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Clinical Blood Flow Quantification With Segmented k-Space Magnetic Resonance Phase Velocity Mapping

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Purpose: To evaluate the accuracy of segmented k-space magnetic resonance phase velocity mapping (PVM) in quantifying aortic blood flow from through-plane velocity measurements.

Materials and Methods: Two segmented PVM schemes were evaluated, one with seven lines per segment (seg-7) and one with nine lines per segment (seg-9), in twenty patients with cardiovascular disease. A non-segmented (non-seg) PVM acquisition was also performed to provide the reference data.

Results: There was agreement between the aortic flow curves acquired with segmented and non-segmented PVM. The calculated systolic and total flow volume per cycle from the seg-7 and the seg-9 scans correlated and agreed with the flow volumes from the non-seg scans (differences < 5%). Sign tests showed that there were no statistically significant differences (*P*-values < 0.05) between the segmented and the non-segmented PVM measurements. Seg-9, which was the fastest among the three sequences, provided adequate spatial and temporal resolution (> 10 phases per cycle).

Conclusion: Segmented k-space PVM shows great clinical potential in blood flow quantification.

Key Words: magnetic resonance imaging; magnetic resonance phase velocity mapping; segmented k-space; blood flow; flow quantification; aorta; velocity

MAGNETIC RESONANCE (MR) phase velocity mapping (PVM) has been widely used clinically to measure blood

velocity and flow. The technique is based on the principle that, by using the proper bipolar magnetic field gradients in the direction of interest, the velocity of the spins in that direction can be encoded within the phase of the detected signal (1). There have been many in vitro and in vivo studies evaluating PVM for flow quantification (2–17). For in vitro studies, the accuracy and precision of PVM was found to be consistently high, with errors lower than 10% (2–7). In vivo studies showed a good correlation between PVM and Doppler ultrasound or other flowmetric techniques (2,4,8–11). PVM has been extensively used to visualize and quantify cardiac, arterial, and venous blood flow with high precision (7,10–17).

Conventional PVM involves a non-segmented gradient-echo sequence with bipolar velocity-encoding gradients in the direction(s) of interest. The acquisition is prospectively or retrospectively ECG-gated to acquire a number of time-phases throughout the cardiac cycle. Only one line of k-space is acquired for each time-phase per cardiac cycle, resulting in a scanning time of several minutes (usually three to five minutes, depending on functional and imaging parameters) for acquisition of a single velocity direction in one imaging slice. Considering that PVM measurements constitute only part of a complete cardiac MR examination and that new diagnostic approaches have involved multi-slice and multidirectional velocity acquisitions (6,18,19), this non-segmented, single k-space line scheme becomes less practical clinically.

With the development of rapid imaging sequences, such as turbo gradient echo (TGE) and echo planar, faster PVM has become possible (20–31). Instead of a single k-space line, a segment of k-space lines are acquired per time-phase and cardiac cycle. As a result, the acquisition can be performed in seconds. Segmented TGE has shown good flow-behavior and can provide high temporal and spatial resolution acquisitions. It is, therefore, promising for reliable rapid flow quantification even in cases of moderately disturbed flow and vessel motion (25–31).

In a recent in vitro study (31), we showed that segmented k-space PVM provides accurate flow results under steady and pulsatile flow conditions (errors < 5%). Segmented PVM measurements with five, seven,

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and nine k-space lines per segment agreed very closely with non-segmented measurements under identical experimental conditions. Therefore, the aim of the present study was to clinically evaluate segmented PVM as a blood flow quantification technique, by measuring blood flow in the human ascending aorta. Of interest was the effect of increasing the number of k-space lines per segment on the accuracy of the results.

MATERIALS AND METHODS

MR PVM measurements were performed in the ascending aorta of twenty patients with cardiovascular disease (16 males, average age 61 years; four females, average age 73 years) using a 1.5 Tesla whole-body MRI scanner (Sonata, Siemens Medical Solutions, Erlangen, Germany) with a maximum gradient strength of 40 mTesla/m. Thirteen patients had aortic dilatation and aneurysms (six with aortic regurgitation) with mid-ascending aorta diameter 4.5–6.0 cm. Six patients had ischemic heart disease (two with mitral regurgitation, one with combined aortic stenosis/aortic regurgitation/mitral regurgitation). One patient had a left ventricular aneurysm. The study protocol was approved by the Institutional Review Board of the Cleveland Clinic Foundation.

Initial true-FISP (fast imaging with steady-state precession) scout images in all three spatial orientations (axial, coronal, and sagittal) showed the location of the ascending aorta. The through-plane velocity of blood was measured in an imaging slice placed perpendicular to the long-axis of the ascending aorta, at the level of pulmonary artery bifurcation. Three PVM acquisitions were performed, at the same location in each subject, using the following three sequences: (a) conventional non-segmented gradient echo (non-seg); (b) segmented TGE with seven k-space lines per time-phase and cardiac cycle (seg-7); and (c) segmented TGE with nine lines per time-phase and cycle (seg-9).

The non-seg scans were 120-180 seconds long, whereas the segmented scans were 10-16 seconds long, depending on the heart rate. Each of the segmented k-space scans was performed during a breathhold. In all sequences, the flow-sensitive and the flowcompensated data were acquired in an interleaved manner in the same cycle. The flip angle was 30°. The slice thickness was 5 mm. The scan percentage factor (percentage of k-space filled) was 75% for the non-seg scans, 55% for the seg-7 scans, and 56% for the seg-9 scans. The 75% rectangular field of view (FOV) was varied from $210 \times 280 \text{ mm}^2$ to $263 \times 350 \text{ mm}^2$, depending on patient size. Combination of the scan percentage factor with the rectangular FOV factor resulted in an acquired number of 144, 105, and 108 k_v -lines for the non-seg, seg-7, and seg-9 scans, respectively. The nominal voxel size varied between $1.5 \times 1.1 \times 5.0 \text{ mm}^3$ and $1.8 \times 1.4 \times 5.0 \text{ mm}^3$ for sequence (a), between $2.0 \times 1.1 \times 5.0 \text{ mm}^3$ and $2.5 \times 1.4 \times 5.0 \text{ mm}^3$ for sequence (b), and between $1.9 \times 1.1 \times 5.0 \text{ mm}^3$ and $2.4 \times 1.4 \times 5.0 \text{ mm}^3$ for sequence (c). Therefore, for a reconstructed matrix of 256 \times 256, the interpolated

voxel size varied from $1.1 \times 1.1 \times 5.0 \text{ mm}^3$ to $1.4 \times 1.4 \times 5.0 \text{ mm}^3$ in all sequences.

The shortest (2.3-3.0 msec) allowable echo time (TE) was used for all measurements. All acquisitions were prospectively ECG-gated, resulting in 19–35, 13–25, and 11–21 time-phases per cycle for the non-seg, seg-7, and seg-9 acquisitions, respectively. The temporal resolution of the segmented acquisitions was improved by using echo view-sharing (32). The resulting resolution was 30 msec, 40–46 msec, and 50–57 msec for the non-seg, seg-7, and seg-9 acquisitions, respectively. The time corresponding to each time-phase was the time of acquisition of the central line of the segment, since that line contained the most useful velocity information of the segment.

In the non-seg sequence, the repetition time (TR) was 15 msec. In seg-7 and seg-9, the time to acquire data for any one k_y -line (actual TR) varied between 5.2 msec and 5.6 msec, depending on imaging parameters such as FOV. The echo was acquired asymmetrically within the data acquisition period. The peak of the echo occurred at data-point 82 of 256 and the missing data points were zero-filled. The velocity encoding value was 250 cm/second.

Each PVM acquisition produced a series of magnitude and phase images. All images were transferred to a work-station (Ultra-10, SUN Microsystems, Inc., Palo Alto, CA). A FORTRAN program was used to examine the images for the presence of any residual phase offsets (5,33). When phase offsets were detected, they were removed based on a two-dimensional regression analysis model, and the phase values of the phase images were converted to velocity values (cm/second), based on their proportionality. The images were further processed using Transform (Version 3.4, Research Systems, Inc., Boulder, CO) to calculate the flow rates and flow volumes. The aortic cross-section was defined as the region of interest, and the velocity was integrated over the aortic cross-sectional area to determine the flow rate (mL/second) through the aorta for each timephase. Integration of the aortic flow curve over systole and over the entire cardiac cycle provided the systolic and total flow volumes (mL/cycle), respectively. When no phase offset was detected, the images were processed using manufacturer software (ARGUS Cardiac Analysis, Siemens Medical Solutions, Erlangen, Germany) in order to automatically calculate the flow rates and flow volumes.

Statistical analysis, including non-parametric sign tests, regression analysis, and t-tests, was performed to compare the flow data from the segmented acquisitions with the flow data from the non-segmented acquisitions. Most of the statistics was performed using Minitab (Version 13, Minitab, Inc., State College, PA).

RESULTS

Figure 1 shows representative magnitude and phase images from the non-seg and seg-9 PVM acquisitions from one subject at peak systole. Both sets of images are of sufficient quality for reliable analysis and flow quantification. Similar image quality was observed in all subjects for all three sequences used.



Figure 1. Magnitude (left) and phase (right) images acquired in the same subject using the non-segmented sequence (top) and the segmented sequence with nine lines per segment (bottom). Both sets of images are of sufficient quality for flow quantification analysis.

As expected, the seg-7 acquisitions provided higher temporal resolution than the seg-9 acquisitions (13–25 time-phases for seg-7 vs. 11–21 time-phases for seg-9); however, seg-9 provided greater scanning speed (13–16 seconds for seg-7 vs. 10–13 seconds for seg-9).

The flow curves obtained using the segmented acquisitions agreed with the curves obtained using the nonsegmented acquisitions, as seen in Figure 2 for four representative cases. In patients with aortic regurgitation, the flow curves showed diastolic reverse flow activity, as expected (e.g., subject 17, Fig. 2). Table 1 shows all of the flow volume results.

The results for the systolic and the total flow volumes from the segmented PVM acquisitions showed a close correlation (Pearson coefficients ≥ 0.94 , Table 2) and agreement (mean difference < 5%) with those from the non-segmented acquisitions. Sign tests showed no statistically significant differences (*P*-values > 0.05) between the segmented and the non-segmented PVM measurements (Table 2). Figures 3 and 4 show the regression analysis results from the comparison between the segmented and the non-segmented systolic and total flow volumes, respectively. In all cases, the regression lines have slopes not significantly different from 1.0, and intercepts not significantly different from zero (as statistically determined via t-tests with P-values > 0.05). In addition, the coefficients of determination are large $(r^2 > 0.88)$, and the standard errors small (Table 2). Good correlation (Pearson coefficient ≥ 0.97 , Table 2) and no statistically significant differences were also found between the seg-7 and seg-9 flow volume results (refer to the regression analysis and sign test results in Fig. 5 and Table 2).

DISCUSSION

Non-segmented MR PVM is an accurate and precise velocimetric and flow quantification technique, and has been used clinically to measure blood flow and characterize blood flow patterns throughout the cardiovascular system. Despite its advantages, it is relatively slow, requiring several minutes for acquisition of a single velocity component in a slice. With the development of segmented k-space acquisition schemes, rapid PVM has become available.

We recently evaluated the in vitro accuracy of segmented PVM with five, seven, and nine lines per segment under a variety of flow conditions (31). The results showed that segmented k-space PVM has high potential for blood flow quantification. Based on the positive in vitro results, the aim of the present study was to clinically evaluate the potential of segmented PVM to quantify blood flow in the aorta of human subjects. We did not evaluate the segmented scheme with five lines per segment clinically, because it would generally require more than 20 seconds per acquisition. A breath-hold of such duration may be difficult to achieve for many patients.

Due to the lack of a universally acceptable goldstandard blood flow quantification technique, in this study we considered non-segmented PVM as the reference technique. This can be justified by the large



Figure 2. Measured aortic flow waveforms in four subjects using the non-segmented sequence, and two segmented sequences with seven and nine k-space lines per segment. A good agreement can be observed in all cases between the three sequences.

number of previous evaluations of the non-segmented technique both in vitro and in vivo. Therefore, a comparison between segmented and non-segmented k-space PVM can provide important information about the reliability of segmented PVM as a flow quantification technique.

Image quality was good for all sequences used, facilitating image analysis and processing. The aortic crosssection could easily be identified for further analysis. Although the in-plane spatial resolution ($\cong 1-3 \text{ mm}^2$) was adequate to obtain the details of the velocity profile for a large vessel such as the ascending aorta, an even higher resolution (however, at the expense of scanning time) would be required for smaller vessels, as previously observed in vitro (31). Higher spatial resolution would also better resolve the velocity gradients immediately adjacent to the vessel wall (important to accurately calculate wall shear stress). However, the effects of reducing voxel size on the signal-to-noise ratio and, thus, image quality in vivo need to be thoroughly investigated.

The measured aortic flow curves in all subjects were very similar for all three sequences (non-seg, seg-7, and seg-9). It should be noted that, in seg-

Table 1 Flow Volume Results

Subject no.	Systolic volume, ml/cycle			Total volume, ml/cycle			
	Non-Seg	Seg-7	Seg-9	Non-Seg	Seg-7	Seg-9	
1	90.1	91.7	90.3	88.1	90.8	86.1	
2	41.8	41.1	42.9	36.6	39.2	36.0	
3	55.8	58.4	59.4	36.8	37.2	33.7	
4	67.8	67.3	69.4	64.6	60.2	66.8	
5	57.5	61.9	58.9	57.0	58.8	58.3	
6	45.3	49	47.9	44.2	50.1	54.2	
7	67.9	67.9	68.9	67.1	64.1	62.6	
8	57.9	59.4	57.3	59.1	43.9	48.7	
9	52.4	49.1	48.5	48.4	47.8	46.4	
10	54.8	55.9	57	53.7	52.7	53.3	
11	63.8	63.7	64.3	58.5	57.4	61.0	
12	72.4	66.1	69.6	77.7	76.1	82.4	
13	68	64	67.2	60.5	67.1	68.4	
14	37.2	40.2	37.2	40.7	36.1	42.7	
15	48.5	48.3	47.3	39.9	36.6	36.3	
16	71.9	70.9	73.9	69.8	67.3	73.2	
17	72.2	77	78.3	53.9	50.5	51.8	
18	72.6	67.8	74.2	53.5	56.8	56.5	
19	66.2	65.1	64	68.0	70.2	71.4	
20	70.5	75.8	72.3	53.8	58.6	57.2	

Non-Seg = non-segmented; Seg-7 = segmented with seven lines per segment; Seg-9 = segmented with nine lines per segment.

Table 2	
Regression Analysis and Other Statistics	

Y (mL/second)	X (mL/second)	Regression equation	R ²	SE (mL/second)	Mean % difference	Sign test <i>P</i> -value	Pearson correlation coefficient
Systolic flow volume							
Seg-7	Non-Seg	Y = 0.95 X + 3.38	0.937	3.2	0.8	1.0	0.97
Seg-9	Non-Seg	Y = 1.01 X - 0.14	0.970	2.3	1.1	0.2	0.99
Seg-9	Seg-7	Y = 1.03 X - 1.40	0.964	2.6	0.5	0.8	0.98
Total flow volume							
Seg-7	Non-Seg	Y = 0.99 X + 0.26	0.885	5.0	-0.8	0.8	0.94
Seg-9	Non-Seg	Y = 1.02 X - 0.57	0.903	4.7	1.2	0.8	0.95
Seg-9	Seg-7	Y = 1.00 X + 1.44	0.942	3.6	2.3	0.8	0.97

SE = standard error; Seg-7 = segmented with seven lines per segment; Seg-9 = segmented with nine lines per segment; Non-Seg=nonsegmented.

mented PVM with N lines per segment (where N is an odd number), the most influential information is contained in the central line of the segment, (N+1)/2, because this is the closest line of the segment to the center of k-space. Therefore, the time point corresponding to each cardiac phase was adjusted to be the time of acquisition of the central k-space line of the segment. As a result, the segmented techniques cannot acquire data at the very beginning of the cardiac cycle when they are prospectively ECG-gated (in this study, the first time phase in the cycle corresponded to 8, 40, and 50 msec after the QRS wave, for the non-seg, seg-7, and seg-9 acquisitions, respectively). This may cause uncertainties in measurements for patients with very rapid early systolic flow acceleration. Use of retrospective ECG-gating should eliminate this problem.

The seg-7 and seg-9 flow volume (systolic and total) results showed strong correlation and agreement with the non-seg flow volume results, suggesting that reliable blood flow quantification is possible using ultra-fast segmented PVM. Increasing the number of k-space lines per segment to nine did not adversely affect the ability to sufficiently resolve the flow waveforms. The seg-9 scans were faster than the seg-7 scans by approximately three seconds, and provided acquisitions with good temporal resolution (> 10 time phases per cycle). Increasing the number of k-space lines per segment above nine will certainly shorten the acquisition even more, but at the expense of temporal resolution. Reducing the temporal resolution may cause uncertainties in determining local velocity values such as peak systolic velocities. Therefore, the number of lines should be further increased only



100 Non-Seg Systolic Volume (ml/beat)



Figure 3. Comparison between the systolic flow volumes measured with the non-segmented and the two segmented sequences (non-Seg: non-segmented acquisition; Seg-7: segmented acquisition with seven k-space lines per segment; Seg-9: segmented acquisition with nine k-space lines per segment). The diagonal line is the line of identity.

Figure 4. Comparison between the total flow volumes measured with the non-segmented and the two segmented sequences (non-Seg: non-segmented acquisition; Seg-7: segmented acquisition with seven k-space lines per segment; Seg-9: segmented acquisition with nine k-space lines per segment). The diagonal line is the line of identity.



Figure 5. Comparison between the flow volumes (systolic and total) measured with the two segmented sequences (Seg-7: segmented acquisition with seven k-space lines per segment; Seg-9: segmented acquisition with nine k-space lines per segment). The diagonal line is the line of identity.

after assuring acquisition of an adequate number of time phases per cycle to prevent loss of valuable information and evaluating the acquisition technique in vitro and in vivo.

In conclusion, segmented k-space magnetic resonance phase velocity mapping in the human ascending aorta produced flow results that agreed with those from non-segmented acquisitions. Increasing the number of lines per segment to nine allowed very fast acquisitions (< 15 seconds) with adequate temporal resolution (less than 60 msec between successive time phases). Segmented k-space MRPVM shows great clinical potential in blood flow quantification.

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