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Early Hemodynamic Response Assessment of Stereotactic Radiosurgery for a Cerebral Arteriovenous Malformation Using 4D Flow MRI

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ABSTRACT

SUMMARY: Brain AVMs treated with stereotactic radiosurgery typically demonstrate a minimum latency period of 1–3 years between treatment and nidus obliteration. Assessment of treatment response is usually limited to evaluation of AVM nidus structural changes using conventional MR imaging and MRA techniques. This report describes the use of 4D Flow MRI to also measure radiation-induced hemodynamic changes in a Spetzler-Martin grade III AVM, which were detectable as early as 6 months after treatment.

ABBREVIATION: SRS = stereotactic radiosurgery

Stereotactic radiosurgery (SRS) is an effective treatment option for brain arteriovenous malformations that are not amenable to microsurgical resection or endovascular embolization. Nidus obliteration is thought to result from radiation-induced endothelial cell proliferation causing progressive vessel wall thickening and eventual occlusion.¹ When successful, this process typically results in a latency period of 1–3 years between treatment and nidus obliteration,² though the nidus can persist in approximately 20% of AVMs even 5 years after treatment.³

Although MR imaging and MRA have widely supplanted DSA as the mainstay modalities for following AVMs posttreatment, prior studies evaluating their ability to assess the response to SRS have only examined AVM structure, specifically nidus volume.^{4–8} Advanced MRA techniques capable of measuring intracranial flow dynamics, such as 4D Flow MRI (<https://www.arterys.com/4d-flow>), have not yet been leveraged to study SRS treatment effects on AVMs.⁹

4D Flow MRI acquires time-resolved phase-contrast data with 3D velocity-encoding throughout the cardiac cycle, pro-

ducing a 4D dataset of velocity vectors. This 4D dataset can be retrospectively analyzed to measure flow in any direction, as well as to calculate numerous other fluid dynamic metrics such as velocity, pressure gradients, vorticity, wall shear stress, turbulence, and so forth.¹⁰ Flow vectors can also be rendered in 3D in various display formats for qualitative visualization.¹¹

We report imaging and hemodynamic data from serial 4D Flow MRI of a Spetzler-Martin grade III AVM treated with SRS as a proof-of-concept application of 4D Flow, and we compare this technique with traditional MR imaging/MRA.

MATERIALS AND METHODS

A 19-year-old man was diagnosed with an unruptured 3.5 × 2.9 × 2.5 cm Spetzler-Martin grade III AVM in the medial left temporal lobe on a trauma work-up. Conventional angiography showed primary arterial supply from the left MCA with secondary contribution from the left posterior cerebral artery (Fig 1A, -B). Venous drainage was solely via the left basal vein of Rosenthal (Fig 1C). The nidus was treated with single-fraction SRS to a prescription dose of 1800 cGy (mean dose, 2000 cGy; maximum, dose 2200 cGy) using a linear accelerator–based volumetric arc therapy technique. The patient experienced no acute or subacute complications.

Baseline MR imaging and serial follow-up MR imaging were performed at 1, 6, 12, and 20 months after SRS. All scans were obtained on a 3T Discovery 750 scanner (GE Healthcare, Milwaukee, Wisconsin), including simple 4-point-encoded 4D Flow MRI with variable-density Poisson disc undersampling,^{11,12} with E-SPiRiT (eigenvector-based iTerative Self-consistent Parallel Imaging Reconstruction) combined parallel imaging and compressed-sensing image reconstruction.^{13,14}

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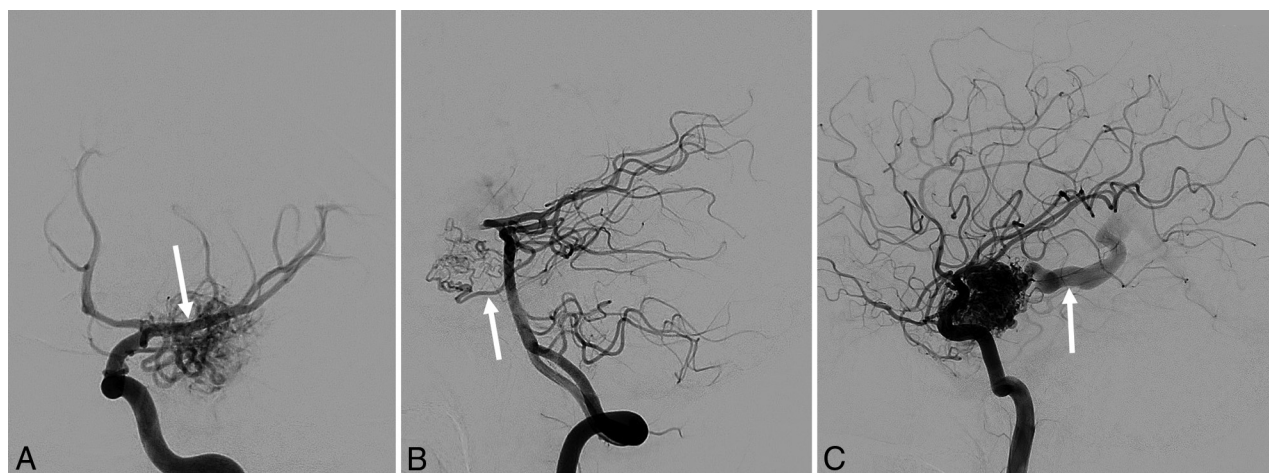


FIG 1. Conventional angiography of a left temporal lobe AVM. Early arterial phases show primarily MCA supply (arrow) on a left frontal oblique left ICA injection (A), with secondary posterior cerebral artery supply via a small branch artery (arrow) on a lateral left vertebral artery injection (B). Late arterial phase (C) shows early opacification of the left basal vein of Rosenthal, the exclusive venous drainage (arrow).

4D Flow MRI acquisition parameters

Parameter	
Contrast volume	15–20 mL of gadobenic acid
Voxel size	$1.12 \times 0.98 \times 1.60$ to $1.16 \times 1.02 \times 1.80$ mm
Flip angle	15°
TR	4.96–5.05 sec
TE	2.40–2.57 ms
Scan time	7 min 45 sec to 9 min 3 sec
Velocity encoding	100–200 cm/s
Bandwidth	62 kHz

The combination of variable-density Poisson undersampling and E-SPiRiT reconstruction provides the mechanism for compressed sensing, which preserves signal to noise while reducing 4D Flow acquisition time. The parallel imaging acceleration factor was 3.6 overall (1.8 in phase-encode and 2.0 in slice-encode directions). Additional specific acquisition parameters are listed in the Table. At each time point, blood flow (liters/minute) was measured in bilateral distal ICAs, bilateral proximal MCAs, and the draining left basal vein of Rosenthal.

RESULTS

In the 20 months following SRS, the patient's left temporal lobe AVM nidus gradually decreased in size from approximately $3.5 \times 2.9 \times 2.5$ cm to $2.0 \times 1.5 \times 1.3$ cm, as measured on 3D time-of-flight MRA. Time-resolved color-coded 3D volumetric renderings of 4D Flow velocity data demonstrated visibly decreasing flow velocity in the left MCA, the dominant arterial supply, denoted by decreasing red and yellow velocity vectors and increasing green velocity vectors (Fig 2). Video renderings of 4D Flow data collected throughout the cardiac cycle are available in the On-line Videos.

Blood flow measurements performed using 4D Flow data initially showed asymmetrically greater flow to the left MCA and ICA compared with the contralateral side 1 month posttreatment, which began to normalize by the 6-month follow-up examination. In contrast, flow in the right anterior circulation slightly increased between the 1- and 6-month follow-ups.

By the 20-month follow-up scan, flow in the left MCA and ICA

had decreased by 51% (0.51 to 0.25 L/min) and 34% (0.59 to 0.39 L/min) compared with initial values, respectively, while the net change in right MCA and ICA flow was <11% (Fig 3A). Blood flow in the draining vein also decreased with time, from 0.25 to 0.07 L/min by the 20-month follow-up, a net reduction of 80% (Fig 3B).

DISCUSSION

Treatment response following SRS therapy for AVMs is most commonly assessed with conventional MR imaging and MRA techniques, which are tailored to evaluate brain parenchyma and vascular anatomy. This case of a patient with an unruptured AVM treated with curative SRS demonstrates the ability of 4D Flow MRI to quantify radiation therapy-induced hemodynamic changes that are not captured by traditional anatomic imaging modalities.

While radiation-induced endothelial hyperproliferation typically results in a 1- to 3-year latency period between SRS and AVM nidus obliteration, we quantified a decrease in blood flow through a left temporal lobe AVM as early as 6 months after treatment, previously unreported using MRA techniques. We also quantified an increase in flow to the contralateral anterior circulation at the 6-month follow-up, which may suggest reversal of a left-to-right steal phenomenon as flow to the AVM decreased.¹⁵ These findings underscore the ability of 4D Flow MRI to quantify flow dynamics across the entire brain in 1 scan and represent the first application of this technique in the evaluation of SRS treatment effects on AVMs.¹⁶

Future studies will follow cohorts of patients to explore the relationship between flow dynamics measured by 4D Flow MRI and long-term outcomes, such as hemorrhage risk, length of latency period, and rate of eventual obliteration. Additional hemodynamic measures, such as intravascular pressure gradients, turbulence, and wall shear stress can be examined as well. Flow-mapping capabilities could evaluate changes in flow connectivity in complex AVMs with multiple feeding and draining vessels and perhaps help to further optimize SRS treatment targets.¹⁷

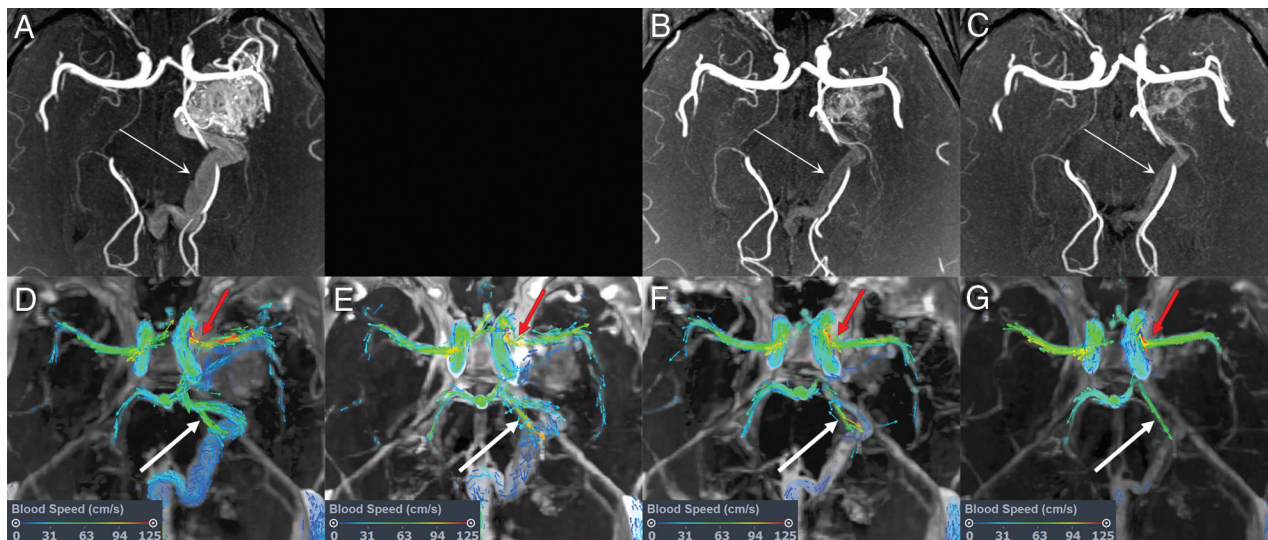


FIG 2. Conventional 3D time-of-flight MRA (upper row) and 4D Flow MRI (lower row) during a 20-month follow-up course after SRS. Scans were acquired at 1 (A and D), 6 (E), 12 (B and F), and 20 (C and G) months following SRS (MRA was not performed at the 6-month follow-up). Conventional sequences show a gradual decrease in size of the left temporal lobe AVM nidus, with decreasing distention of the draining vein (thin white arrow). 4D Flow MRI 1 month after SRS (D) shows asymmetrically greater flow in the left MCA (thick red arrow) and posterior cerebral artery feeding vessels (thick white arrow), as well as in the distended draining vein. The left MCA flow velocity gradually decreases with time (D–G).

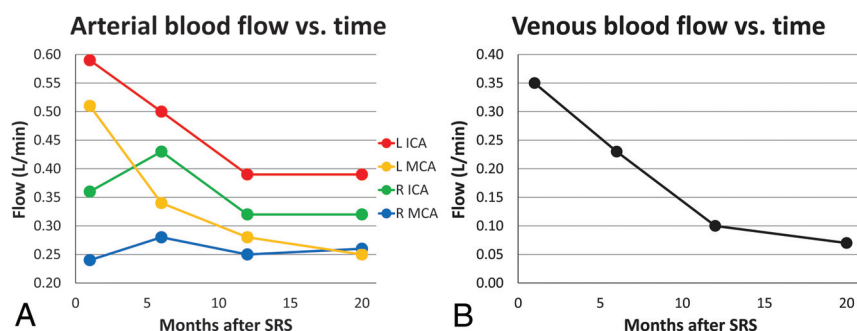


FIG 3. Blood flow in the anterior circulation (A) and draining vein (B) measured by 4D Flow over a 20-month follow-up after SRS. At the 6-month follow-up, arterial flow decreased most dramatically in the left MCA. Flow in the left MCA, left ICA, and draining vein continued to decrease throughout the posttreatment course.

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