

ORIGINAL RESEARCH

Effect of aneurysm and ICA morphology on hemodynamics before and after flow diverter treatment

Ignacio Larrabide,^{1,2,3} Arjan J Geers,^{1,2} Hernán G Morales,^{1,2} Martha L Aguilar,^{1,2} Daniel A Rüfenacht⁴

► Additional material is published online only. To view please visit the journal online (<http://dx.doi.org/10.1136/neurintsurg-2014-011171>).

¹Networking Biomedical Research Center on Bioengineering, Biomaterials and Nanomedicine (CIBER-BBN), Barcelona, Spain
²Center for Computational Imaging & Simulation Technologies in Biomedicine (CISTIB), Universitat Pompeu Fabra, Barcelona, Spain
³PLADEMA-CONICET and Universidad Nacional del Centro, Tandil, Buenos Aires, Argentina
⁴Hirslanden Clinic, Zürich, Switzerland

Correspondence to

Dr I Larrabide, PLADEMA-CONICET, Universidad Nacional del Centro de la Provincia de Buenos Aires, c/Pinto 399, B7000GHG Tandil, Buenos Aires 7000, Argentina; ignacio.larrabide@gmail.com

Received 11 February 2014

Revised 5 March 2014

Accepted 7 March 2014

ABSTRACT

Background Flow diverter (FD) treatment aims to slow down blood flow inside the aneurysm and increase the average time that blood resides in the aneurysm.

Objective To investigate the relationship between vessel and aneurysm morphology and their influence on the way in which braided FDs change intra-aneurysmal hemodynamics.

Materials and methods Twenty-three patient-specific intracranial aneurysm models at the supraclinoid segment of the internal carotid artery were studied. Vessel and aneurysm morphology was quantified and blood flow was modeled with computational fluid dynamics simulations. The relation between morphologic variables and the hemodynamic variables, *WSS* (wall shear stress) and *totime* (ratio between the aneurysm volume and inflow at the aneurysm neck), was assessed statistically.

Results Intra-aneurysmal flow was less dependent on the vessel than on aneurysm morphology. In summary, after treatment with a FD, a greater aneurysm flow reduction and redirection to the vessel main stream should be expected for (a) aneurysms located further away from the curvature peak, (b) aneurysms on the inner side of the bend, (c) aneurysms with no proximal stenosis, and (d) larger aneurysms.

Conclusions Although the change in intra-aneurysmal hemodynamics after FD treatment strongly depends on the morphology of the aneurysm, the hemodynamic effect of a FD is also linked to the parent vessel morphology and the position and orientation of the aneurysm with respect to it.

INTRODUCTION

The study of intracranial aneurysm hemodynamics has grown considerably in recent years. Hemodynamics and morphology have proved to be powerful indicators of aneurysm rupture.^{1 2} The pioneering work of Ujiie *et al* showed a link between aneurysm morphology and local hemodynamics.³ Owing to its close link to hemodynamics, aneurysm morphology has often been considered when planning treatment.^{2 4-7} Studies have shown that local morphology and configuration of the aneurysm relative to the parent vessel also play a key role in intra-aneurysmal hemodynamics and should be considered.^{8 9} For instance, Passerini *et al* concluded that internal carotid arteries (ICAs) with “double bends” are less prone to develop intracranial aneurysms.¹⁰

Commercially available, braided flow diverters (FDs) provide a new alternative endovascular approach to treating aneurysms that are otherwise too risky or difficult to manage with conventional techniques.¹¹ Although animal experiments and clinical studies have already demonstrated the impressive effectiveness of FDs, their effects on local hemodynamics are not yet fully understood.^{12 13} Nevertheless, delayed and complete angiographic occlusion is reported in most cases.¹⁴⁻¹⁶ Recent studies estimated treatment success based on angiographic images, and used computational fluid dynamics (CFD) to investigate the relation between aneurysm size and shape and flow reduction after FD insertion.^{17 18} CFD provides a non-invasive way of studying and predicting the effect of these devices on local hemodynamics.^{19 20}

The combined study of aneurysm and vessel morphology in relation to FD treatment is relevant, but not yet developed. The purpose of this study is to provide a better insight into the relation between vessel and aneurysm morphologic variables, and intra-aneurysmal hemodynamic changes produced with braided, closed cell design FD treatment.

MATERIALS AND METHODS

Twenty-three aneurysm models were used in this study. This population was defined as explained below and was drawn from the @neurIST database. It was collected from different centers and processed by different observers following the same data protocol.²¹ Anatomic models were obtained from diagnostic three-dimensional rotational angiography images, acquired through an Integris Allura System (Philips Healthcare, Best, The Netherlands) or an AXIOM Artis (Siemens Medical Solutions, Erlangen, Germany). Voxel sizes in the reconstructed 3D images ranged from 0.208×0.208×0.208 mm³ to 0.378×0.378×0.378 mm³. These data correspond to those of a previous study we carried out, but where the questions asked and hypotheses were different.²²

For an initial set of 40 aneurysms located at the supraclinoid segment of the ICA, three clinicians independently selected the most appropriate endovascular treatment. The criteria followed were (1) the absence of branching arteries (typically an ophthalmic artery or anterior choroidal artery) near to the aneurysm that could be occluded by the device and (2) an aneurysm that was suboptimal for treatment with coils because of its morphology and/or

To cite: Larrabide I, Geers AJ, Morales HG, *et al*. *J NeuroIntervent Surg* Published Online First: [please include Day Month Year] doi:10.1136/neurintsurg-2014-011171

neck width. A subset of 23 aneurysms was chosen for treatment with FD by at least two clinicians. The remaining aneurysms (17) were considered more suitable for conventional treatment by two or more clinicians, and thus excluded from the study. Eleven aneurysms (48%) were located at the ophthalmic artery, nine (39%) at the posterior communicating artery and three (13%) at the anterior choroidal artery. Aneurysm sizes ranged between 1.3 and 22 mm. Seventy per cent of patients were female. Patient age was 50 (± 8) years at the time of aneurysm discovery. [Figure 1](#) presents the 21 ICAs studied.

Images were segmented using a geodesic active regions method, and a 3D triangulated surface model was generated.²³ Triangle removal, hole filling and smoothing to preserve volume were employed to remove imperfections in the vascular models. Geometric models of the FDs, consisting of 48 wires on a single layer with a strut thickness of 60 μm were created and deployed in the vascular models using a virtual stenting method.²⁴ The braiding angle was 60° in all cases. This was used as a reference and not intended to match any particular device available in the market. Although the braiding angle is preferably preserved after deformation, it is not forced by the deployment algorithm and might vary depending on the position on the stent mesh and the vessel/aneurysm geometry treated.

Aneurysm morphologic characterization

AngioLab software (Universitat Pompeu Fabra, Barcelona, Spain), which has been proved to have minimal interobserver variability, was used for manual isolation of the aneurysms.^{25 26} Morphologic variables were automatically computed from the isolated aneurysm dome ([figure 2A](#)). Only the morphologic variables most often reported and associated with aneurysm rupture and treatment planning, were selected for analysis ([table 1](#)).^{3 4}

Vessel morphologic characterization

The vasculature surrounding the aneurysm was characterized using the Vascular Modeling Toolkit ([figure 2B,C](#)). The morphologic characterization was based on the work of Piccinelli *et al.*^{27 28} The vessel morphology was characterized by three variables—namely, d^{peak} ([figure 2C](#)), α^n ([figure 2D](#)) and α^t

([figure 2E](#)), which are described in [table 1](#). Further details on the computation of these variables can be found online as supplementary content (see online file supplementary content_1).

CFD modeling/simulation

Volumetric meshes, with unstructured tetrahedral and eight-node prism elements, were generated using the commercial software ICEM CFD, V12.1 (ANSYS, Canonsburg, Pennsylvania). To ensure CFD simulations independent from mesh element size, a mesh independency analysis was carried out. The convergence criterion of mesh independence was that variables of interest—namely, wall shear stress (WSS), inflow at the aneurysm neck and intra-aneurysmal velocity had to be within 2.5% error from the finest tested mesh. Convergence was reached with an element size around the stent strut of 0.016 mm (ie, about 12 elements along the perimeter), three prism layers with a total height of 0.3 mm defined everywhere but in the region of the FDs, and a global tetrahedral element size of 0.3 mm. The total number of mesh elements ranged between 0.4 and 3.4 million elements for the untreated cases and between 2.6 and 15.2 million elements for the treated ones.

Following previous studies and because we were interested only in the effect of the FD inside the aneurysm, the portions of the stent lying on the vessel wall were removed to reduce mesh size and computational time, and only the portion of the stent covering the aneurysm neck was kept.¹⁹

Unsteady CFD simulations were created with CFX, V12.1 (ANSYS). Blood was modeled as an incompressible Newtonian fluid (density=1066 kg/m³, viscosity=0.0035 Pa·s). The vessel wall was assumed to be rigid with a no-slip boundary condition. A flow rate waveform was imposed at the inlet, and pressure waveforms at the outlets, all derived from a 1D mathematical model of the systemic arterial tree.²⁹ A straight extension was added to the vascular model inlet and a flat velocity profile was imposed at the inlet of the extension. The combination of the extension and the considered vascular region causes the effects of the imposed velocity profile to vanish at the location of the aneurysm.³⁰ The shape and average flow rate of the waveforms ([figure 3](#)) were the same for all cases so that geometrical effects stood out. Cardiac

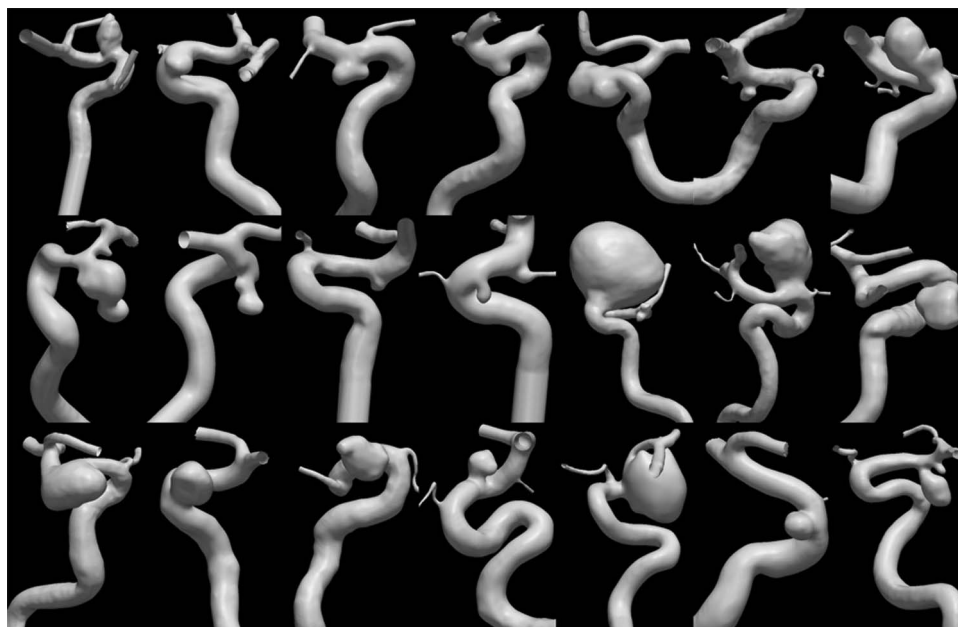


Figure 1 Visualization of the 23 aneurysms used in this study. Two cases present two aneurysms.

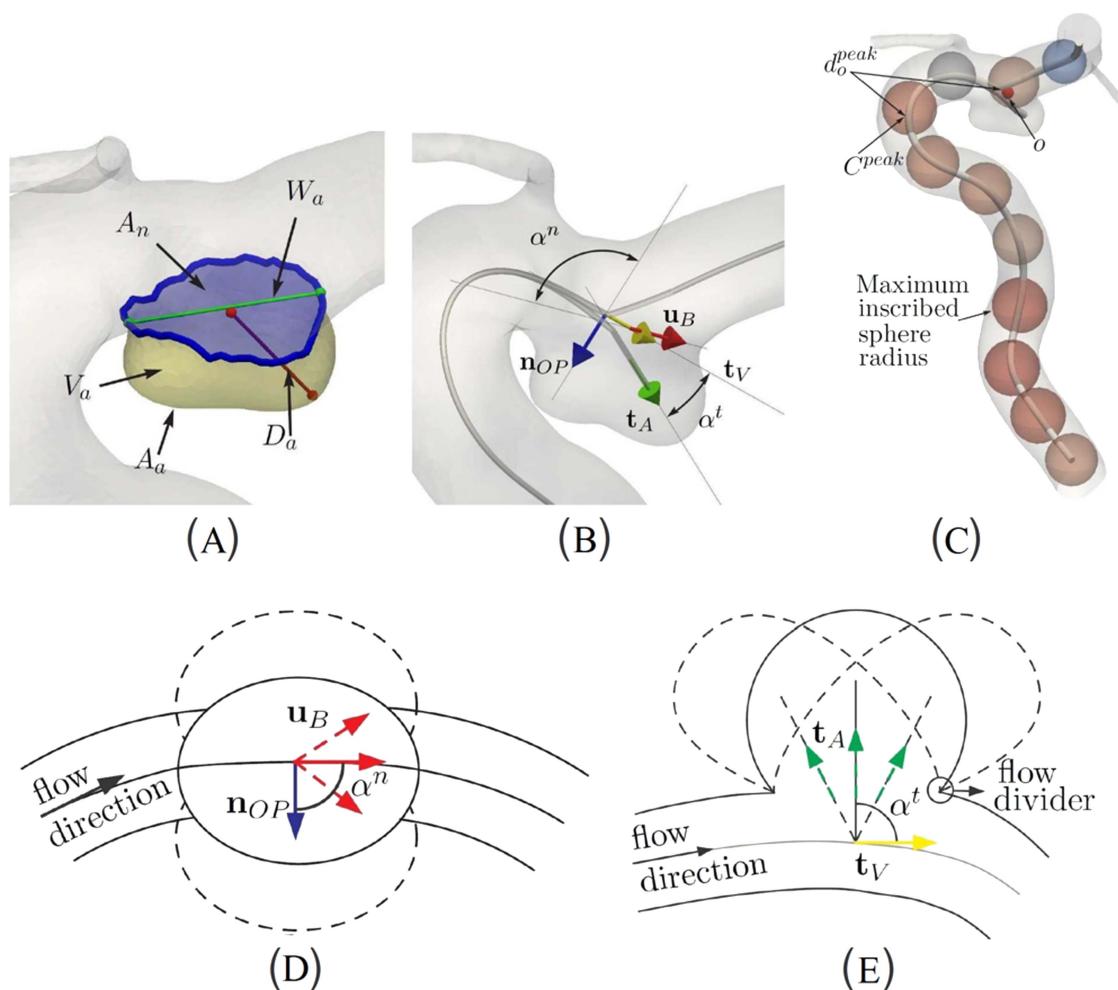


Figure 2 (A) Graphical representation of the aneurysm sac morphologic variables (detailed in [table 2](#)). (B) Local vectors describing the aneurysm orientation with respect to the parent vessel. n_{OP} is the normal vector of the local osculating plane and pointing towards the bifurcation apex at zero abscissa, and t_A is defined as the centerline tangent direction, one inscribed sphere radius away from the zero abscissa along the aneurysm centerline into the aneurysm. Also, $\alpha^n = \alpha(n_{OP}, u_B)$ and $\alpha^t = \alpha(t_V, t_A)$ are indicated. (C) Graphical representation of the vessel morphologic variables studied. C^{peak} indicates the location of the curvature peak before the aneurysm, o is the bifurcation reference system origin, d^{peak} is the distance between these two along the centerline. Maximum inscribed sphere radius, which is an approximation of the vessel radius, is shown as semi-transparent spheres along the centerline. (D) Schematic view of the aneurysm with the associated n_{OP} and the change of u_B (three positions) as the location of the aneurysm on the parent vessel changes. (E) Schematic view of the aneurysm showing t_V and the change of t_A (three positions) as the orientation of the aneurysm with respect to parent vessel changes. The aneurysm bifurcation apex is also indicated.

cycles of 0.8 s each were discretized in time steps of 0.005 s. To reduce initial transients, three complete cardiac cycles were computed and data of the third were analyzed.

Data analysis

The hemodynamic variables—namely, WSS and *totime*, were studied and correlated with aneurysm and vessel morphologic variables. WSS was averaged in space and time on the aneurysm wall.^{30 31} *totime* is defined as the ratio between the aneurysm volume (V_a) and inflow at the aneurysm neck (Q_a).^{32 33} These variables were recorded before ($x_{untreated}$) and after ($x_{treated}$) virtual treatment. All hemodynamic changes (x_{change}) between untreated and treated models reported in this work were calculated as

$$x_{change} = x_{untreated} - x_{treated} \quad (1)$$

where a negative change in the hemodynamic variables indicates an effective increase after the insertion.

Data analysis was performed using the R statistical software package.³⁴ Because data were non-parametric, Spearman's correlation test was used and the coefficient ρ is reported. For all tests, statistical significance was considered for $p < 0.05$ (indicated with *). Correlations were recorded between morphologic variables and both untreated hemodynamics and hemodynamic change due to FD insertion (Eq. 1).

The mean porosity of the FDs (ϕ_{FD}) was quantified by measuring the voids of the FD mesh from an image of the aneurysm neck for each case.

RESULTS

Figure 1 shows the 21 cases hosting 23 aneurysms used in this study. Mean, SD, minimum and maximum of all morphologic variables are reported in [table 2](#). As supplemental content of this article, is provided a text file with All data used for the data analysis can be found online in file supplementaryContent_2-data.

New devices

Table 1 Aneurysm and vascular morphology variables observed and analyzed

Index	Description
Aneurysm morphology, measured on the extracted aneurysm dome surface	
D_a	Aneurysm depth, measured from the neck centroid to the furthest point on the dome
A_a	Aneurysm dome area
V_a	Aneurysm volume
W_a	Aneurysm neck maximum width
A_n	Aneurysm neck area, defined as the surface of the aneurysm ostium
AR_a	Aspect ratio, defined as D_a/W_a
Vessel morphology, measured on the parent vessel of the aneurysm	
d^{peak}	Describes the position of the aneurysm in the parent vessel, is the distance from o to the peak curvature C^{peak} before the aneurysm along the centerline
α^n	Describes the orientation of the aneurysm, is the angle between the local osculating plane normal n_{OP} on the parent vessel at zero abscissa and the bifurcation reference system up normal u_B
α^t	Describes the orientation of the aneurysm, is the angle between the tangent t_V at the parent vessel at zero abscissa and the aneurysm direction t_A
Stent porosity, measured across the neck of the aneurysm	
ϕ_{FD}	Porosity of the flow diverter (FD) measured on the patch covering the neck as $100 \times (\text{area of voids})/(\text{total area})$

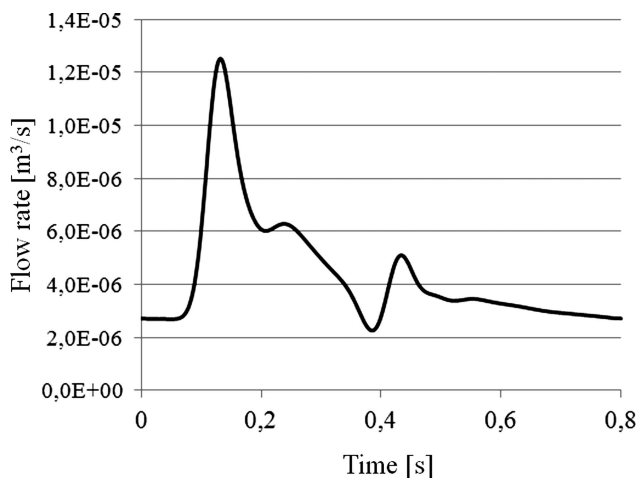
Morphology and hemodynamics in ICA aneurysms

Table 3 summarizes the correlation results between morphologic and hemodynamic variables.

Statistically significant correlations were found between *totime* and aneurysm morphological variables (D_a , A_a , V_a , A_n , and AR_a , with $\rho=0.52$ to $\rho=0.86$). WSS was found to decrease ($\rho=-0.42$ to $\rho=-0.50$) with increasing D_a , A_a , V_a , and A_n .

totime correlated significantly with α^n ($\rho=-0.73$) and α^t ($\rho=-0.72$). This means that when the bifurcation apex is towards the outer side of the bend (larger α^n), *totime* is smaller. As the aneurysm leans upstream (larger α^t), *totime* is also found to be smaller.

Three cases presented four aneurysms with WSS >10 Pa (see figure 4, one aneurysm in case A, two aneurysms in case B and one aneurysm in case C), which is an order of magnitude above physiological values.³⁵ A severe stenosis (~70%) before the

**Figure 3** Flow rate waveform used at the inlets of all the models. Inlet boundary conditions were imposed at the supraclinoid segment of the internal carotid artery.**Table 2** Aneurysm and vessel morphologic variables summary. All variables are expressed as mean±SD, minimum and maximum

Variable (unit)	Mean±SD	(min, max)
Aneurysm morphology		
D_a (mm)	6.16±4.56	(0.98, 20.60)
A_a (mm ²)	179.55±301.3	(3.68, 1413.69)
V_a (mm ³)	409.07±1079.15	(0.86, 5179.40)
A_n (mm ²)	23.74±26.30	(1.58, 98.65)
AR_a (-)	1.16±0.65	(0.33, 2.73)
Vessel morphology		
d^{peak} (mm)	-9.62±5.20	(-18.50, -2.56)
α^n (degrees)	51.2±25.4	(12.6, 106.0)
α^t (degrees)	64.6±24.5	(30.1, 127.9)

aneurysms was found in two of them (cases B and C). In these cases, upstream stenosis induced an increase in flow velocity (see streamlines in figure 4), which increased WSS and decreased *totime*. The case without stenosis (case A figure 4) corresponds to a small aneurysm where the main flow jet on the parent vessel penetrates the aneurysm. As counter examples, cases D, E, and F shown in figure 4 correspond to median WSS for this population (~3.10 Pa). None of these presented a severe stenosis.

Figure 5 presents streamlines and velocity contours at peak systole for cases A to F both before and after the FD insertion. It can be seen that although the jet going into the aneurysm is dumped after the insertion of the FD, the flow jet direction and the vortex pattern inside the aneurysm remain almost identical.

Figure 6 presents a scatterplot relating depth (D_a) and α^t , showing that larger aneurysms tend to lean downstream (smaller α^t). Additionally, larger aneurysms present larger *totime* (table 3). Moreover, it was found that aneurysms with smaller α^t will have smaller *totime* (confirmed by $\rho=-0.72$ for α^t in table 3).

Morphology in relation to hemodynamic changes after FD insertion

Table 4 presents correlations between morphologic variables, stent porosity, and reduction of hemodynamic variables (Eq. 1) after FD insertion.

FD insertion increased *totime* for all cases and its increment was larger for higher values of the aneurysm morphologic variables. WSS reduction inside the aneurysm was found to significantly correlate to all aneurysm morphologic variables ($\rho=-0.49$ to $\rho=-0.55$) except for AR_a .

When the aneurysm bifurcation apex lies towards the outer side of the vessel (larger α^n), the FD is less effective in reducing WSS and increasing *totime*. When aneurysms lean upstream (larger α^t), the increment in *totime* is larger.

The mean±SD porosity ϕ_{FD} among all cases was 72.83 ± 5.47 , ranging from 59.51% and 80.85%. Figure 7 presents two cases with high porosity (top row), corresponding to larger vessel diameters. The two cases with low porosity (bottom row) correspond to the vessels with small diameter, which presented stenosis as shown in figure 4 and a denser FD mesh across the aneurysm neck. For smaller vessel diameters, the FD mesh is denser locally, resulting in a lower porosity. Images have different scales, but the thickness of stent struts is the same in all cases.

Significant correlation ($\rho=-0.53$) was found between ϕ_{FD} and WSS. Increasing FD porosity produced less reduction of WSS. Higher WSS reduction was seen for cases with elevated WSS

Table 3 Morphologic versus pretreatment hemodynamic variables in ICA aneurysms correlations

	Aneurysm morphology					Vessel morphology		
	D_a	A_a	V_a	A_n	AR_a	d^{peak}	α^n	α^t
WSS	-0.49*	-0.50*	-0.49*	-0.42*	-0.36	-0.33	0.29	0.36
totime	0.86*	0.82*	0.82*	0.52*	0.73*	0.33	-0.73*	-0.72*

*Indicates statistically significant correlation ($p < 0.05$).
ICA, internal carotid artery; WSS, wall shear stress.

(>10 Pa) before FD insertion, which corresponded to stenosed cases also presenting low FD porosity.

DISCUSSION

The 23 cases in this study presented a wide range of sizes (D_a from 0.98 mm to 20.60 mm) and shapes (both saccular and fusiform aneurysms). CFD analysis was performed on these cases before and after the virtual implantation of a FD, representing, braided, closed cell devices of an average porosity of 72.83 ± 5.47 , ranging from 59.51% to 80.85%. Flow is highly dependent on geometry, and the aneurysm morphology (neck area, aneurysm volume, etc) has a stronger influence on intra-aneurysmal flow than vessel morphology. Sato *et al* observed that a change in aneurysm position and orientation

relative to the parent vessel induces substantial changes in aneurysmal flow patterns.³⁶

A computational study, as presented here, is primarily oriented to analyze the effect under controlled conditions (ie, inflow, FD design). It benefits from these controlled conditions, enabling exploration of the proposed relationship between geometry and FD effect on hemodynamics.

The results obtained in this part of the study reproduce previous research, on aneurysm morphology and hemodynamics, which proves the correctness of the methodology.^{3-5 10 28 36 37}

This same methodology was used to make a link between aneurysm and vessel morphology with hemodynamic changes induced by FD treatment, which is the main contribution and novelty of this work.

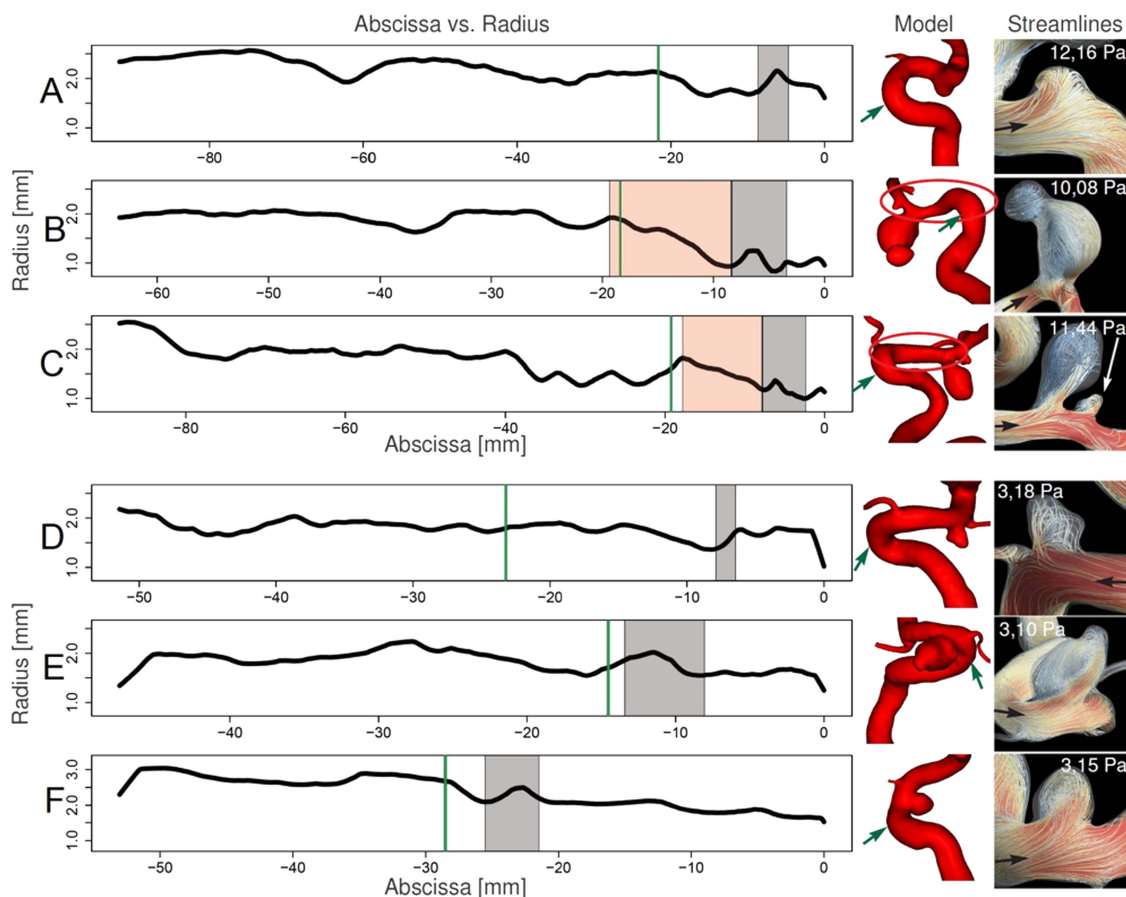
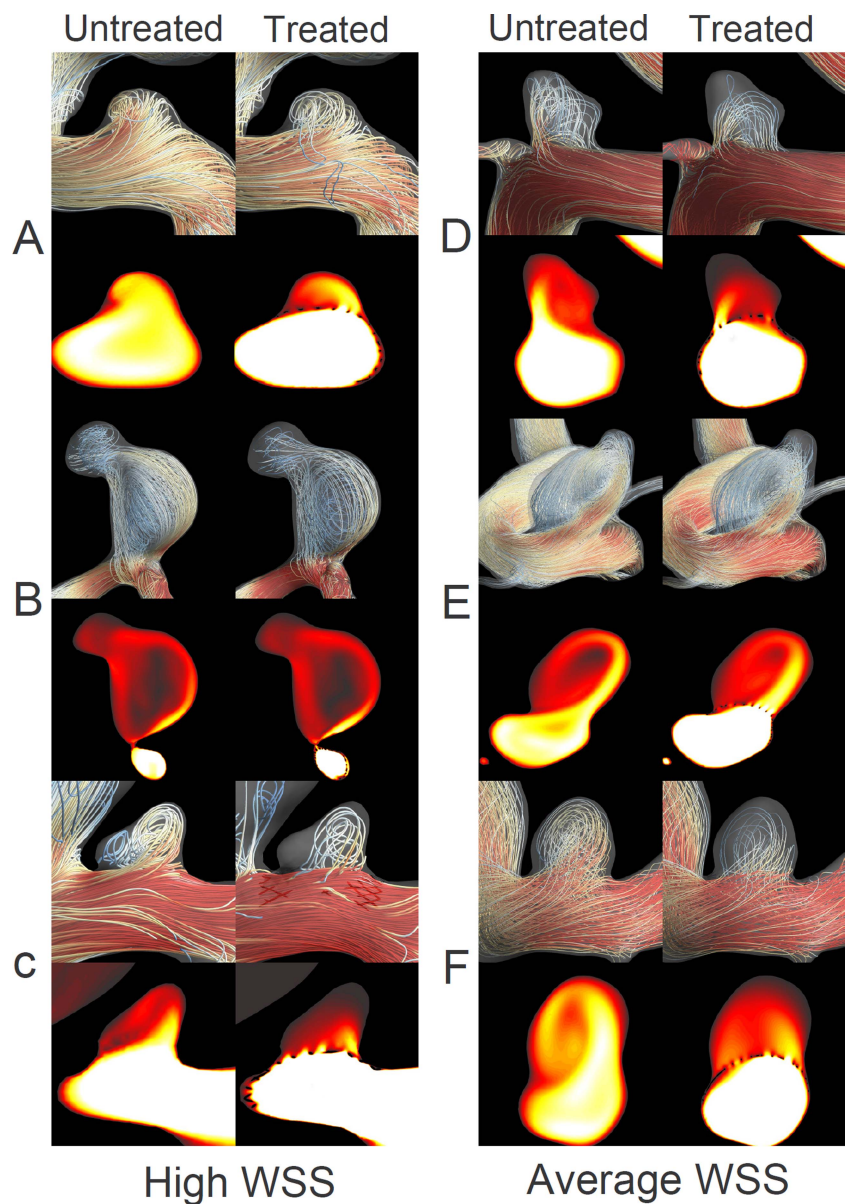


Figure 4 Position on the centerline abscissa versus maximum inscribed sphere radius. The green arrow indicates the peak curvature location. Aneurysm location is indicated in gray. A, B, and C are cases with high wall shear stress (WSS). D, E, and F are the three cases around the median WSS. The red region indicates the location of an acute stenosis with a fourfold reduction of cross-sectional area. On the right, are presented the geometrical models of each case with an oval highlighting the stenosis region for cases B and C and a green arrow indicating the location of the curvature peak. The black arrow indicates the direction of flow and the WSS before flow diverter insertion is also indicated for each case.

Figure 5 Streamlines and velocity contours at peak systole for cases A to F presented in [figure 4](#) both, before and after insertion of the flow diverter. The main jet stream entering the aneurysm is dumped but its direction and the vortex structure inside the aneurysm are not altered after the treatment. *WSS*, wall shear stress.



Hemodynamics in ICA aneurysms in relation to morphology

The *WSS* was found to decrease with increasing aneurysm morphology variables (D_a , A_a , V_a , and A_n), with statistically significant correlation between them (see [table 3](#)). An association between aneurysm size and shape and *WSS* was made by Ujii *et al* and Raghavan *et al*, and for the first time, morphological variables were considered in the assessment of intra-aneurysmal rupture risk.^{3 4} Our findings confirm that larger aneurysms have a lower *WSS* and larger *totime* than small aneurysms.

Two cases had a stenosis just before the aneurysm. These cases showed a fourfold reduction in cross-sectional area in the affected segment (~10 mm long), and a fourfold increase in the mean velocity. As a result, the velocity inside the aneurysm increased, inducing an elevated *WSS* (>10 Pa).

As indicated in previous studies, configuration of the aneurysm with respect to the parent vessel is important in intra-aneurysmal flow.^{5 36 37} In particular, when the aneurysm is located on the outer side of the bend, the inflow is higher. Our results confirm this finding for ICA aneurysms with similar shape and configuration to those in the mentioned studies. The smaller turnover time in aneurysms with higher α^n is produced

by the larger inflow when the aneurysm is located on the outer side of the vessel.

Our results suggest that the angle between aneurysm and parent vessel is also relevant. The angle α^n is similar to the inflow angle defined by Baharoglu *et al*.⁵ In that study, the inflow angle is associated with higher energy transmission to the dome and with a higher chance of rupture. Our analysis of α^t in relation to D_a showed that as aneurysms are larger they tilt upstream (larger α^t , or similarly, larger aneurysm). Larger aneurysms also presented larger *totime* and therefore, aneurysms tilting upstream showed a higher *totime*, an effect of the combination of size and orientation.

Reduction of hemodynamic variables after FD insertion in relation to morphology

The hemodynamic effect of FDs was characterized by analysis before and after treatment. Previous studies have shown that the presence of a FD reduces *WSS* inside an aneurysm, which is confirmed by our results.³⁸⁻⁴⁰ Additionally, we observed that aneurysms with relatively high *WSS* before treatment had a larger reduction in *WSS*.

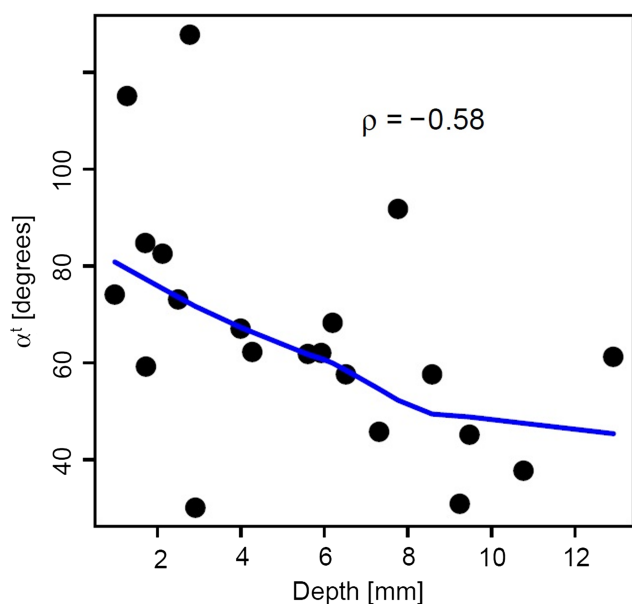


Figure 6 Scatterplot comparing depth (D_a) and α^t . The corresponding ρ for Spearman's test is shown. Also, a locally weighted polynomial regression line (blue) is shown where significant correlation was found.

FDs introduced into small vessels presented a low porosity (figure 7). This occurs because the number of wires of the FD (48 wires, 24 turning left and 24 turning right) and the width of the wires is constant and therefore, when the vessel has a smaller diameter, the porosity of the stent mesh is lower.

The d^{peak} had an impact on WSS reduction. We observed that aneurysms that were closer to the C^{peak} (lower d^{peak}) had less WSS reduction ($\rho = -0.49$). Closer to the peak, inertial effect due to vessel curvature is stronger, and secondary flow due to the Dean effect is more evident.⁸ Also as outlined by Thevenin *et al*, the effect of flow diverters is stronger on straight tubes than on curved ones, which is equivalent to the aneurysm being located far away or near the curvature peak, respectively.⁴¹

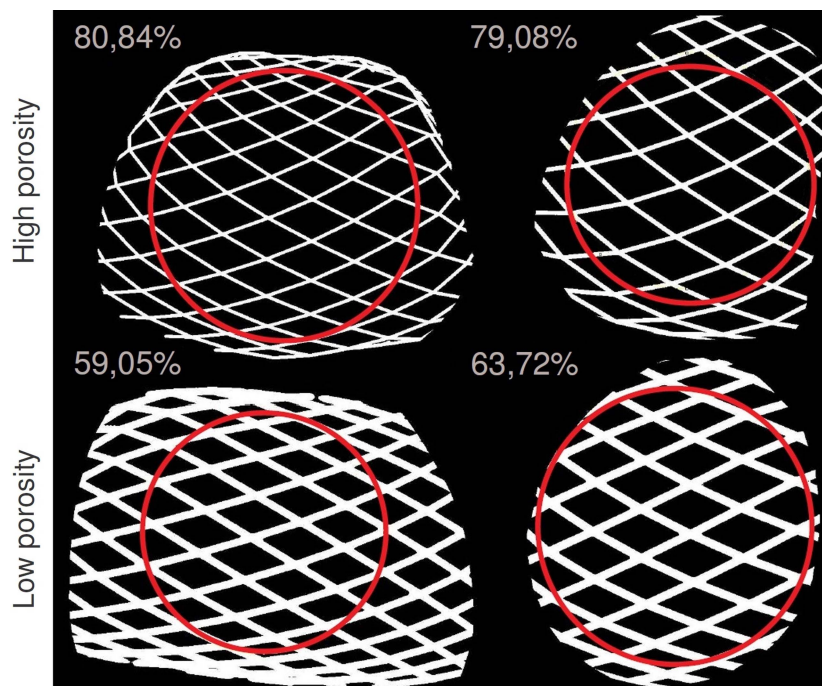
Aneurysms located on the outer side of the vessel bend had a smaller increase of *totime*. These cases are also prone to have shorter *totime* before the FD insertion ($\rho = -0.73$). Sato *et al* similarly found that the inflow into an aneurysm located on the outer side of a vessel is larger than the inflow into an aneurysm with equivalent shape and size on the inner side of the vessel.^{8,9} Additionally, after the insertion of the FD the absolute increase in *totime* is lower for aneurysms on the outer side of the bend. Owing to inertia, more flow of the parent vessel reaches the aneurysm when it is located on the outer side of the bend and

Table 4 Morphologic versus post-treatment hemodynamic variables reduction in ICA aneurysms correlations

	Aneurysm morphology					Vessel morphology			Porosity ϕ_{FD}
	D_a	A_a	V_a	A_n	AR_a	d^{peak}	α^n	α^t	
WSS	-0.51*	-0.54*	-0.55*	-0.49*	-0.37	-0.49*	0.35	0.42*	-0.53*
totime	-0.79*	-0.78*	-0.77*	-0.59*	-0.53*	-0.16	0.62*	0.65*	-0.08

*Indicates statistically significant correlation ($p < 0.05$).
ICA, internal carotid artery; WSS, wall shear stress.

Figure 7 Computation of stent porosity. Red circles delineate the measured region and the numbers indicate the flow diverter porosity.



therefore, the *totime* increase after FD insertion is smaller for such aneurysms.⁴¹

The FD does not alter the direction of the aneurysm inflow jet. It was observed that although the flow velocity is diminished inside the aneurysm and the inflow jet is dumped after the FD insertion, the main flow direction is not altered and the flow patterns inside the aneurysm are not modified substantially in most cases. Because the change induced by the FD mainly affects the intensity of aneurysm inflow, depending on the parent vessel and aneurysm geometry, and the patient flow conditions, the sole use of a single FD might not be enough to substantially decrease intra-aneurysmal flow below the critical threshold that would ensure aneurysm occlusion.

In this work, we studied the absolute rather than the relative differences in flow from baseline after FD placement. In a previous study, it was found that the relative change in hemodynamic variables was larger for smaller aneurysms.²² However, the hemodynamic conditions required for the occlusion of an aneurysm by hemostatic thrombosis most often cited in literature are absolute and not relative to the pretreatment condition.⁴² For these reasons absolute values of flow variables at baseline and post-treatment reduction were considered. It is important to notice that although the effect was larger for small than for large/giant aneurysms, some of the small aneurysms presented higher WSS and shorter *totime* after treatment than most of the large/giant aneurysms did before the treatment.

General considerations and applications

From the results and discussion presented in this paper, a series of considerations should be taken into account by the clinician. In summary, after treatment with a FD, a larger reduction in aneurysm flow and redirection to the vessel main stream should be expected for (a) aneurysms located further away from the C^{peak} , (b) aneurysms on the inner side of the bend, (c) aneurysms with no proximal stenosis, and (d) larger aneurysms. These aspects should be taken into consideration for their impact on the choice of treatment and on the post-treatment evolution and follow-up.

Limitations of the study

Different assumptions and hypotheses were considered for the development of this study. First, only a portion of the FD inserted into the parent vessel was modeled. Previous studies have shown that the parent vessel undergoes an increase in resistance due to the presence of a FD, so including only a portion of the FD might have led to small inaccuracies in the calculated pressure drop. Second, the vascular wall was modeled as rigid, implying an overestimation of pressure.¹⁴ Third, the resistance by peripheral vascular beds was not taken into account, and only pressures from a 1D arterial tree model were considered at the outlets. We assume that the inaccuracies in pressure conditions are small and will not have affected the conclusions of our study.

All cases in this study corresponded to the supraclinoid segment of the ICA to reduce differences in flow conditions and highlight the effect of geometry. The reader should be aware of this choice as the conclusions of this study may not hold for other locations.

Further aspects of patient-specific conditions of the coagulation system, including use of antiaggregants, were not considered in the simulations. This additional complexity will influence the event of aneurysm thrombosis in a given patient; however, it is unlikely that it will significantly affect flow produced by the FD.

CONCLUSIONS

In this study, the relation between parent vessel and aneurysm morphology, and intra-aneurysmal hemodynamics before and after FD treatment were investigated. The study included aneurysms located at the supraclinoid segment of the ICA with identical flow boundary conditions to identify differences due to morphology.

Aneurysm position and orientation with respect to its parent vessel were shown to be important, but aneurysm morphology showed higher correlation with WSS and *totime*. The presence of a stenosis before the aneurysm induced an increased intra-aneurysmal WSS and a decreased *totime*.

After FD insertion, intra-aneurysmal flow reduction was also found to depend on aneurysm position and orientation with respect to the parent vessel, but again, this effect was subordinate to aneurysm morphology. Finally, the vessel diameter had an impact on stent porosity, which is a determinant of flow reduction after treatment.

Contributors IL designed the study, analyzed the data, drafted the manuscript. AJG processed the data, analyzed the data, revised the manuscript. HGM MLA, DAR analyzed the data, revised the manuscript. All authors gave final approval of the manuscript.

Funding From the Networking Biomedical Research Center on Bioengineering (IL, AJG, HGM, MLA), Biomaterials and Nanomedicine (CIBER-BBN) and Universitat Pompeu Fabra, Barcelona, Spain; National Scientific and Technical Research Council (CONICET) - Argentina (IL) and Hirslanden Clinic (DAR), Zürich, Switzerland. This research has been partially funded by the Industrial and Technological Development Center (CDTI) under the CENIT-CDTEAM and CENIT-cvREMOD programs, the European Commission project @neurIST (IST-2005-027703).

Competing interests None.

Provenance and peer review Not commissioned; externally peer reviewed.

Data sharing statement An electronic copy summarizing the numerical data studied in this work is provided as supplementary material. These data remain owned by the authors and the associated rights reserved.

REFERENCES

- 1 Cebral JR, Lohner R. Efficient simulation of blood flow past complex endovascular devices using an adaptive embedding technique. *Med Imaging, IEEE Trans* 2005;24:468–76.
- 2 Xiang J, Natarajan SK, Tremmel M, *et al.* Hemodynamic-morphologic discriminants for intracranial aneurysm rupture. *Stroke* 2011;42:144–52.
- 3 Ujiie H, Tachibana H, Hiramatsu O, *et al.* Effects of size and shape (aspect ratio) on the hemodynamics of saccular aneurysms: a possible index for surgical treatment of intracranial aneurysms. *Neurosurgery* 1999;45:119–29; discussion 129–30.
- 4 Raghavan ML, Ma BS, Harbaugh RE. Quantified aneurysm shape and rupture risk. *J Neurosurg* 2005;102:355–62.
- 5 Baharoglu MI, Schirmer CM, Hoyt DA, *et al.* Aneurysm inflow-angle as a discriminant for rupture in sidewall cerebral aneurysms: morphometric and computational fluid dynamic analysis. *Stroke* 2010;41:1423–30.
- 6 Shojima M, Oshima M, Takagi K, *et al.* Magnitude and role of wall shear stress on cerebral aneurysm: computational fluid dynamic study of 20 middle cerebral artery aneurysms. *Stroke* 2004;35:2500–5.
- 7 Cebral JR, Mut F, Weir J, *et al.* Quantitative characterization of the hemodynamic environment in ruptured and unruptured brain aneurysms. *AJNR Am J Neuroradiol* 2011;32:145–51.
- 8 Imai Y, Sato K, Ishikawa T, *et al.* Inflow into saccular cerebral aneurysms at arterial bends. *Ann Biomed Eng* 2008;36:1489–95.
- 9 Sato K, Imai Y, Ishikawa T, *et al.* The importance of parent artery geometry in intra-aneurysmal hemodynamics. *Med Eng Phys* 2008;30:774–82.
- 10 Passerini T, Sangalli LM, Vantini S, *et al.* An integrated statistical investigation of internal carotid arteries of patients affected by cerebral aneurysms. *Cardiovasc Technol* 2011;3:26–40.
- 11 Nelson PK, Lylyk P, Szikora I, *et al.* The pipeline embolization device for the intracranial treatment of aneurysms trial. *AJNR Am J Neuroradiol* 2011;32:34–40.
- 12 Sadasivan C, Lieber BBB, Gounis MJM, *et al.* Angiographic quantification of contrast medium washout from cerebral aneurysms after stent placement. *AJNR Am J Neuroradiol* 2002;23:1214–0.
- 13 Klisch J, Turk A, Turner R, *et al.* Very late thrombosis of flow-diverting constructs after the treatment of large fusiform posterior circulation aneurysms. *AJNR Am J Neuroradiol* 2011;32:627–32.

- 14 Cebal JR, Mut F, Raschi M, *et al.* Aneurysm rupture following treatment with flow-diverting stents: computational hemodynamics analysis of treatment. *AJNR Am J Neuroradiol* 2011;32:27–33.
- 15 Kulcsar Z, Houdart E, Bonafé A, *et al.* Intra-aneurysmal thrombosis as a possible cause of delayed aneurysm rupture after flow-diversion treatment. *AJNR Am J Neuroradiol* 2010;23:20–5.
- 16 Sadasivan C, Cesar L, Seong J, *et al.* Treatment of rabbit elastase-induced aneurysm models by flow diverters: development of quantifiable indexes of device performance using digital subtraction angiography. *IEEE Trans Med Imaging* 2009;28:1117–25.
- 17 Larrabide I, Aguilar ML, Morales HG, *et al.* Intra-aneurysmal pressure and flow changes induced by flow diverters: relation to aneurysm size and shape. *AJNR Am J Neuroradiol* 2013;34: 816–22.
- 18 Pereira VM, Bonnefous O, Ouared R, *et al.* A DSA-based method using contrast-motion estimation for the assessment of the intra-aneurysmal flow changes induced by flow-diverter stents. *AJNR Am J Neuroradiol* 2013;34: 805–15.
- 19 Appanaboyina S, Mut F, Löhner R, *et al.* Simulation of intracranial aneurysm stenting: techniques and challenges. *Comput Methods Appl Mech Eng* 2009;198:3567–82.
- 20 Kulcsár Z, Augsburger L, Raymond P, *et al.* Flow diversion treatment: intra-aneurysmal blood flow velocity and WSS reduction are parameters to predict aneurysm thrombosis. *Acta Neurochir (Wien)* 2012;154:1827–34.
- 21 Villa-Uriol M, Berti G, Hose D, *et al.* @neurIST complex information processing toolchain for the integrated management of cerebral aneurysms. *Interface Focus* 2011;1:308–19.
- 22 Larrabide I, Aguilar ML, Morales HG, *et al.* Intra-aneurysmal pressure and flow changes induced by flow diverters: relation to aneurysm size and shape. *AJNR Am J Neuroradiol* 2013;34: 816–22.
- 23 Hernandez M, Frangi AF. Non-parametric geodesic active regions: method and evaluation for cerebral aneurysms segmentation in 3DRA and CTA. *Med Image Anal* 2007;11:224–41.
- 24 Larrabide I, Kim M, Augsburger L, *et al.* Fast virtual deployment of self-expandable stents: method and in vitro evaluation for intracranial aneurysmal stenting. *Med Image Anal* 2012;16:721–30.
- 25 Larrabide I, Villa-Uriol MC, Cárdenes R, *et al.* AngioLab-A software tool for morphological analysis and endovascular treatment planning of intracranial aneurysms. *Comput Methods Programs Biomed* 2012;108:806–19.
- 26 Larrabide I, Villa-Uriol MC, Cárdenes R, *et al.* Three-dimensional morphological analysis of intracranial aneurysms: a fully automated method for aneurysm sac isolation and quantification. *Med Phys* 2011;38:2439–0.
- 27 Piccinelli M, Veneziani A, Steinman DADA, *et al.* A framework for geometric analysis of vascular structures: application to cerebral aneurysms. *IEEE Trans Med Imaging* 2009;28:1141–55.
- 28 Piccinelli M, Bacigaluppi S, Boccardi E, *et al.* Geometry of the internal carotid artery and recurrent patterns in location, orientation, and rupture status of lateral aneurysms: an image-based computational study. *Neurosurgery* 2011;68:1270–85; discussion 1285.
- 29 Raymond P, Merenda F, Perren F, *et al.* Validation of a one-dimensional model of the systemic arterial tree. *Am J Physiol Heart Circ Physiol* 2009;297:H208–22.
- 30 Tremmel M, Xiang J, Hoi Y, *et al.* Mapping vascular response to in vivo hemodynamics: application to increased flow at the basilar terminus. *Biomech Model Mechanobiol* 2010;9:421–34.
- 31 Szymanski MP, Metaxa E, Meng H, *et al.* Endothelial cell layer subjected to impinging flow mimicking the apex of an arterial bifurcation. *Ann Biomed Eng* 2008;36:1681–9.
- 32 Park SI, Kim BM, Kim DI, *et al.* Clinical and angiographic follow-up of stent-only therapy for acute intracranial vertebrobasilar dissecting aneurysms. *AJNR Am J Neuroradiol* 2009;30:1351–6.
- 33 Kim M, Levy EI, Meng H, *et al.* Quantification of hemodynamic changes induced by virtual placement of multiple stents across a wide-necked basilar trunk aneurysm. *Neurosurgery* 2007;61:1305–12.
- 34 Team RDC. R: a language and environment for statistical computing. *Vienna Austria R Found Stat Comput* 2008;1:1–105.
- 35 Malek AM. Hemodynamic shear stress and its role in atherosclerosis. *JAMA J Am Med Assoc* 1999;282:2035–42.
- 36 Sato K, Imai Y, Ishikawa T, *et al.* The importance of parent artery geometry in intra-aneurysmal hemodynamics. *Med Eng Phys* 2007;30:774–82.
- 37 Imai Y, Sato K, Ishikawa T, *et al.* Inflow into saccular cerebral aneurysms at arterial bends. *Ann Biomed Eng* 2008;36:1489–95.
- 38 Augsburger L, Raymond P, Rufenacht DA, *et al.* Intracranial stents being modeled as a porous medium: flow simulation in stented cerebral aneurysms. *Ann Biomed Eng* 2010;39:850–63.
- 39 Liou TM, Li YC. Effects of stent porosity on hemodynamics in a sidewall aneurysm model. *J Biomech* 2008;41:1174–83.
- 40 Hassan T, Ahmed YM, Hassan AA. The adverse effects of flow-diverter stent-like devices on the flow pattern of saccular intracranial aneurysm models: computational fluid dynamics study. *Acta Neurochir (Wien)* 2011:1633–40.
- 41 Thevenin D, Seshadhri S, Janiga G. Impact of stents and flow diverters on hemodynamics in idealized aneurysm models. *J Biomech Eng* 2011;133:071005.
- 42 Wootton DM, Ku DN. Fluid mechanics of vascular systems, diseases, and thrombosis. *Annu Rev Biomed Eng* 1999;1:299–329.



Effect of aneurysm and ICA morphology on hemodynamics before and after flow diverter treatment

Ignacio Larrabide, Arjan J Geers, Hernán G Morales, et al.

J NeuroIntervent Surg published online April 1, 2014
doi: 10.1136/neurintsurg-2014-011171

Updated information and services can be found at:
<http://jnis.bmj.com/content/early/2014/04/01/neurintsurg-2014-011171.full.html>

These include:

Data Supplement

"Supplementary Data"
<http://jnis.bmj.com/content/suppl/2014/04/01/neurintsurg-2014-011171.DC1.html>

References

This article cites 39 articles, 13 of which can be accessed free at:
<http://jnis.bmj.com/content/early/2014/04/01/neurintsurg-2014-011171.full.html#ref-list-1>

P<P

Published online April 1, 2014 in advance of the print journal.

Email alerting service

Receive free email alerts when new articles cite this article. Sign up in the box at the top right corner of the online article.

Topic Collections

Articles on similar topics can be found in the following collections
[New devices](#) (79 articles)

Advance online articles have been peer reviewed, accepted for publication, edited and typeset, but have not yet appeared in the paper journal. Advance online articles are citable and establish publication priority; they are indexed by PubMed from initial publication. Citations to Advance online articles must include the digital object identifier (DOIs) and date of initial publication.

To request permissions go to:
<http://group.bmj.com/group/rights-licensing/permissions>

To order reprints go to:
<http://journals.bmj.com/cgi/reprintform>

To subscribe to BMJ go to:
<http://group.bmj.com/subscribe/>

Notes

Advance online articles have been peer reviewed, accepted for publication, edited and typeset, but have not yet appeared in the paper journal. Advance online articles are citable and establish publication priority; they are indexed by PubMed from initial publication. Citations to Advance online articles must include the digital object identifier (DOIs) and date of initial publication.

To request permissions go to:

<http://group.bmj.com/group/rights-licensing/permissions>

To order reprints go to:

<http://journals.bmj.com/cgi/reprintform>

To subscribe to BMJ go to:

<http://group.bmj.com/subscribe/>