

Responsabile scientifico:

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Main topic

▲ Aneurismi Cerebrali

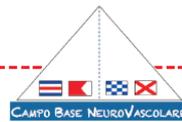


Intraoperative measurement of arterial blood flow in complex cerebral aneurysms' surgery

*Studio flussimetrico intra-operatorio nel clipping
degli aneurismi complessi*

Alessandro Della Puppa





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NO CONFLICT OF INTEREST TO DISCLOSE



Summary

Monitoring in aneurysm surgery

Flow-assisted microsurgical clipping

Flowmetry in complex aneurysms

Is it enough?

Neurophysiological monitoring and ICG

Multimodal strategy

New perspectives about flowmetry

Monitoring in aneurysm surgery

Monitoring in aneurysm surgery

Why?

Surgical Outcome

Authors	Period	n. patients	n. centers	Deficits post-op	Ischemic injury	Monitoring technique
Higashida et al. (USA)	2000	2535	429	11%	6%	No
Kim et al (Korea)	2006	1696	48	7%	7%	No
Amin-Hanjani et al (USA)	2000-2003	103	1	2%	2%	Flowmetry (All patients)
Kjung-Il Jo et al (Korea)	2008-2012	610	1	2%	4.5%	SSEP+MEPs (All patients)
Choi HH etal	2014-2017	386	1	1.8%	2.1%	SSEP+MEPs (All patients)

Causes of cerebral ischemia during aneurysm surgery

- 1 *Artery clip stenosis or occlusion*
- 2 *Temporary clipping*
- 3 *Retraction injury or ischemia*
- 4 *Intraoperative aneurysm rupture*
- 5 *Mechanical injury to artery*
- 6 *Decrease in mean systemic blood pressure*
- 7 *Intraoperative vasospasm*
- 8 *Embolic intraoperative events*

Aims of intra-operative monitoring in aneurysm surgery

To improve patient outcome by decreasing ischemic injury related to surgery

. Identification of potentially wrong current strategy

*. Surgical strategy can be changed within right time
(reversible phase)*

Causes of cerebral ischemia during aneurysm surgery

1 *Artery clip stenosis or occlusion*

2 *Temporary clipping*

3 *Retraction injury or ischemia*

4 *Intraoperative aneurysm rupture*

5 *Mechanical injury to artery*

6 *Decrease in mean systemic blood pressure*

7 *Intraoperative vasospasm*

8 *Embolic intraoperative events*



**Efficacy of
Intra-operative
Monitoring**

Monitoring in aneurysm surgery

Why?

How?

Monitoring Techniques for Prevention of Procedure-Related Ischemic Damage in Aneurysm Surgery

Susanna Bacigaluppi^{1,2}, Marco Fontanella^{3,4}, Pirjo Manninen⁵, Alessando Ducati³, Giovanni Tredici¹, Fred Gentili⁶

Prevention of Procedure-Related Ischemic Damage in Aneurysm Treatment

Monitoring Modality	Detection Characteristics										
	Depth	Area	Parent Vessel Stenosis	Perforator Occlusion	Sac Remnant	Feature Monitored	Frequency	Timing	Open/Closed Sky	Invasiveness	Cost
DSA	S and D	B	Yes	Yes	Yes	Flow	1–2 times	UEFU (intraop)	O and C	High	High
CT angiography (intraop)	S and D	B	Yes	No	Yes	Flow	1–2 times	UEFU (intraop)	O and C	Low	High
ICG videoangiography	S	B	Yes	Yes*	Yes	Flow	Repeatable	intraop UEFU	O	Low	Low
CT perfusion (intraoperative)	S and D	B	Yes, indir	Yes, indir	No	Flow	1–2 times	UEFU (intraop)	O and C	Low	High
Micro-Doppler	S	R	Yes, indir	Yes	Yes	Flow	Repeatable	intraop UEFU	O	Low	Low
Transient time flowmeter	S	R	Yes, indir	No	No	Flow	Repeatable	intraop UEFU	O	Low	Low
Laser speckle contrast analysis	S	B	Yes, indir	Yes, indir	No	Flow	Intermittent	Throughout	O	Low	Low
Thermal diffusion flowmetry	S	R	Yes, indir	Yes, indir	No	Flow	Continuous	Throughout	O	Low	Low
Microdialysis and brain tissue P _{O2} measurement	S	R	Yes, indir	Yes, indir	No	fx (metab)	Continuous	Throughout	O and C	High	Low
Near-infrared spectroscopy	S	R	Yes, indir	Yes, indir	No	fx (metab)	Continuous	Throughout	O and C	Low	Low
SSEP	S and D	B	Yes, indir	Yes, indir	No	fx (Nt)	Continuous	Throughout	O and C	Low	Low
BAEP	S and D	B	Yes, indir	Yes, indir	No	fx (Nt)	Continuous	Throughout	O and C	Low	Low
MEP	S and D	B	Yes, indir	Yes, indir	No	fx (Nt)	Intermittent	Throughout	O and C	Low	Low
EEG	S	B	Yes, indir	Yes, indir	No	fx (Nt)	Continuous	Throughout	O and C	Low	Low
ECOG	S (and D)	B	Yes, indir	Yes, indir	No	fx (Nt)	Continuous	Throughout	O	Low	Low

Surgical goals and intra-operative technologies

Flow monitoring techniques

- *Vessel flow visualization*
 - *ICG-videoangiography*
 - *Intra-operative DSA*
- *Ultrasonometry flow evaluation*
 - *Micro-Doppler*
 - *Transit Time Flowmetry*
- *Perfusion CT-scanning*

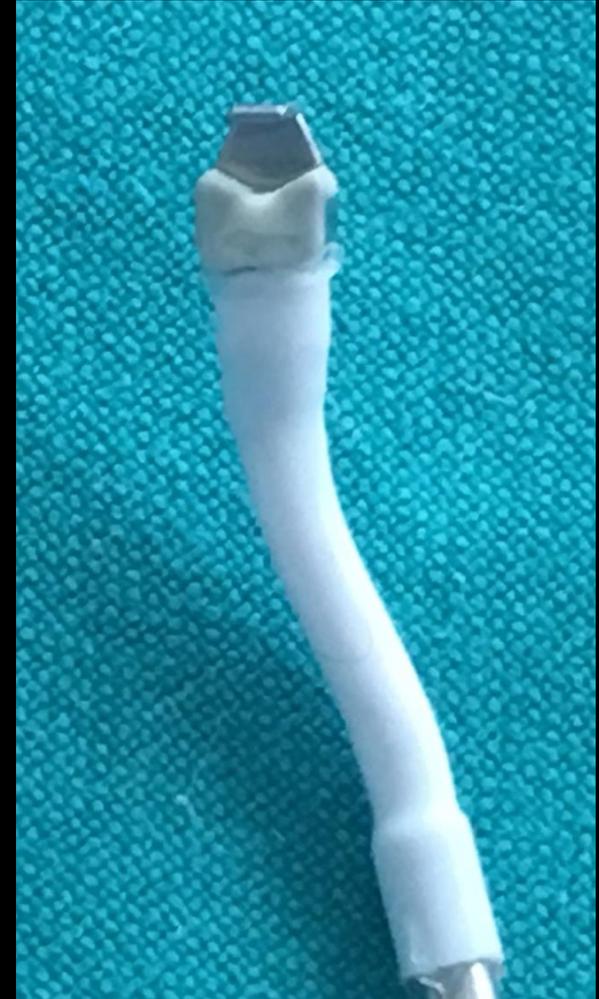
Function monitoring techniques

- *Neurophysiologic Monitoring*
(*Motor & Somato-Sensory EP*)
- *Detection of Metabolic Activity*
(*Microdialysis*)

Monitoring in aneurysm surgery
Flow-assisted microsurgical clipping

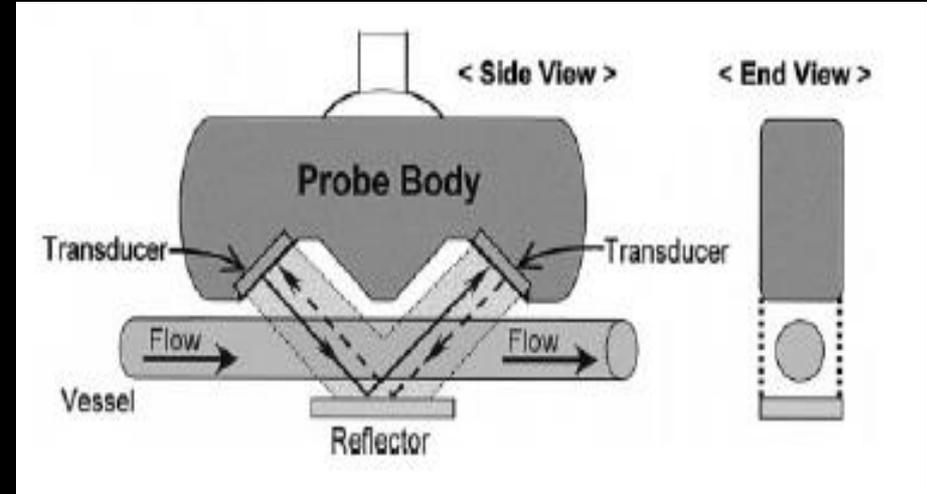
The flow probe device

Flow sensing perivascular microprobe

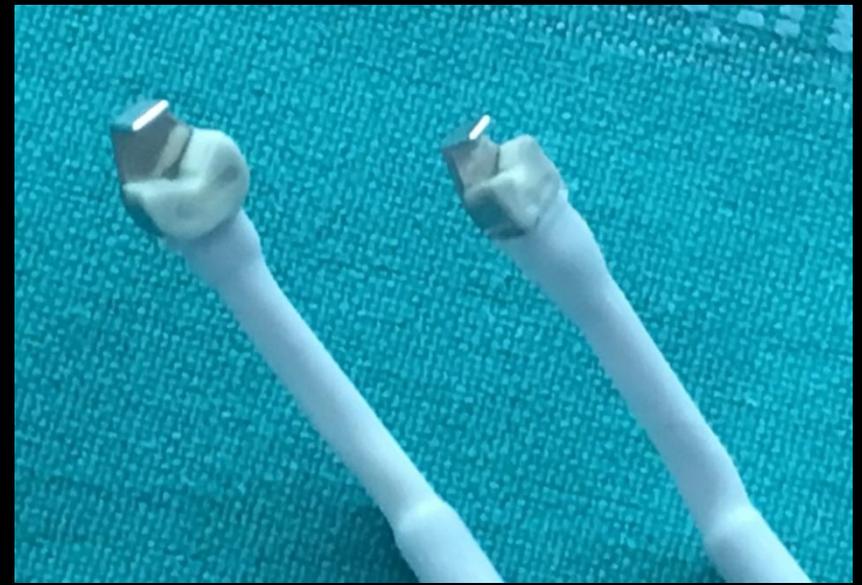
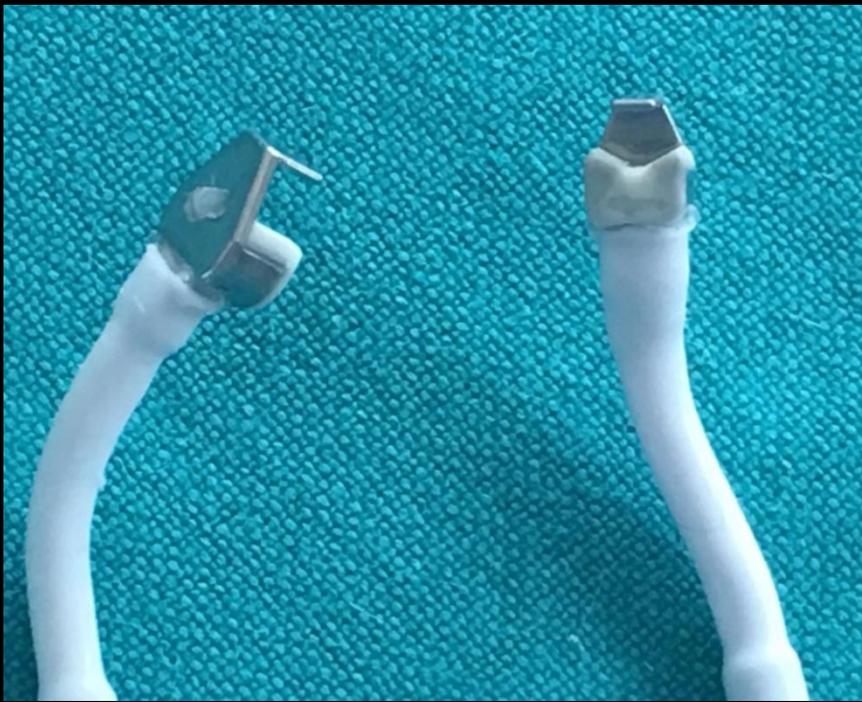


Electronic flow detection unit

The body of the probe consists of two ultrasonic transducers and a fixed acoustic reflector

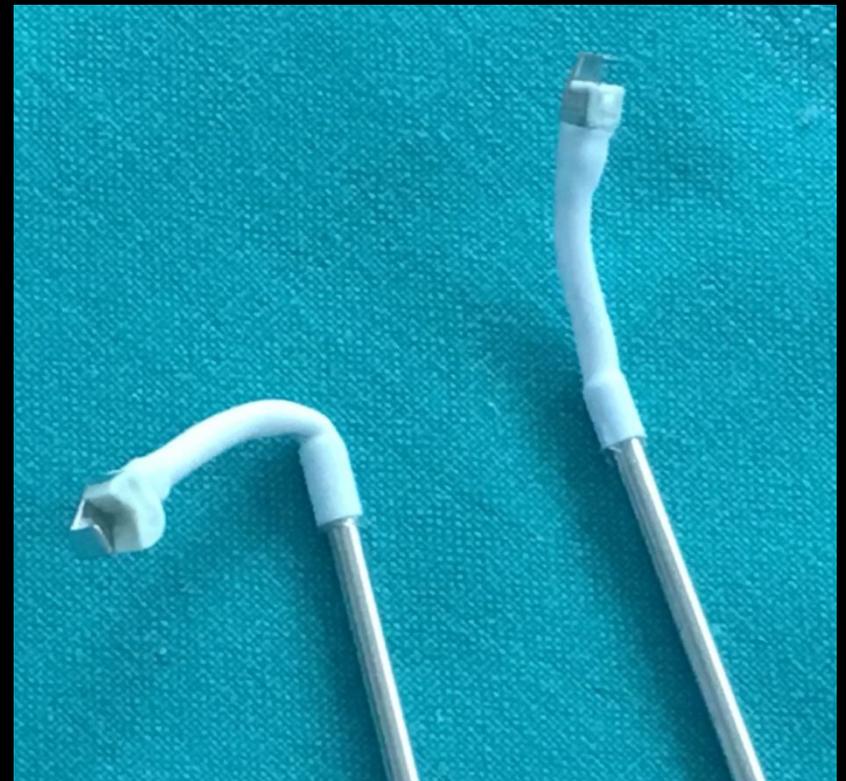


The two transducers emit and receive ultrasonic beams; once an ultrasonic beam starts from the first transducer, it reaches the reflector plate and subsequently the second transducer, and then the same happens from the second transducer to the first

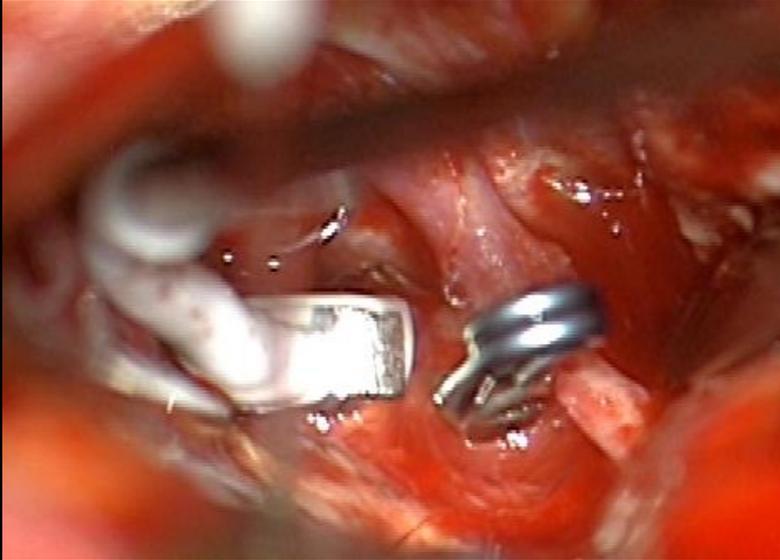


The probe is manufactured in a variety of sizes, ranging from 1.5 to 3 mm in diameter

The probe can be adapted to vessel features

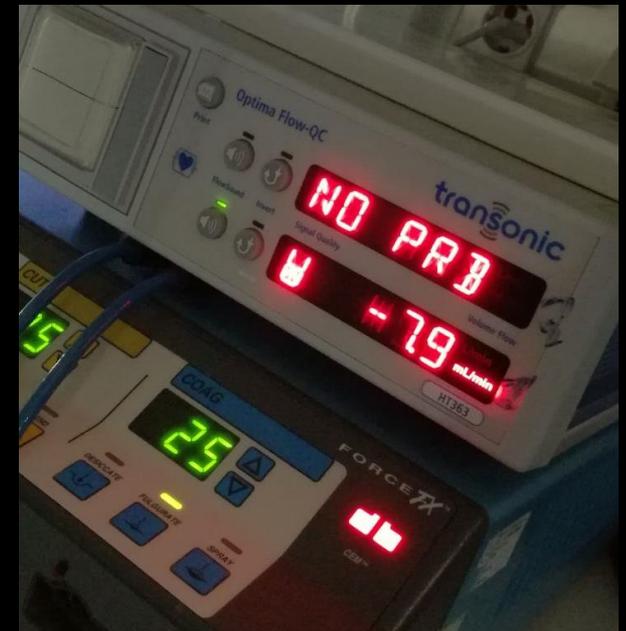


The flowmetry



Once the probe is placed around the vessel, the ultrasound beams have to intersect the vessel upstream or downstream to reach the transducers.

The ultrasound *transit time* is affected by the motion of flow through the vessel, and the difference between the upstream and downstream transit time is used to obtain the volume of flow (in mL/min) through the vessel insonated, which is calculated by the electronic flow detection unit



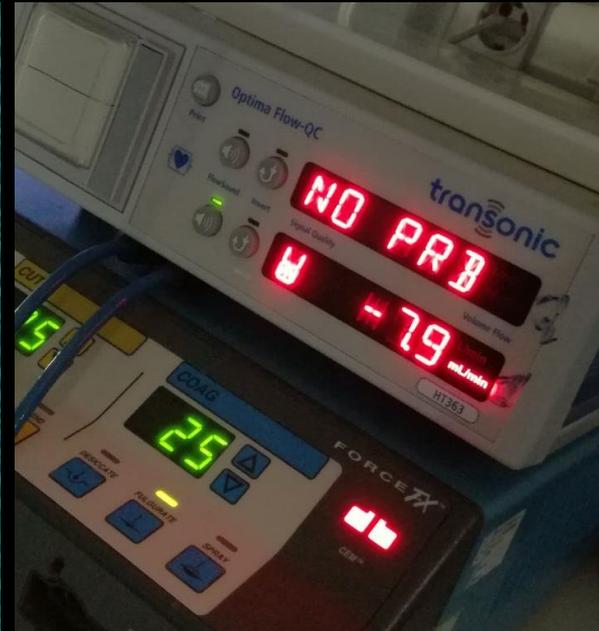
The flowmetry

Reading is influenced by:

- Systemic blood pressure
- End Tidal (PaCO₂)

Reading is not influenced by:

- Wall thickness
- Heart rate
- Hematocrit



Real time flow evaluation
Quantitative assessment (ml/min)
Can be easily repeated on demand

THE UTILITY OF INTRAOPERATIVE BLOOD FLOW MEASUREMENT DURING ANEURYSM SURGERY USING AN ULTRASONIC PERIVASCULAR FLOW PROBE

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OBJECTIVE: Inadvertent vessel compromise is one major cause of unfavorable outcome from aneurysm surgery. Existing strategies for intraoperative assessment of this complication have potential limitations and disadvantages. We assessed the utility of quantitative intraoperative flow measurements using the Transonic ultrasonic flow probe (Transonic Systems, Inc., Ithaca, NY) during aneurysm surgery.

METHODS: Of all aneurysms treated surgically at our institution from 1998 to 2003, 103 patients with 106 aneurysms were identified in whom intraoperative flow measurements were available for analysis. We assessed the frequency of flow compromise and clip repositioning and correlated these with postoperative angiography and stroke.

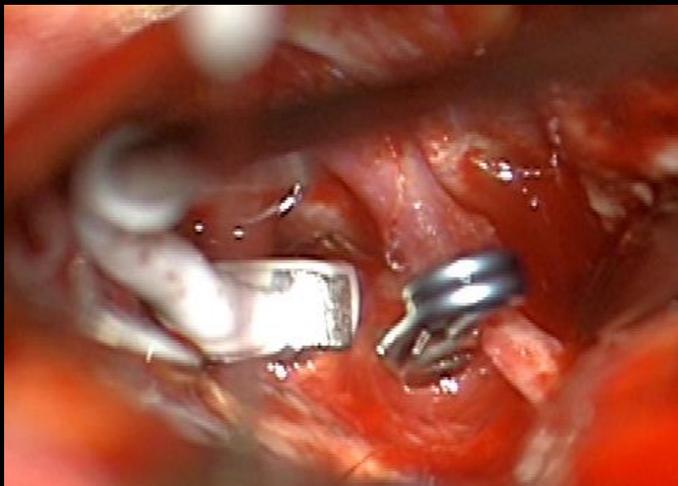
RESULTS: Significant (>25%) reduction in flow rate was apparent in 33 (31.1%) cases, and resulted in clip repositioning in 27 (25.5%), with return to baseline flow except for two cases with vessel thrombosis/dissection. In the other six cases, flow reduction was owing to spasm resolving with papaverine (n = 3) or responded to retractor repositioning (n = 3). In another six (5.7%) cases, unnecessary clip repositioning was avoided (n = 3) or safe occlusion of the parent vessel for trapping of the aneurysm was allowed by confirming adequate distal flow (n = 3). Aneurysms of the basilar, middle cerebral, anterior communicating, or carotid terminus were more likely to be associated with flow compromise (odds ratio, 4.3; *P* = 0.03). Postoperative angiography corroborated vessel patency in all cases, and no unexpected large vessel occlusions or strokes were evident.

CONCLUSION: Use of the ultrasonic flow probe provides real-time immediate feedback concerning vessel patency. Vessel compromise is easier to interpret than with Doppler, and faster/less invasive than intraoperative angiography. Intraoperative flow measurement is a valuable adjunct for enhancing the safety of aneurysm surgery.

KEY WORDS: Aneurysm, Angiography, Blood flow, Flow probe

Flow drop <25% → No Stroke!

Aneurysm exposure → Flowmetry



Clipping

Flowmetry



Flow drop >25%

Flow drop <25%

Clip repositioning

Flowmetry

Flow drop >25%

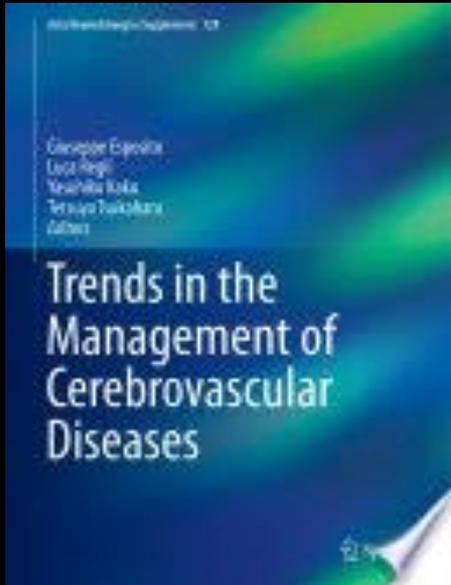
Flow drop <25%

Aneurysm
complete
exclusion

Video #1

Flowmetry Technique

(Right MCA unruptured aneurysm)



Intraoperative Measurement of Arterial Blood Flow in Aneurysm Surgery



Alberto Pasqualin, Pietro Meneghelli, Angelo Musumeci, Alessandro Della Puppa, Giacomo Pavesi, Giampietro Pinna, and Renato Scienza

	Ruptured	Unruptured	Total
<i>MCA</i>	92	161	253
<i>ACoA</i>	74	44	118
<i>ICA</i>	22	44	66
<i>Distal ACA</i>	13	7	20
<i>Posterior circulation</i>	2	5	7
<i>TOTAL</i>	203	261	464

Table 1 Intraoperative flowmetry for cerebral aneurysms; treated in Verona and Padua (2001–2010)

	Ruptured	Unruptured	Total
MCA	83	94	177
ACoA	66	27	93
ICA	19	35	54
Distal ACA	9	5	14
Posterior circulation	2	5	7
TOTAL	179	166	345

MCA Middle cerebral artery, *ACoA* Anterior communicating artery, *ICA* Internal carotid artery, *ACA* Anterior cerebral artery

Table 2 Intraoperative flowmetry for cerebral aneurysms; treated in Verona (2011–2016)

	Ruptured	Unruptured	Total
MCA	9	67	76
ACoA	8	17	25
ICA	3	9	12
Distal ACA	4	2	6
Posterior circulation	–	–	–
TOTAL	24	95	119

MCA Middle cerebral artery, *ACoA* Anterior communicating artery, *ICA* Internal carotid artery, *ACA* Anterior cerebral artery

Trends in the Management of Cerebrovascular Diseases

Intraoperative Measurement of Arterial Blood Flow in Aneurysm Surgery



Alberto Pasqualin, Pietro Meneghelli, Angelo Musumeci, Alessandro Della Puppa, Giacomo Pavesi, Giampietro Pinna, and Renato Scienza

Table 4 Basal flow values detected on cerebral arteries in patients with unruptured and ruptured aneurysms (Verona and Padua common experience, 2001–2010)

	Ruptured aneurysms	Unruptured aneurysms	<i>p</i> value
	Flow values ± SD	Flow values ± SD	
A1 tract	28.6 ± 10.9	30.9 ± 14.5	NS
A2 tract	24.7 ± 11.5	22.8 ± 10.3	NS
M1 tract	39.7 ± 14.5	39.2 ± 16.0	NS
M2 tract	21.7 ± 11.7	21.0 ± 10.6	NS
M3 tract	18.8 ± 11.0	13.5 ± 5.6	NS
Supraclinoid ICA	50.2 ± 23.0	42.0 ± 19.9	NS
Pericallosal A.	15.1 ± 4.6	14.9 ± 6.3	NS
PICA		12.2 ± 7.1	–

SD standard deviation, *NS* $p > 0.05$



Intraoperative Measurement of Arterial Blood Flow in Aneurysm Surgery



Alberto Pasqualin, Pietro Meneghelli, Angelo Musumeci, Alessandro Della Puppa, Giacomo Pavesi, Giampietro Pinna, and Renato Scienza

Clip repositioning was needed in 18% of patients
48% ruptured vs 52% unruptured

Clip Replacements were done

In the SAH group in 62% on MCA, 31% on ACoA, and 7% on ICA aneurysms

In the unruptured aneurysms in 82% in MCA, 12% in ACoA, 6% in pericallosal aneurysms

Complete flow restoration was obtained in 75% of patients

Monitoring in aneurysm surgery
Flow-assisted microsurgical clipping
Flowmetry in complex aneurysms

Last 5 years personal experience on flowmetry

Retrospective analysis 2013-2017

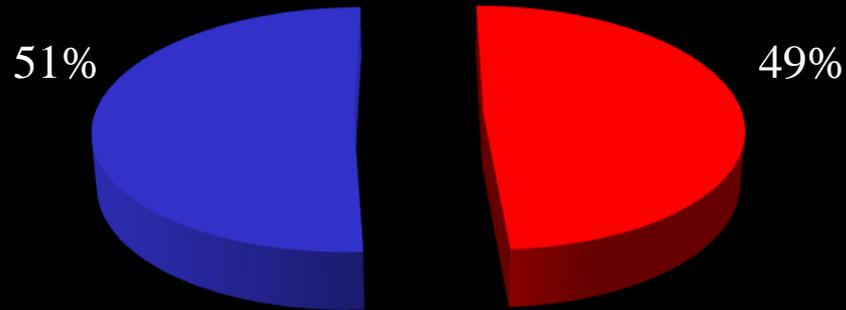
183 patients/207 aneurysms

1020 flow assessments by flowmetry

Last 5 years personal experience

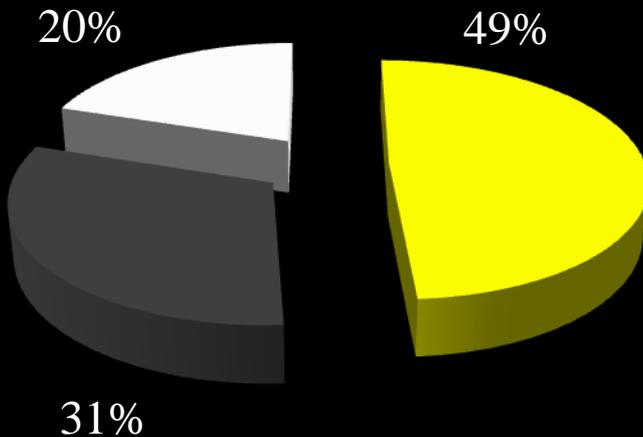
Aneurysms

■ Ruptured ■ Unruptured



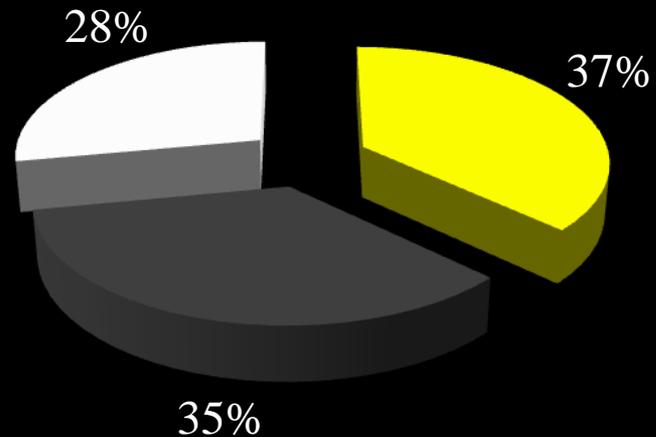
Aneurysm Location

■ MCA
■ ACA
■ ICA

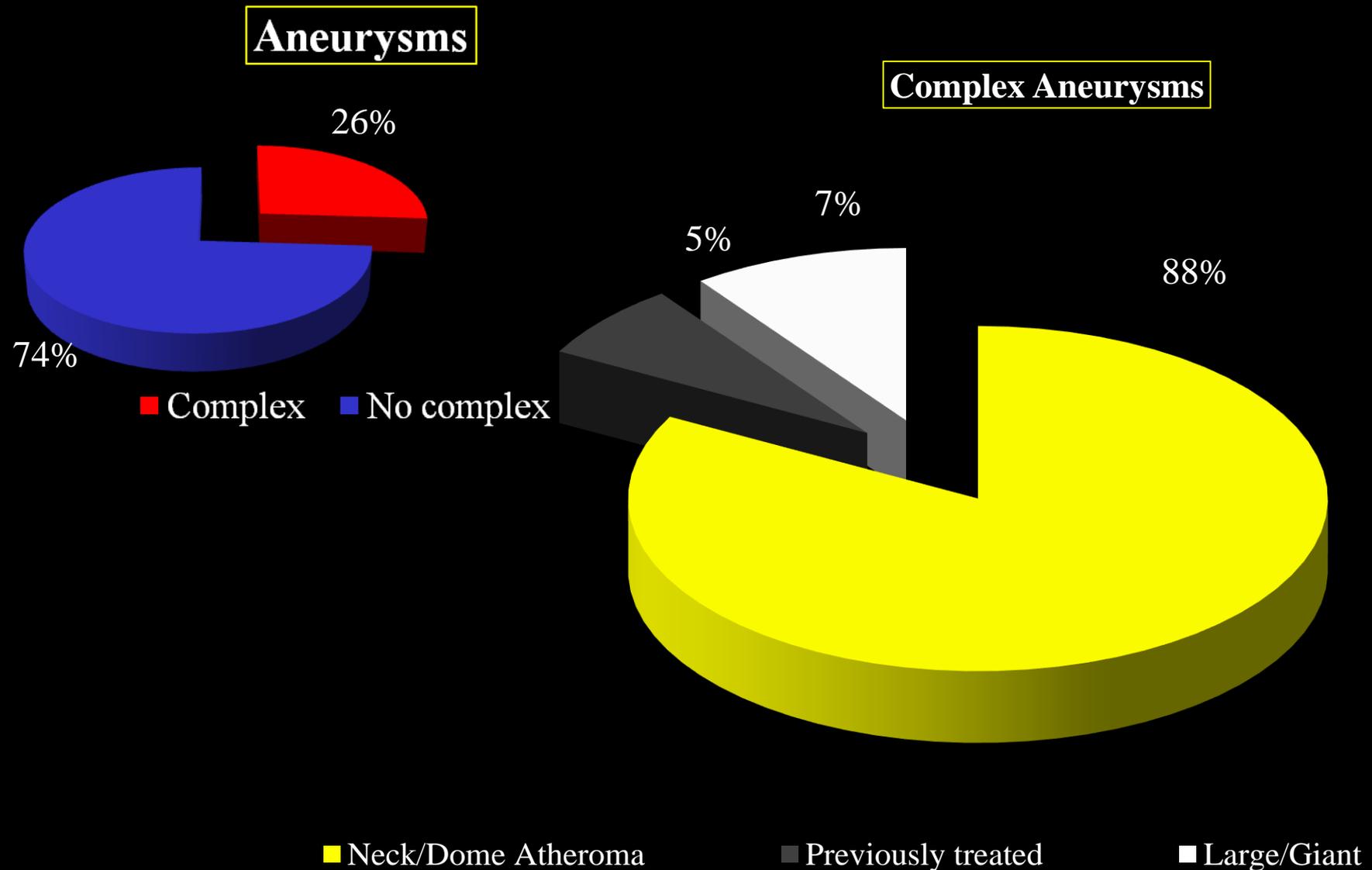


Aneurysm Size

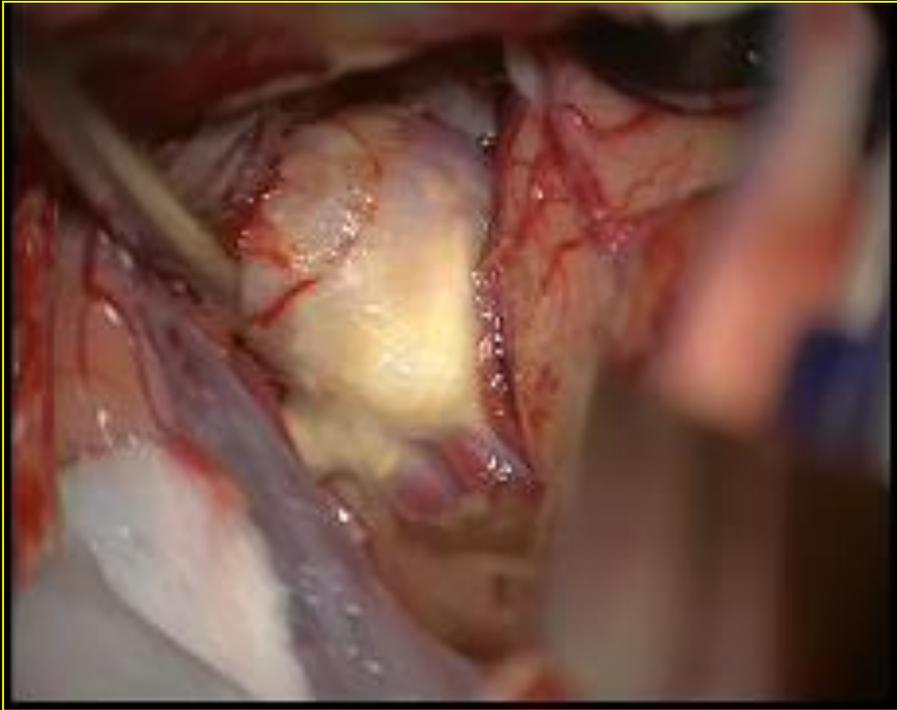
■ <7mm
■ 7-12mm
■ >12mm



Last 5 years personal experience – Complex Aneurysms



Neck/Dome Atheroma (Calcification)





Key points

High risk of vessel strain/distorsion due to neck/dome stiffness → vessel occlusion during clip positioning

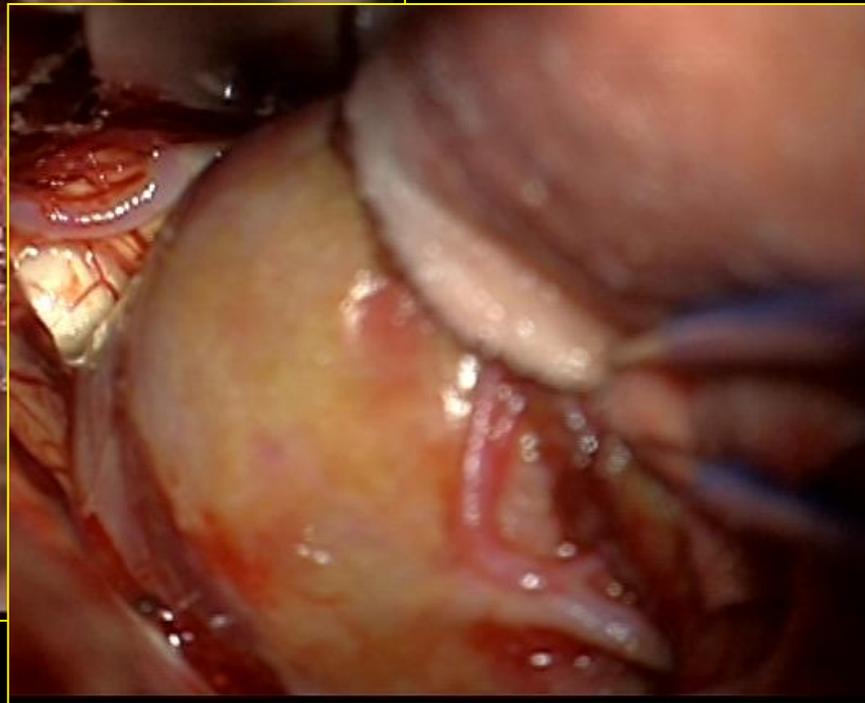
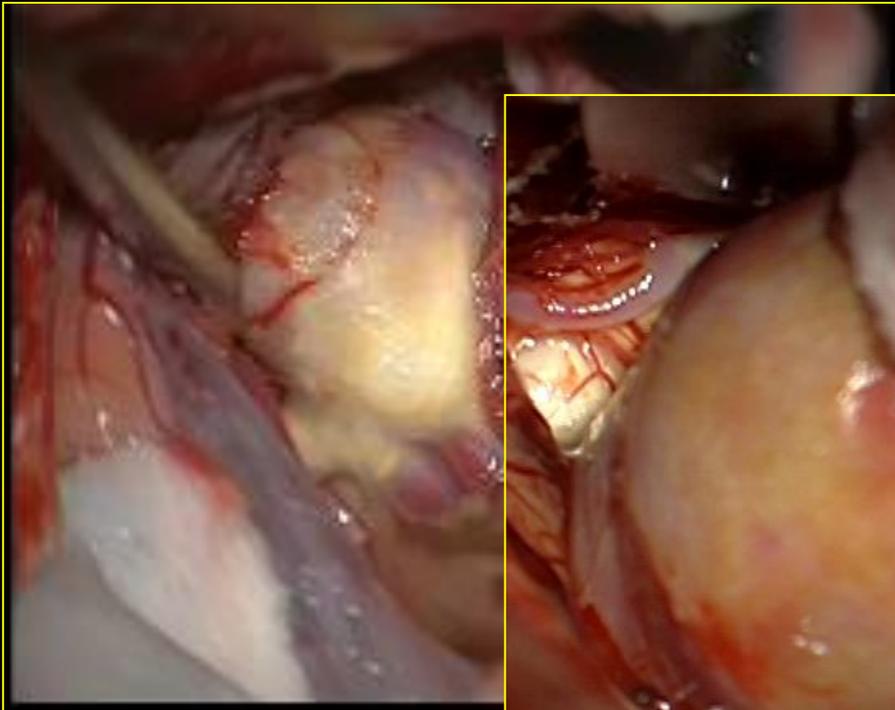
Video #2

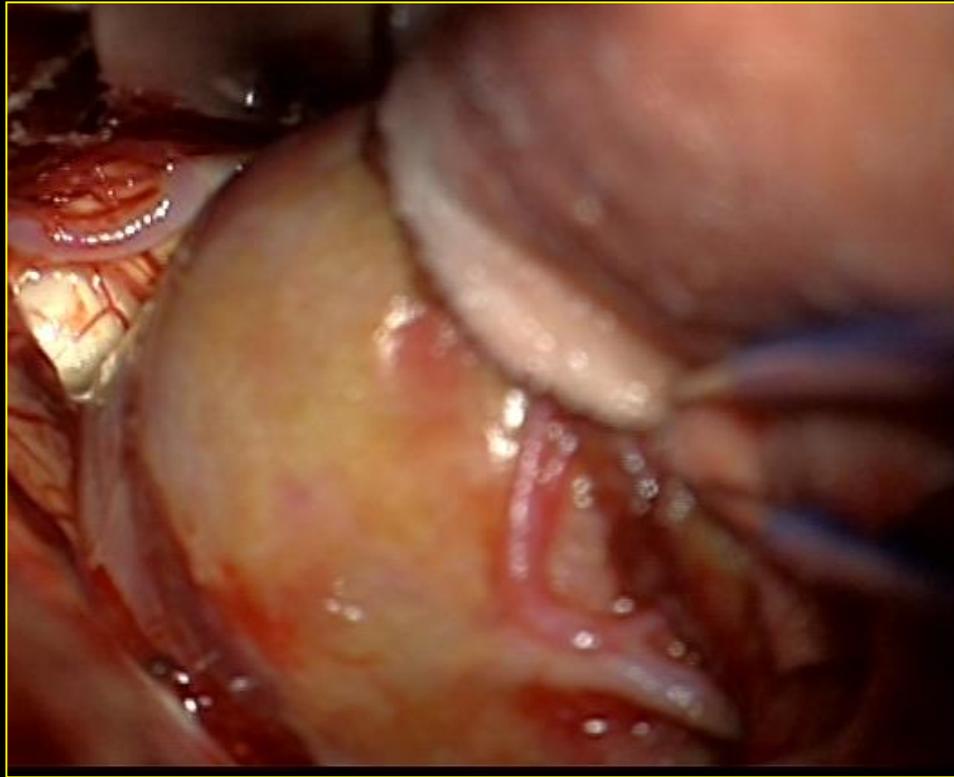
*Neck/Dome atheroma
(Left MCA unruptured aneurysm)*

Video #3

*Neck/Dome atheroma
(ACoA unruptured aneurysm)
(with single right ICA)*

Neck/Dome Atheroma (Calcification)
Large/Giant Aneurysms





Key points

1. High flow/Dome thrombus push clip down on the neck occluding branch arteries
2. Need of neck reconstruction after thrombus reduction/removal

Video #4

Giant aneurysm

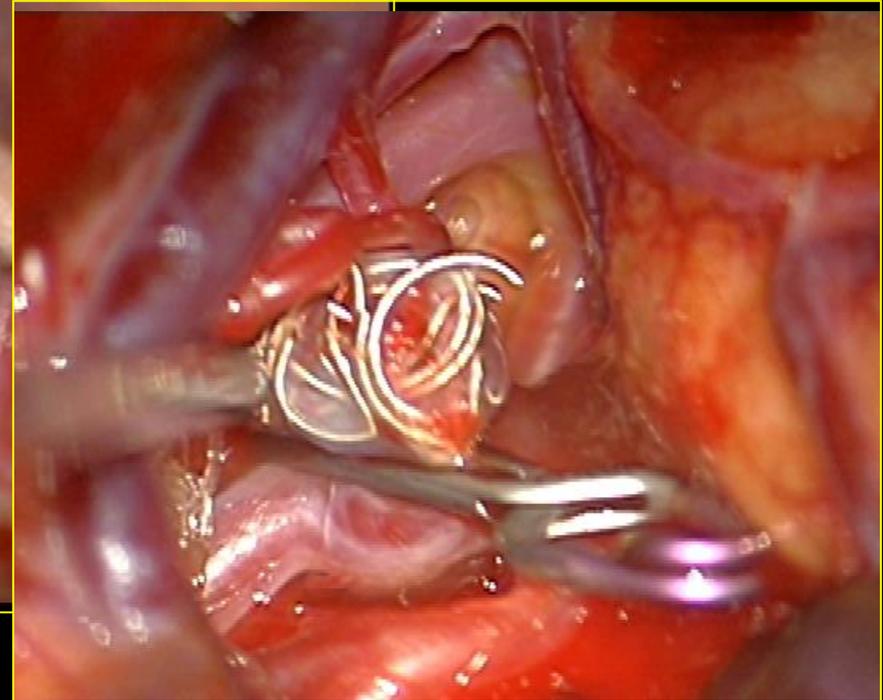
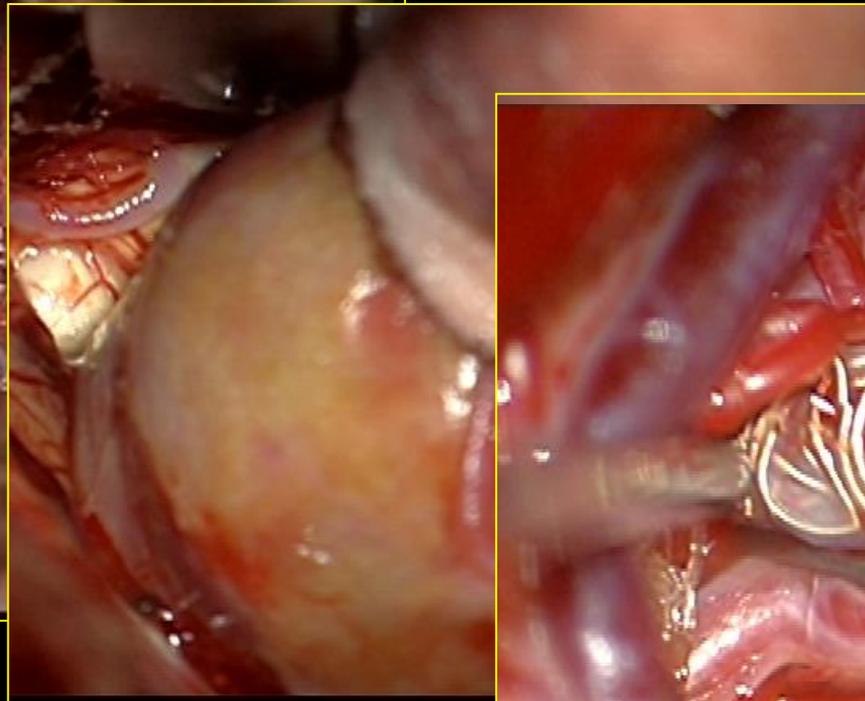
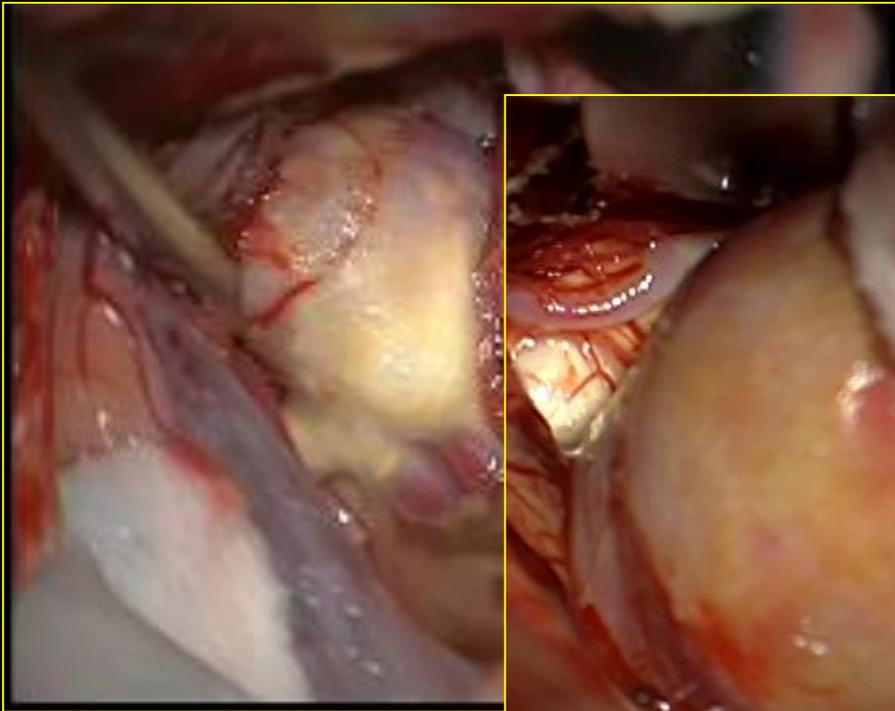
(Left MCA unruptured aneurysm)

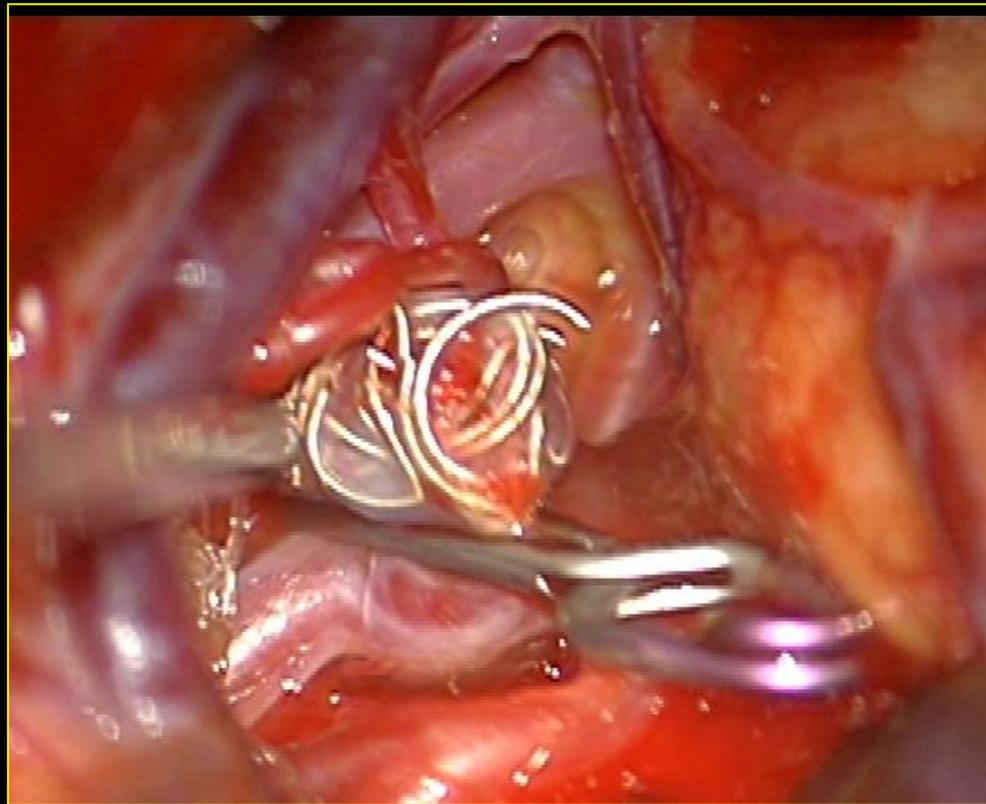
Video #5

Large aneurysm

(Left COFT unruptured aneurysm)

Neck/Dome Atheroma (Calcification)
Large/Giant Aneurysms
Previously treated Aneurysms



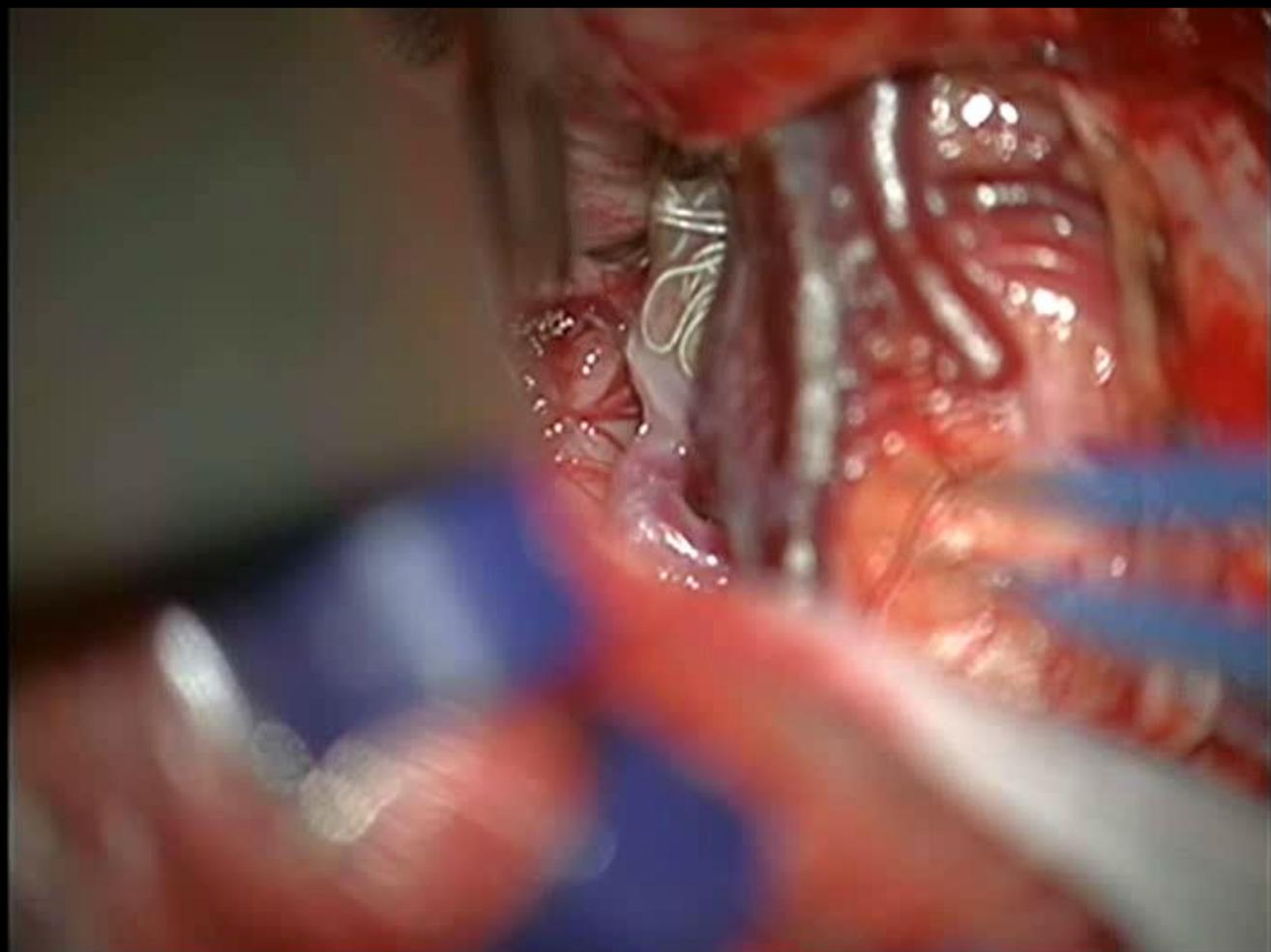


Key points

1. Small space on the neck for clip placement
2. Dome/Neck wall stiffness
3. Scar tissue around dome/neck and adhesion of coils to brain parenchyma through dome wall

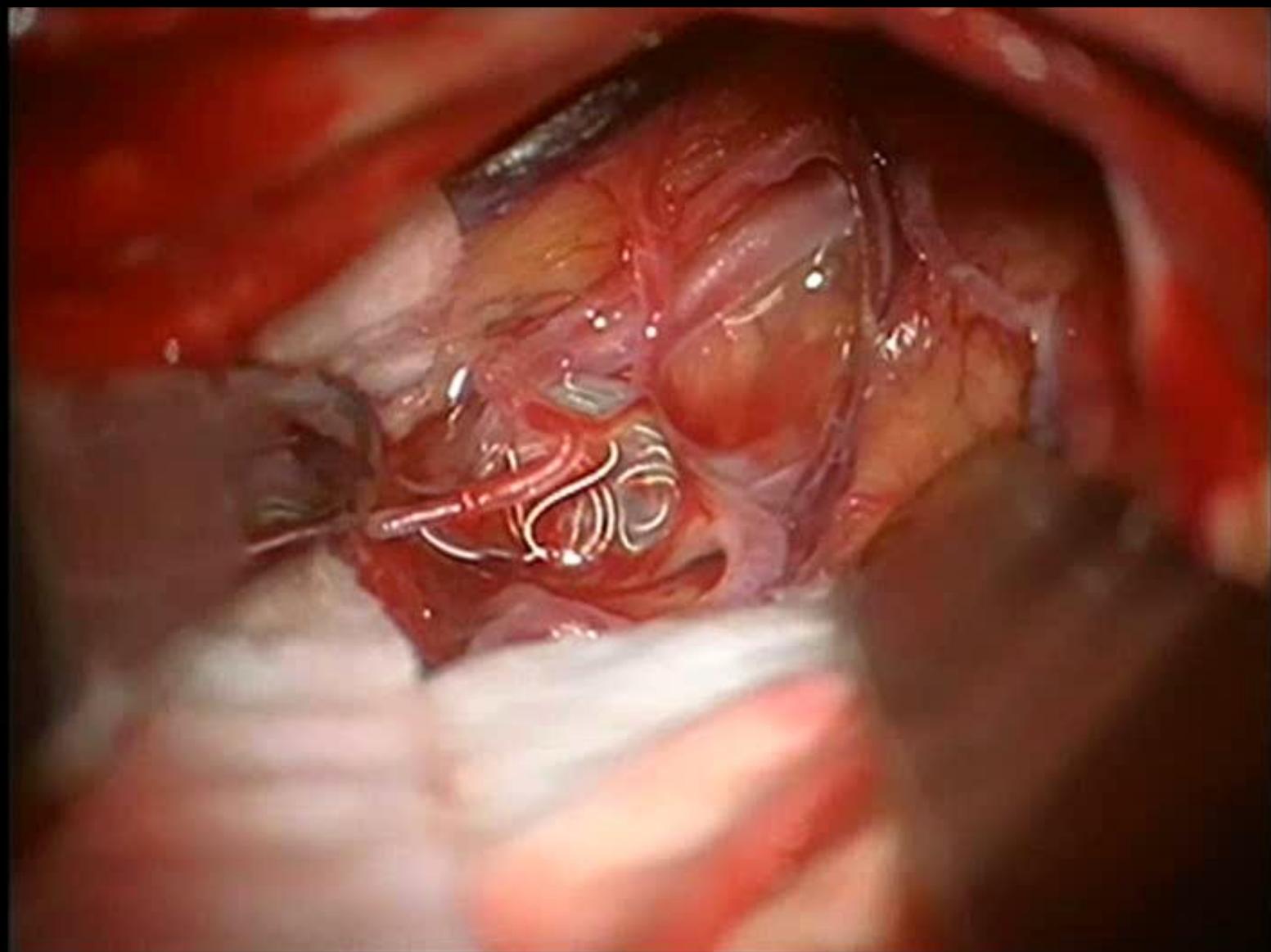
Video #6

*Previously coiled aneurysm
(Right MCA bleeding aneurysm)*



Video #7

*Previously coiled aneurysm
(Left MCA bleeding aneurysm)*



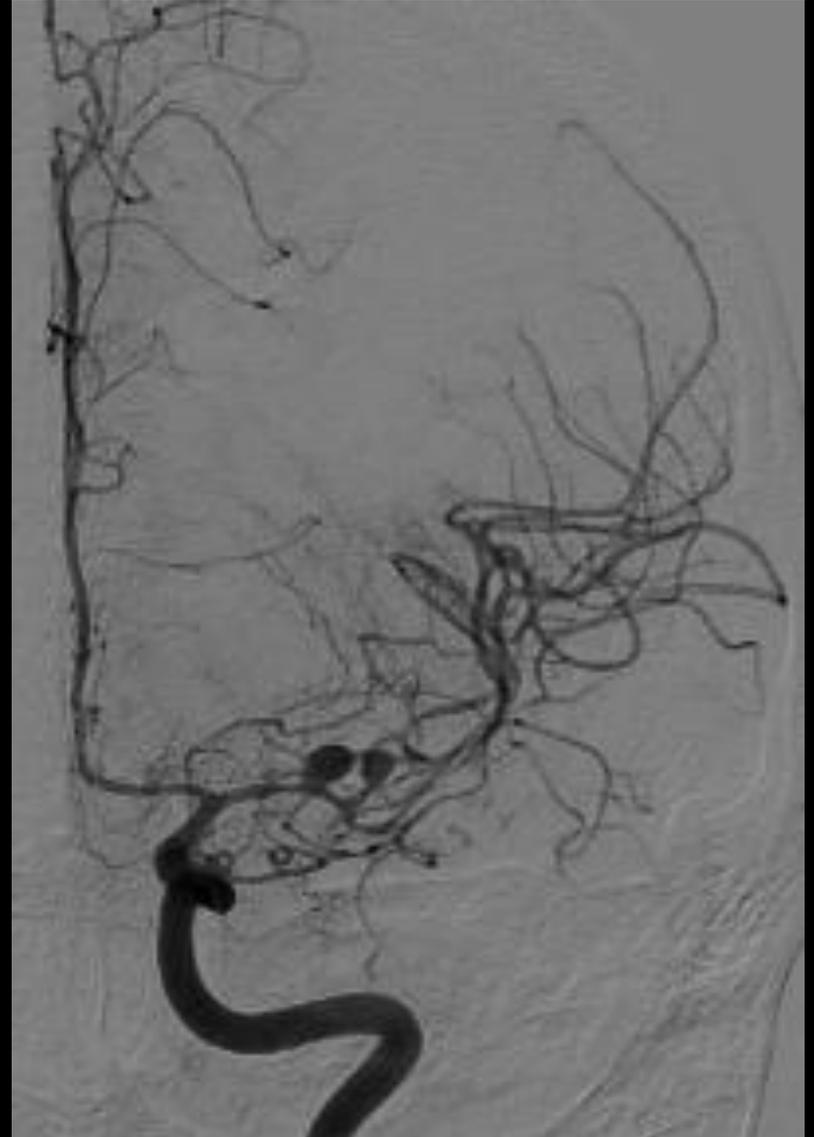
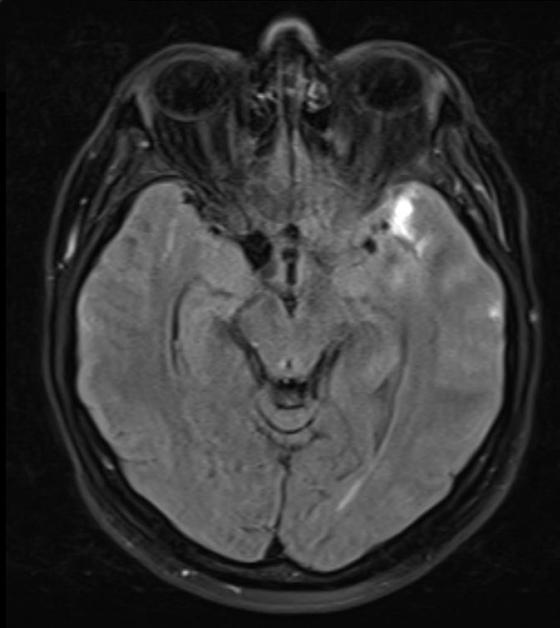
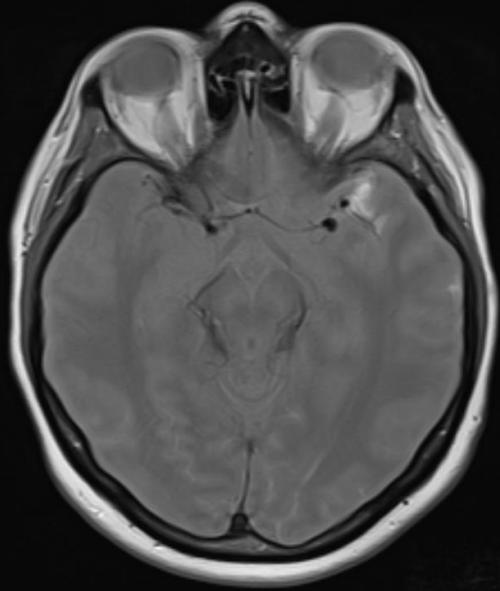
Monitoring in aneurysm surgery
Flow-assisted microsurgical clipping
Flowmetry in complex aneurysms
Is it enough?

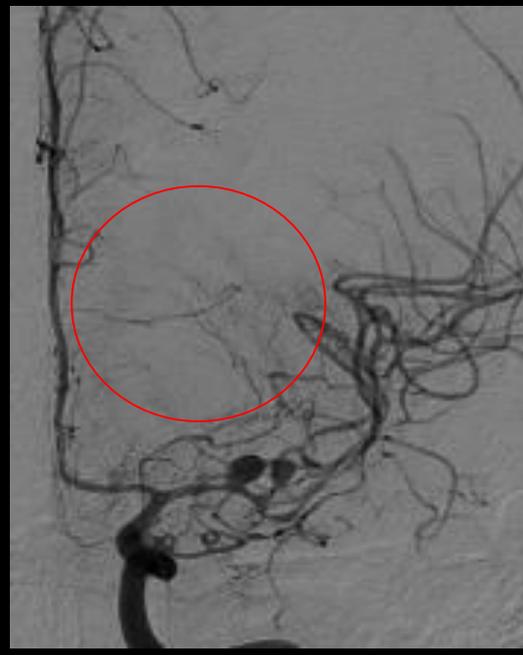
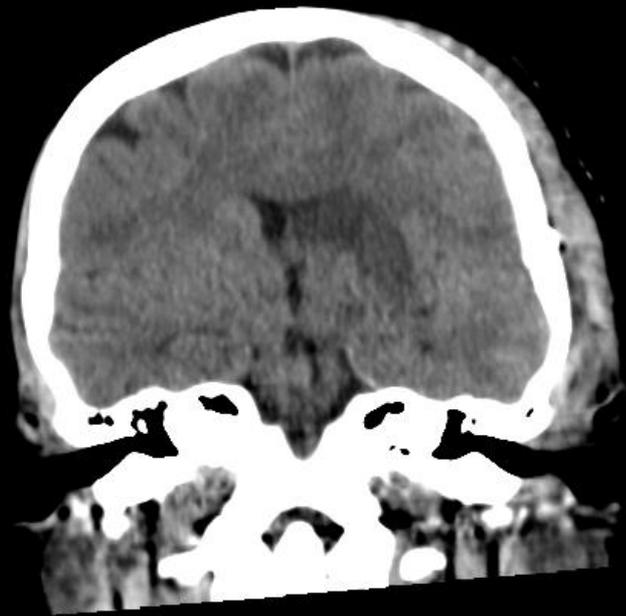
F/63Y SAH H&H:1

MCA bleeding bifurcation + M1 unruptured aneurysms

ICG-VA + flowmetry

No IOM





Strengths and weaknesses of intraoperative tools

	Advantages	Main drawbacks
Flowmetry	Quantitative assessment Safe, fast, easy Time No large exposure Can be repeated	Perforating arteries not assessed Learning curve
Intra-op DSA	Arteries non exposed in the surgical field	Invasive, Time consuming Team and equipment Time Repeated assessment is difficult
Microvascular Doppler	Real time assessment	Qualitative assessment
ICG-VA	perforator patency?	Can only assess vessels directly visible through the operating microscope field, Calcifications. Clot. Time Qualitative assessment
IOM	High predictivity	Influenced by anesthetic technique, prone to false negatives

Strengths and weaknesses of intraoperative tools

	Intraop DSA	Micro Doppler	Flowmetry	ICG	EPs
<u>Incomplete clipping</u>					
<i>Dome remnant</i>	++	(+)	-	++	-
<i>Neck remnant</i>	++	-	-	+	-
<u>Branching vessel</u>					
<i>Stenosis</i>	+	(+)	++	(+)	(+)
<i>Occlusion</i>	++	++	++	++	+
<i>Perforator occlusion</i>	(+)	(+)	-	+	++
<u>Time and procedure risk</u>	Yes	No	No	No	No

Raabe et al, J Neurosurg Sci 2016
(partially modified)

++ Very High + High (+) Low - Not applicable

Monitoring in aneurysm surgery

Flow-assisted microsurgical clipping

Flowmetry in complex aneurysms

Is it enough?

Neurophysiological monitoring and ICG

Our strategy – Neurophysiological monitoring

- . MEPs SSEPs monitoring*
- . Cortical vs Scalp electrodes*



Video #8

*MEPs role in perforators occlusion detection
(Right P-Comm unruptured aneurysm)*

ICG-VideoAngiography (ICG-VA)



Indocyanine green (ICG) dye is a near-infrared fluorescent tricarbo-cyanine dye

The absorption and emission peaks of ICG dye are 805 and 835 nm, respectively

After ICG is intravenously injected, it binds almost completely to globulins, preferentially to α 1-lipoproteins, within 1 to 2 seconds.

The normal vascular permeability is preserved, and the dye remains in the intravascular compartment.

The ICG dye is not metabolized in the body, and it is eliminated almost entirely via the liver.

The dye has a plasma half-life of 3 to 4 minutes

The recommended dose of ICG dye for videoangiography is 0.2 to 0.5 mg/kg, and the total daily dose should not exceed 5 mg/kg.

ICG-VideoAngiography (ICG-VA)

First published data in 2003[^]

- . 4-9% remnant detected[^]
- . 7% detection of main artery occlusion[°]
- . 3% detection of perforating artery occlusion[°]
- . 92-98% concordance with DSA[^]

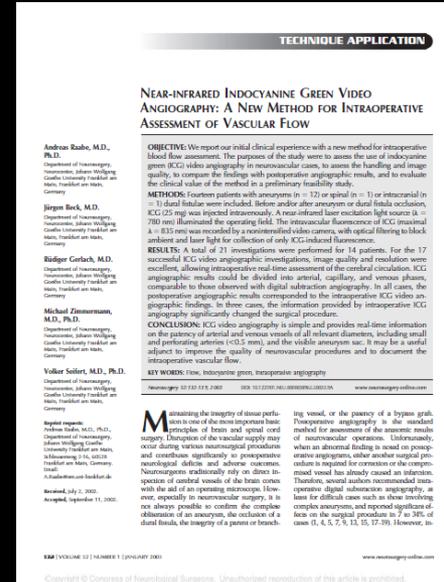
Prospective study in 2005*

Intra-op ICG vs Post-op DSA

- . Suboptimal or wrong clip position in 8 of total 10%*

Recent data

- . ICG lead to wrong clipping in 5-10% **
- . ICG cannot detect 3-7% additional findings showed by DSA**



[^] Raabe et al, Neurosurg 2003

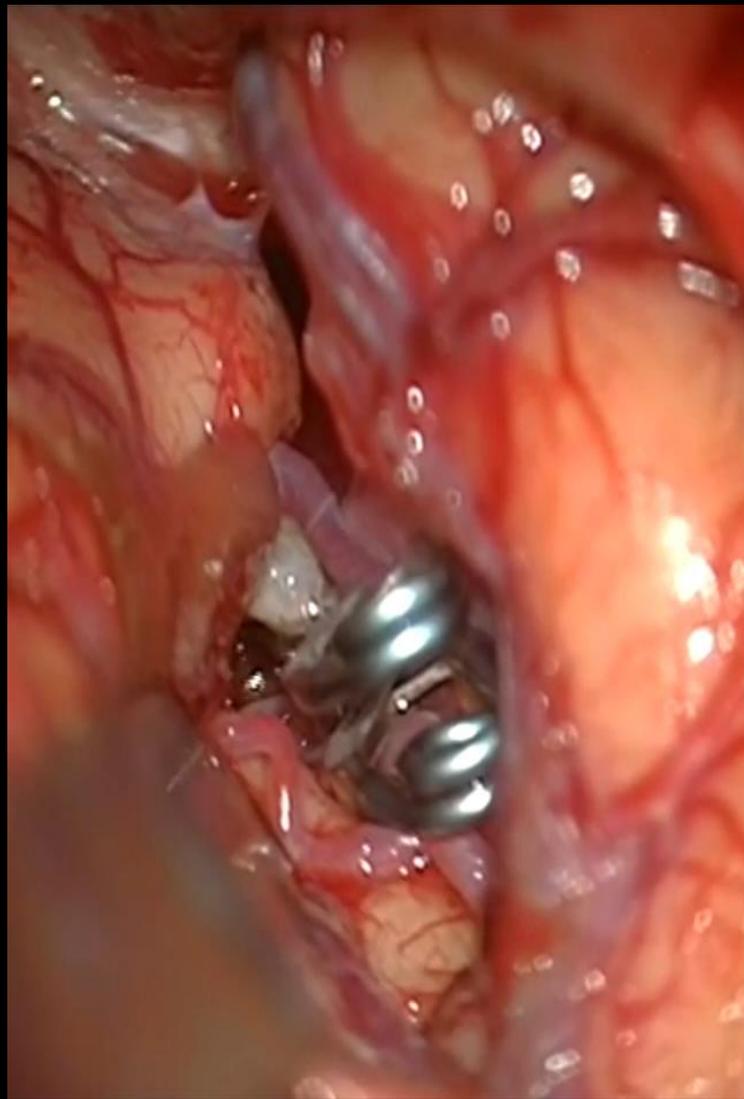
[°] De Oliveira et al, Neurosurg 2006

Long Replay 1 / 5 - 00:10.2 / 01:07



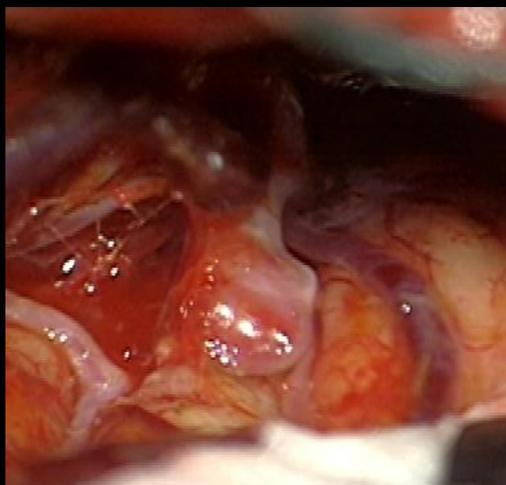
Video #9

*Indocyanine Green Videoangiography (ICG-VA)
(Right MCA unruptured aneurysm)*



Monitoring in aneurysm surgery
Flow-assisted microsurgical clipping
Is it enough?
Neurophysiological monitoring
Multimodal strategy

Aneurysm exposure



Flowmetry

ICG-VA (optional)

Clipping

Flowmetry



Flow drop >25%

Flow drop <25%

Clip repositioning

Flowmetry

Flow drop >25%

Flow drop <25%

ICG-VA

(squeezing)

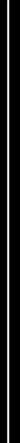
Complete exclusion

Remnant

Sac opening/puncturing



Neurophysiological Monitoring



Microsurgical Clipping of Intracranial Aneurysms Assisted by Neurophysiological Monitoring, Microvascular Flow Probe, and ICG-VA: Outcomes and Intraoperative Data on a Multimodal Strategy

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85 pts/96 aneurysms

Flowmetry + EPs + ICG

Aims

. Strategy results

Remnant, Strokes (clip related)

. Impact on surgery

Clip repositioning

*. Relationship CR/Aneurysm features
(Atheroma, Location, Size)*

Table 1. Patients and Aneurysms

Clinical Case Features	n (%)
Total number of aneurysms	96
Presentation	
Ruptured aneurysms	46 (47.9%)
Unruptured aneurysms	50 (52.1%)
Sex	
Male	28 (29.2%)
Female	57 (59.4%)
Neck-dome Atheroma	
Yes	26 (27.1%)
No	70 (72.9%)
Size	
<7 mm	11 (38.5%)
7–12 mm	18 (48.0%)
13–24 mm	13 (13.5%)
Aneurysm location	
Proximal group	45 (46.9%)
A1 segment	1 (1%)
Anterior communicating complex	16 (16.7%)
M1 segment	1 (1%)
ICA-posterior communicating	15 (15.6%)
ICA bifurcation	8 (8.3%)
ICA-anterior choroidal	2 (2.1%)
ICA-ophthalmic	2 (2.1%)
Distal group	51 (53.1%)
Pericallosal	2 (2.1%)
MCA bifurcation	49 (51%)

ICA, internal carotid artery; MCA, middle cerebral artery.

Strategy results

Aneurysm remnant	1.1%
Clip-related strokes	10.4%
Symptomatic	2.08%
Asymptomatic	8.32%

Table 2. Patients with Postoperative Stroke

Causes of Clip Repositioning

Patient Number	SAH	EPs	Flowmetry	ICG-VA	Ischemic Territory	Postoperative Deficits	Aneurysm Location	Atheroma
1	Yes	—	—	—	Hypotalamic	—	Bleeding ACoA + MCA bif	No
2	No	—	—	—	Fronto-orbitaly	—	ACoA + MCA bif	No
3	No	—	+	—	Frontal M2	Motor	MCA bif	Yes
4	Yes	—	+	—	Frontal M2	Language	MCA bif	Yes
5	Yes	—	+	—	Frontal M2	—	MCA bif	No
6	No	—	+	—	Basal ganglia	—	M1	No
7	Yes	+	—	—	Basal ganglia	—	P-comm	No
8	Yes	—	—	—	Hypotalamic	—	ACoA	No
9	Yes	—	—	—	Hypotalamic	—	ACoA	No
10	Yes	—	—	—	Hypotalamic	—	P-comm	No

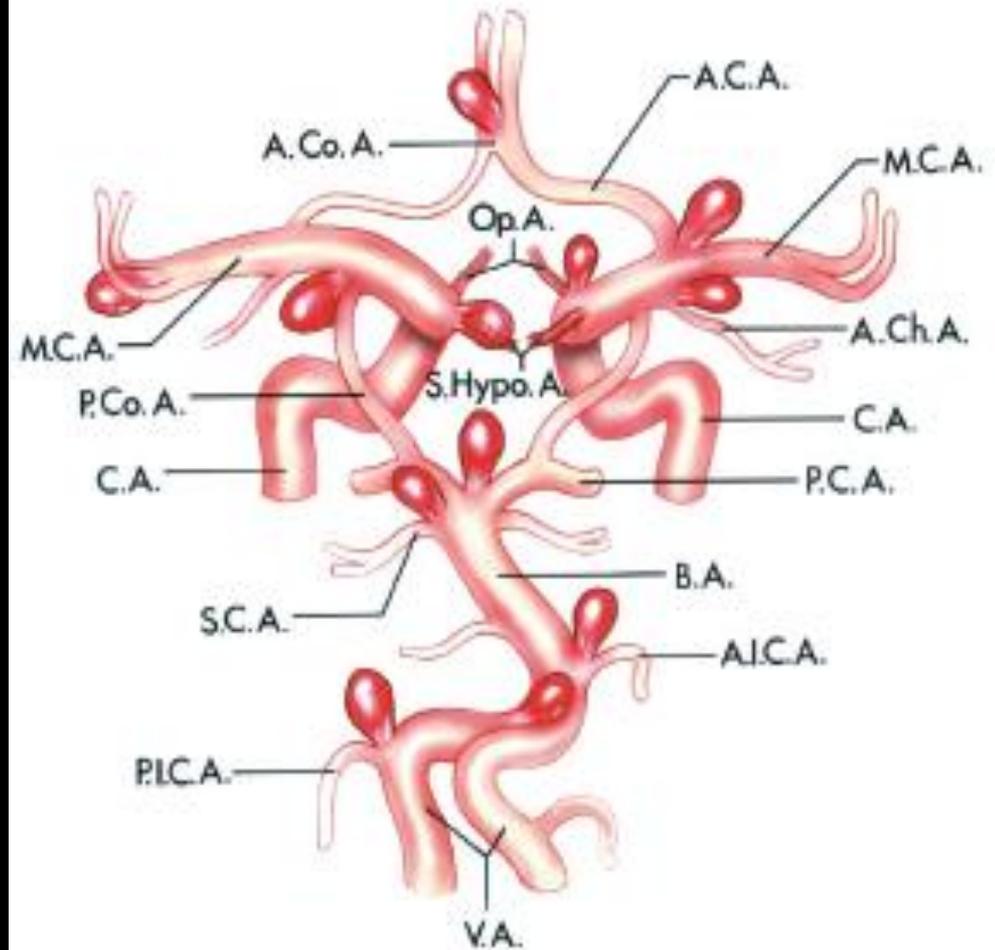
SAH, subarachnoid hemorrhage; EP, evoked potentials; ICG-VA, indocyanine green video angiography; ACoA, anterior communicating artery; MCA, middle cerebral artery; ICA bif, internal carotid artery bifurcation; P-comm, posterior communicating artery.

Clip repositioning rate 40.6%

MEP	9.3%
Flowmetry	22.91%
ICG-VA	8.3%
ICG	1.05%
Squeezing	7.25%

Monitoring technique depending on aneurysm feature

	Location		Atheroma
	Proximal	Distal	
MEP	p<0.05	-	-
Flowmetry	-	p< 0.01	p< 0.01
ICG-VA	-	-	p<0.05



A Tailored strategy?

- . Atheroma → Flowmetry, ICG*
- . Distal (MCA and pericallosal) → Flowmetry*
- . Perforators → EPs*

Monitoring in aneurysm surgery

Flow-assisted microsurgical clipping

Flowmetry in complex aneurysms

Is it enough?

Neurophysiological monitoring and ICG

Multimodal strategy

New perspectives about flowmetry

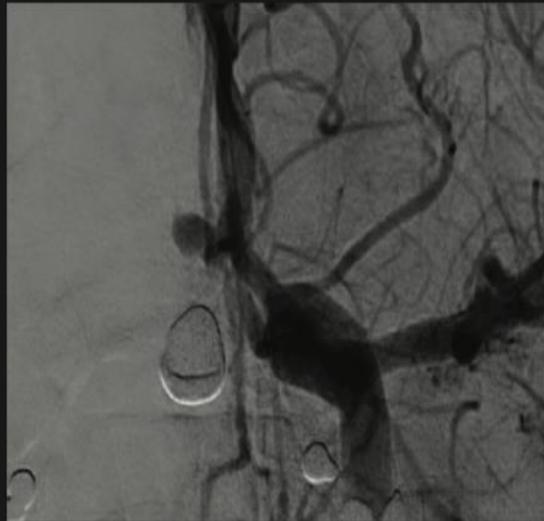
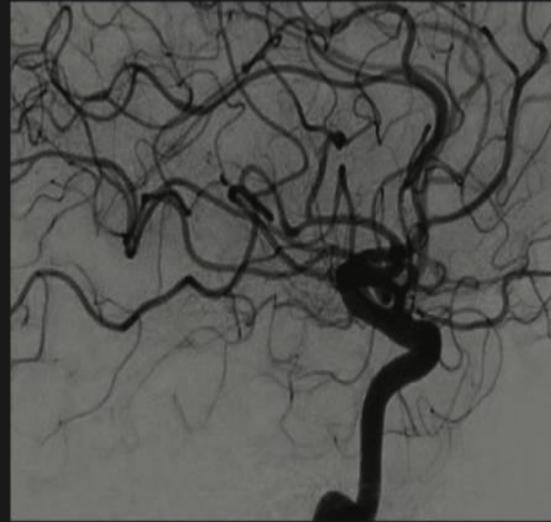


Video #10

*Small M2 flow evaluation with microprobe
(Right MCA unruptured aneurysm)*

Video #11

*Heubner Artery flow evaluation with microprobe
(ACoA unruptured aneurysm)*



Video #12

*P -Comm flow evaluation with microprobe
(P-Comm unruptured aneurysm)*

Final Considerations

- . Neurological outcome and quality of life represent primary aims of modern cerebrovascular surgery*
- . Intra-operative technologies can reduce post-operative stroke occurrence*
- . Flowmetry has a crucial role in reducing post-operative stroke due to main branches occlusion*
- . Best monitoring technique can depend on aneurysm features*
- . A “tailored monitoring strategy” can be pursued*