was set to VENC = 150 cm/s. The velocity-encoding gradients were combined with the refocusing part of the slice-selective gradient, as described previously for conventional cine techniques (31). However, the actual implementation did not realize a further overlap with the two frequency-encoding gradients to retain more flexibility for the incorporation of flow-encoding gradients along arbitrary directions.

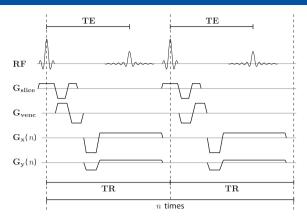


Figure 1. Schematic radiofrequency (RF) pulse and gradient sequence for real-time velocity-encoded phase-contrast MRI of through-plane flow. The method is based on highly undersampled radial fast low-angle shot (FLASH) (TR/TE=3.44/2.76 ms; flip angle, 10°) and relies on two interleaved image acquisitions (n=7 spokes per image) with bipolar velocity-encoding gradients G_{VENC} of opposite polarity (VENC=150 cm/s). These gradients are combined with the refocusing part of the motion-compensated slice-selective gradient G_{slice} , but do not overlap with the frequency-encoding gradients G_x and G_y to allow for additional flow-encoding gradients along arbitrary directions. The implementation yields a temporal resolution of 48 ms per phase-contrast map.

Gradient switching times were kept to a normal (rather than fast) mode, which refers to slew rates below the technical possibilities of the MRI system used. This choice avoids peripheral nerve stimulation under all experimental circumstances. Accordingly, the Zero-Max version resulted in minimum values of TR/TE = 3.15/2.46 ms and a temporal resolution of 44 ms for phase-contrast MRI. The corresponding results for the Min-Max version were TR/TE = 3.44/2.76 ms and a resolution of 48 ms.

All real-time applications relied on image acquisitions with only seven spokes (144 complex data points each) covering a FOV of $256 \times 256\,\mathrm{mm}^2$ at an in-plane resolution of 1.8 mm and with a slice thickness of 6 mm (flip angle, 10°). Successive acquisitions employed five different sets of complementary spokes, as described previously (30). Their purpose is twofold: first, the strategy generates slightly different streaking artifacts from frame to frame that are more easily treatable than constant artifacts and, second, combined datasets (i.e. 35 spokes) are exploited for immediate online control during scanning using gridding reconstructions (see below). In our current setting, these images are replaced within a few minutes by the respective inverse reconstructions performed offline.

Image reconstructions

Fully sampled radial FLASH images served as multislice localizer scans, as well as for comparative phantom studies. Such images were reconstructed by conventional gridding. Similarly, online monitoring of velocity-encoded magnitude images and phase-contrast maps during real-time flow MRI was accomplished by gridding of combined datasets with 35 spokes and a sliding window factor of five. Thus, these gridding images cover a five-fold longer acquisition time than the respective nonlinear inverse reconstructions, but offer the same frame rate. Although partially compromised by streaking artifacts and enhanced noise, such reconstructions provide sufficient quality to supervise the actual measurement. In order to gain real-time speed, gridding



reconstructions employed an online channel compression (30) based on a principal component analysis of the first few scans. It reduces the data from the 32-channel acquisition to the most relevant six or eight components ('virtual channels') and thus ensures sufficient reconstruction speed for immediate visual control.

For offline reconstruction using regularized nonlinear inversion, the incoming data were automatically exported to a computer equipped with eight GTX580 graphical processing units, each providing 512 processing cores (Nvidia, Santa Clara, CA, USA). Extending the original algorithm to jointly estimate image content and coil sensitivity profiles (19) and its adaptation to real-time imaging (27,28), the present framework allows for the reconstruction of phase-sensitive images before taking a phase-difference map. Although still applying a temporal median filter to the reconstructed magnitude images (i.e. to reduce residual streaking artifacts), the phase-contrast maps are calculated from the unfiltered complex-valued images. To remove unwanted phase components from coil sensitivities or off-resonance effects, each reconstructed image was multiplied with all coil sensitivities to obtain separate images for all channels. Phase-difference maps were then calculated for each individual channel by multiplying the first flow-encoded image with the complex conjugate of the associated second flow-encoded image. The final phase-contrast map was determined by summing the results of all channels and extracting the phase (32), which also ensures that contributions from individual channels are weighted with their respective energy in each pixel.

Once the offline calculation of a real-time MRI series had been completed, the magnitude images and corresponding phasecontrast maps were re-imported into the MRI scanner by exploiting the manufacturer's mechanism of a 'retrospective reconstruction'. However, instead of again reconstructing the raw data which are temporally stored on the scanner, the formalism was used to create DICOM images with the correct header information from the offline reconstructions for storage in the system's databank. To date, where offline reconstructions still take some time, this strategy provides a simple and almost fully automatic by-pass of the normal reconstruction process with only a single manual intervention (i.e. starting the re-import on the scanner). In fact, most nonlinear inverse reconstructions of a real-time heart study are already available after the end of the examination, so that the images may immediately be re-imported onto the scanner for further evaluation.

The total reconstruction time for two flow-encoded images and the corresponding phase-contrast map is currently about 0.2 s using four parallel graphical processing units and a grid size of 432×432 (acquisition matrix of 288×288) with an additional oversampling factor of 1.5 for reconstruction). This includes a reduction of the original 32-channel dataset to 12 virtual channels by a principal component analysis and the use of six Newton steps for the iterative optimization (28).

Flow calibration

Although commercial phase-contrast flow MRI sequences employ similar, if not identical, combinations of flow-encoding and imaging gradients, we performed independent calibration experiments to evaluate the accuracy of the entire acquisition and analysis process, and to ascertain the equivalence of phase-contrast maps that were obtained by gridding of fully sampled datasets with those calculated by nonlinear inversion of highly undersampled datasets.

Flow calibrations used a 16-mm-diameter stiff tube to simulate the properties of the aorta. Flow at different rates was provided by a computer-controlled immersion pump (Lux Plus KTW270, Herzog, Göttingen, Germany) and independently measured by an external flow meter (FCH-C-PA, BioTech, Vilshofen, Germany) outside the main magnetic field of the scanner. Studies of steady flow were performed with both the Zero-Max and Min-Max flowencoding schemes at temporal resolutions of 44 and 48 ms, respectively.

Pulsatile flow was generated by a repetitive pump protocol with brief periods of high velocity (0.1–0.4 ms), followed by a lower velocity for a total cycle time of 2.0 s. High- and low-flow conditions referred to pump voltages differing by a factor of two. Respective phase-contrast MRI studies were performed with the Min-Max encoding scheme.

Human studies

Because the calibration measurements resulted in a lower phase offset for stationary spins when using the Min-Max version (see Results), all real-time phase-contrast MRI studies of the ascending aorta were performed with polarity-alternated bipolar velocity-encoding gradients at a temporal resolution of 48 ms in a transverse orientation. These real-time acquisitions were accompanied by ECG recordings to provide an ECG stamp, i.e. the relative time to the last R wave, to each magnitude image and phase-contrast map.

The total measurement protocol comprised real-time phase-contrast MRI movies during free breathing (about 30 s), as well as during breath hold (about 15 s), in inspiration and expiration. Three free-breathing studies were interleaved with two repetitions of two breath hold scans (reversed order of inspiration and expiration), yielding a total of seven experimental runs. Pauses between scans were at least 30 s. The total examination time, including scans for positioning (sequential multislice radial FLASH MRI), was about 15 min.

Flow analysis

All flow-encoded phase-contrast maps were analyzed using ARGUS supplied by the manufacturer (Siemens Healthcare, Erlangen, Germany). In addition, human flow MRI data were also evaluated by QFlow 5.2 (Medis, Leiden, the Netherlands) which offers greater flexibility and extended parametric access. Both programs define a region of interest by complementing a manual outline of the aorta in a single magnitude image with an automatic segmentation and propagation to all images and maps of a series. In human studies, the automatic segmentation by QFlow 5.2 was visually controlled and manually corrected if necessary.

Of note, instead of looking at a single synthetic cardiac cycle from an ECG-synchronized cine recording (i.e. about 20 images), the present analyses comprised 15–20 consecutive heartbeats (each with about 20 images). Although this procedure offers access to the physiological variance of flow parameters from heartbeat to heartbeat, it is also possible to combine all (about 300–400) images into a single dataset with individual images sorted according to the ECG. At least for ARGUS, however, this combination strategy turned out to be more vulnerable to segmentation errors.

The main flow parameters derived from real-time phase-contrast MRI movies were the maximum 'peak' velocity (for ARGUS: mean of one pixel and four neighbors), the stroke volume defined as the output of the left ventricle into the ascending aorta during



one cardiac cycle, and the flow rate as the mean cardiac output per minute. In contrast with conventional cine recordings, where, for example, the latter parameter must be extrapolated from a single synthetic cardiac cycle to a full minute, real-time evaluations of much longer series of heartbeats allow for more accurate estimates. In fact, it is even possible to extend the measurement to a minute. Finally, color-coded visualization of velocity distributions within the ascending aorta was accomplished using routines written in MATLAB (MathWorks, Natick, MA, USA).

RESULTS AND DISCUSSION

Flow calibration

During experimental optimization of the flow-encoding radial FLASH MRI sequences, the residual phase error for stationary spins, i.e. a nonzero mean signal in a stationary region of a phasecontrast map, turned out to be three times smaller for the Min-Max than for the Zero-Max version. A likely explanation for this smaller phase offset is the larger similarity of the overlapping slice-refocusing and flow-encoding gradients when using two polarity-alternated bipolar gradients of equal strength. However, the Zero-Max scheme may lead to a larger disparity, so that residual gradient imperfections (e.g. as a result of eddy currents) are less well compensated in a phase-difference map. When averaging over an area comparable with that of the human aorta, the observed phase offsets corresponded to about 0.38 L/min for the Zero-Max and only 0.12 L/min for the Min-Max version. Thus, at the expense of a slightly longer acquisition time, we therefore decided to exclusively use bipolar velocity-encoding gradients with opposite polarity for human applications. Moreover, the results presented here are not corrected for residual phase offset.

Figure 2 summarizes the results of a Min-Max calibration study for a 16-mm-diameter flow phantom with constant flow at different rates. The recordings of a flow meter are compared with MRI flow rates derived from gridding reconstructions of fully sampled radial FLASH images, as well as nonlinear inverse reconstructions of highly undersampled acquisitions, with identical parameters as in the human studies, at a resolution of 48 ms. In order to apply the same evaluation procedure as *in vivo*, the images

were labeled with a simulated ECG signal and analyzed by ARGUS. The results demonstrate excellent agreement of MRI and flow meter values for the range of 4–8 L/min. The observation of a slight overestimation for the lower values of 2–3 L/min may be explained by the fact that voxels in the border region experience a disproportionate signal increase from moving spins compared with the contribution from stationary spins (33). Of note, there seems to be no difference between the two types of acquisition, so that real-time MRI studies based on nonlinear inverse reconstructions of highly undersampled datasets may be performed with the same accuracy as conventional phase-contrast recordings.

The calibration studies were complemented by real-time phase-contrast MRI of pulsatile flow generated by protocols with two different velocities (as controlled by the pump voltage) and different durations for a total cycle time of 2.0 s (same phantom as above). The protocols and resulting flow parameters are summarized in Table 1. Although no independent assessment of the dynamic flow velocities was available, the estimated flow rates from four consecutive cycles are again in excellent agreement with the experimentally determined values. The calibration results for both steady and pulsatile flow therefore confirm the quantitative reliability of the proposed real-time phase-contrast MRI technique for the measurement of flow rates.

Human studies

Focusing on through-plane flow in the ascending aorta, this preliminary real-time flow MRI study was successful in all subjects. The data were not affected by putative motion problems caused by free breathing, although it cannot be excluded that future applications to other vessels may require more sophisticated strategies that detect respiratory variations directly from the data. All real-time phase-contrast MRI data (six subjects, seven runs each, 15–20 heartbeats each, about 5000 images and phase-contrast maps) were analyzed with both ARGUS and QFlow 5.2 (not shown) and revealed excellent agreement.

Figure 3 shows a selected magnitude image and corresponding phase-contrast map of the ascending aorta from a real-time movie acquisition (temporal resolution of 48 ms) at a time point with

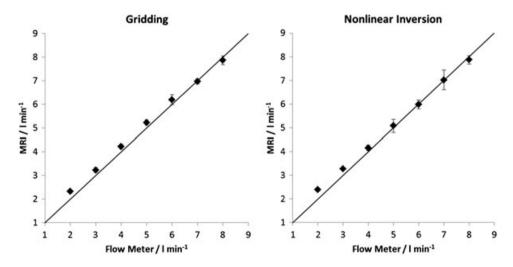


Figure 2. Flow rates (L/min) for a calibration phantom (16-mm-diameter tube) with constant flow measured by phase-contrast MRI (VENC = 150 cm/s) using either fully sampled radial fast low-angle shot (FLASH) with gridding (225 spokes, other parameters as in Fig. 1) or undersampled acquisitions (seven spokes) with nonlinear inverse reconstruction. Calibration values refer to an external flow meter (accuracy, ± 0.05 L/min). MRI flow rates represent the mean \pm standard deviation (SD) of data from 10 simulated heartbeats. The diagonal line refers to exact calibration.



Table 1. Pulsatile flow calibration									
Repetitive protocol		Flow velo	city (cm/s)	Flow rate (L/min)					
High flow	Low flow	High flow	Low flow	MRI	Exp.				
0.4 s	1.6 s	$\textbf{72.2} \pm \textbf{0.3}$	49.7 ± 0.5	$\textbf{5.35} \pm \textbf{0.03}$	5.30				
0.2 s	1.8 s	$\textbf{58.2} \pm \textbf{2.0}$	$\textbf{44.4} \pm \textbf{0.2}$	$\textbf{4.63} \pm \textbf{0.08}$	4.63				
0.1 s	1.9 s	$\textbf{52.5} \pm \textbf{1.1}$	$\textbf{44.4} \pm \textbf{0.1}$	$\textbf{4.38} \pm \textbf{0.07}$	4.47				

MRI-derived flow parameters are given as the mean \pm standard deviation (SD) averaged across four repetitive 2.0-s cycles. Flow velocities represent peak and baseline values during respective periods. High and low flow conditions refer to pump voltages differing by a factor of two.

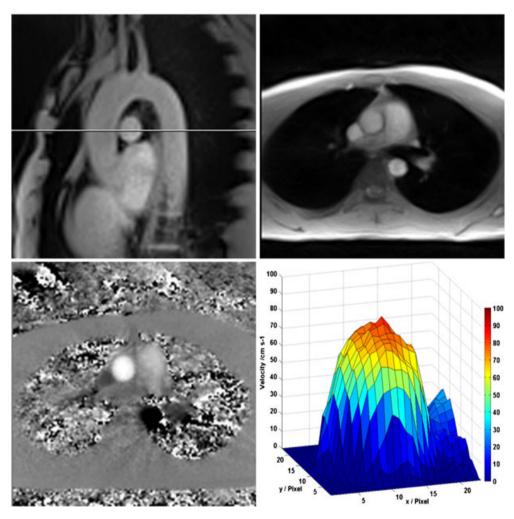


Figure 3. Real-time velocity-encoded MRI of through-plane flow in the ascending aorta (resolution, 48 ms) of a healthy subject. Top left: aortic arch with transverse slice orientation [radial fast low-angle shot (FLASH) localizer]. Top right: selected magnitude image, as well as corresponding phase-contrast map (bottom left) and flow velocities within the aorta (bottom right) (color-coded range from 0 to 100 cm/s). Other parameters as in Fig. 1.

strong systolic flow. In general, the image quality is surprisingly good in view of the extreme undersampling by a factor of 32, that is seven spokes relative to a formally required number of $144 \times \pi/2 = 226$ spokes. In particular, the phase-contrast map is only marginally affected by residual streaking artifacts (outside the area of interest), and the phase differences for stationary spins are rather homogeneous and close to zero. A dynamic representation of phase-contrast maps from 10 consecutive heartbeats of this dataset is available as Supporting Information (Movie S1). The

bottom right map in Fig. 3 visualizes the two-dimensional flow profile within the ascending aorta by color-coded velocities ranging from 0 (blue) to $100\,\mathrm{cm/s}$ (red).

Figure 4 depicts the dynamic velocity distributions of flow in the ascending aorta at a resolution of 48 ms. The series of 12 consecutive profiles (total duration, 576 ms) covers the early phase of a single cardiac cycle. In this subject, the velocity increases to a peak value of 80 cm/s (red) within about 150 ms (three frames). Its slightly slower decline is accompanied by the



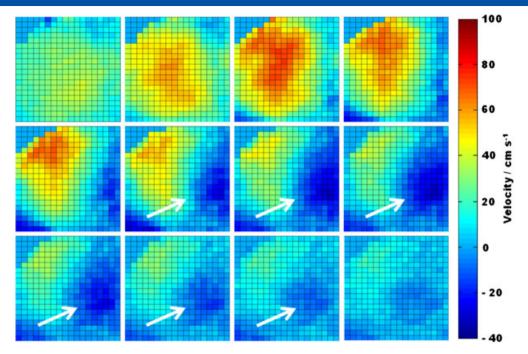


Figure 4. Real-time flow velocities (color-coded range from - 40 to 100 cm/s) within the ascending aorta (18 \times 20 pixels) at 12 consecutive time points during the early phase of a single cardiac cycle (resolution, 48 ms; total duration, 12 \times 48 ms = 576 ms; different subject from Fig. 3). Arrows indicate back flow (blue). Other parameters as in Fig. 1.

occurrence of back flow with velocities of up to 40 cm/s (arrows, dark blue). A dynamic visualization of real-time velocity profiles from 20 consecutive heartbeats is available as Supporting information (Movie S2).

Figure 5 demonstrates the variability of peak flow velocities and stroke volumes as a function of time (20 cardiac cycles) under the condition of free breathing. The lower portions of Fig. 5

contain temporal signal intensity profiles of the anterior chest wall from corresponding magnitude images that indicate inspiration (arrows) and expiration phases. Although peak velocities vary during the respiratory cycle (here about four to five cardiac cycles) by more than 10% between expiration and inspiration, the stroke volume remains more stable. The results indicate that the lowest peak velocities are obtained during expiration.

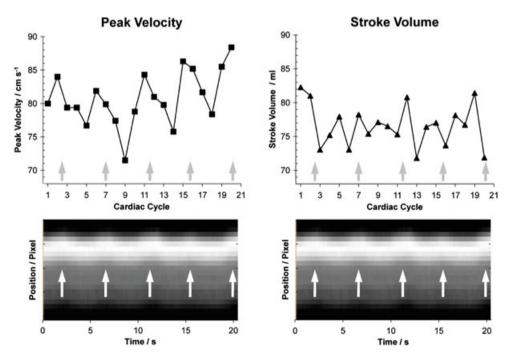


Figure 5. Real-time peak flow velocities (cm/s) in the ascending aorta and related stroke volumes (mL) over 20 consecutive heartbeats (resolution, 48 ms; same subject as in Fig. 3). The bottom parts refer to temporal signal intensity profiles of the anterior chest wall taken from the corresponding magnitude images. Arrows indicate inspiration. Other parameters as in Fig. 1.



Quantitative results for all subjects are summarized in Table 2 for free breathing. All flow parameters [mean ± standard deviation (SD)] are averaged across 20 heartbeats and three repetitions (corresponding to 60 heartbeats). The observed differences for the peak velocity between subjects most probably represent the physiological variation of hemodynamics in response to parameters such as, for example, body composition, blood pressure and state or alertness during MRI. As such, our small group of subjects varied considerably with respect to age, body weight and physical fitness, so that individual variances (i.e. intrasubject SD values) for peak velocities, stroke volumes and flow rates are much smaller than the differences between the corresponding mean values of individual subjects.

Mean flow parameters averaged across subjects are shown in Table 3 for free breathing and breath holding under inspiration and expiration. Peak velocities in the ascending aorta tend to be higher during breath hold than during free breathing, whereas stroke volumes remain remarkably constant. The former observation matches the finding of a respiratory-dependent variation of peak velocities as seen in Fig. 5. Both absolute and individually normalized flow rates increase during expiration, as expected for a lowered intrathoracic pressure. The considerably larger SD value for the stroke volume under breath hold with inspiration is most probably a result of an inconsistent performance of our subjects.

Altogether, the flow parameters for free breathing seem to better reflect a normal physiological state than those obtained during breath hold. This observation is supported by rather small SD values for peak velocities and flow rates under free breathing, despite the heterogeneity of the subjects. Larger studies are needed to confirm these initial findings.

CONCLUSIONS

This work describes the development and implementation of a real-time method for velocity-encoded phase-contrast MRI with a temporal resolution below 50 ms. Calibration studies confirmed the accuracy of flow rates obtained from highly undersampled radial FLASH acquisitions with nonlinear inverse reconstruction. The evaluation of real-time flow parameters for the ascending aorta of healthy human subjects revealed general agreement with literature data: for example, compare refs. (34,35) using MRI and non-MRI techniques. Moreover, the present results support the notion that free breathing is a preferred condition for the physiological characterization of cardiovascular flow. It is expected that real-time access to flow parameters (over any chosen period of time) will offer new potential for clinical assessments of pathological deviations or irregularities within single or serial heartbeats.

Although beyond the scope of this methodological work, a limitation of our study is the small number of subjects and the restriction to flow in the ascending aorta. A clinical evaluation of real-time flow measurements in direct comparison with ECG-gated phase-contrast MRI will address this shortcoming in the near future. Another restriction that needs to be taken into account is the available software for flow analysis. Although existing packages may be applicable, as used here, suitable adjustments will considerably accelerate the processing of data from a large number of heartbeats and more easily provide information about physiological or pathological variations. Pertinent improvements will be a prerequisite for the clinical acceptance of real-time flow MRI examinations.

Table 2. Flow parameters for the ascending aorta of six healthy subjects as determined by real-time velocity-encoded phase-contrast MRI during free breathing

Subject	Age (years)	Heart rate (beats/min)	Peak velocity (cm/s)	Stroke volume (mL)	Aortic area (cm²)	Flow rate (L/min)
1	24	62 ± 2	81 ± 9	102 ± 8	$\textbf{6.9} \pm \textbf{0.3}$	$\textbf{6.4} \pm \textbf{0.5}$
2	37	60 ± 4	73 ± 6	97 ± 13	$\textbf{6.9} \pm \textbf{0.5}$	$\textbf{5.8} \pm \textbf{0.8}$
3	57	68 ± 3	54 ± 3	96 ± 9	10 ± 0.4	$\textbf{6.5} \pm \textbf{0.6}$
4	28	64 ± 4	68 ± 4	74 ± 9	$\textbf{6.4} \pm \textbf{0.2}$	4.7 ± 0.6
5	22	89 ± 7	59 ± 4	68 ± 10	$\textbf{6.1} \pm \textbf{0.4}$	6.0 ± 0.9
6	25	57 ± 3	79 ± 6	76 ± 4	5.2 ± 0.3	4.3 ± 0.2

Values are given as mean \pm standard deviation (SD) averaged across 20 heartbeats and three repetitions (i.e. a total of 60 heartbeats).

Table 3. Mean flow parameters for the ascending aorta as determined by real-time velocity-encoded phase-contrast MRI during free breathing as well as breath hold in inspiration and expiration

Breathing condition	Peak velocity (cm/s)	Stroke volume (mL)	Flow rate (L/min)	Normalized flow rate (%)
Free	68.8 ± 10.6	$\textbf{85.7} \pm \textbf{13.6}$	$\textbf{5.6} \pm \textbf{0.9}$	100
Inspiration	$\textbf{73.9} \pm \textbf{15.0}$	$\textbf{87.4} \pm \textbf{23.6}$	$\textbf{5.8} \pm \textbf{1.3}$	102 ± 9
Expiration	74.5 ± 15.6	$\textbf{85.3} \pm \textbf{11.9}$	$\textbf{6.1} \pm \textbf{1.1}$	109 ± 15

Values are given as mean \pm standard deviation (SD) averaged across six subjects, as well as 20 heartbeats and three repetitions for free breathing (60 heartbeats) or 15 heartbeats and two repetitions for breath holding (30 heartbeats).



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