

Dr. Robin Novakovic-White Disclosures

No, nothing to discloseYes, please specify:

Company Name	Honoraria / Expenses		Funded Researc h	Royaltie s/ Patent		Ownersh ip/ Equity Position	Employee	Other (please specify)
AHA - S:VIN Journal								Commission Editor
Aidoc		X			X			
SVIN								President-Elect

Stroke Agenda

Objectives:

- Stroke reality and treatment options for ischemic stroke
- Once upon a thrombectomy..... a long, long, time ago!
 - ✓ Indication for thrombectomy in anterior circulation LVO
- Could there be a worse fate? Basilar artery occlusion
 - ✓ Indication for thrombectomy in BAO
- When is enough enough? Large core infarcts in anterior LVO
 - ✓ Indication for thrombectomy large core anterior LVO
- How far can we go? MeVO and DMVO
 - ✓ Indication for thrombectomy in MeVO??? Wait is there any?

US Incidence of Stroke

Volume 151, Issue 8, 25 February 2025; Pages e41-e660

2025 Heart Disease and Stroke Statistics: A Report of US

AHA STATISTICAL UPDATE

Circulation

and Global Data From the American Heart Association

In the US, stroke:

- Is a leading cause of disability
 - o 795,000 strokes occur annually
 - 07% US adults ≥60 yo living with brain injury from stroke

Every 40 seconds someone in the US has a stroke

- Fifth leading cause of death
 - o Deaths from stroke: 165,393 (2022) and 162,639 (2023)

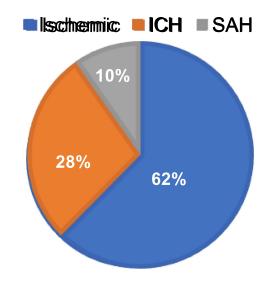


Every 3 minutes and 14 seconds someone in the US dies from a stroke

- SS Martin, et al. 2025 Heart Disease and Stroke Statistics: A Report of US and Global Data From the American Heart Association (Circulation 2025).
- Centers for Disease Control and Prevention. National Center for Health Statistics. Mortality data on CDC WONDER. https://wonder.cdc.gov/mcd.html (accessed
- W Andres, et al. Trends in the prevalence of stroke among community-dwelling individuals in the US 1999-2018 (JAMA Neurol, 2023).

Stroke Incidence

GEORAKETRPKE ITYPES



- SS Martin, et al. 2025 Heart Disease and Stroke Statistics: A Report of US and Global Data From the American Heart Association (*Circulation* 2025).
- GBD 2019 Stroke Collaborators. Global, regional, and national burden of stroke and its risk factors, 1990-2019: a systematic analysis for the Global Burden of Disease Study 2019 (*Lancet Neurol.* 2021).



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Stroke All Subtype Outcomes

- In a meta-analysis of 55 studies:
 - 56.7% (95% CI, 48.3%–65.1%) of people returned to work after stroke at 1 year
 - 66.7% (95% CI, 60.2%–73.2%) at 2 years
- Functional and cognitive impairment, and dementia are common after stroke.
 - P Duong, et al. Operational definitions and estimates of return to work poststroke: a systematic review and metaanalysis (Arch Phys Med Rehabil 2019).
 - S Koton, et al. Association of ischemic stroke incidence, severity, and recurrence with dementia in the Atherosclerosis Risk in Communities Cohort Study (JAMA Neurol 2022).





https://www.stroke.org/en/help-and-support/resource-library/prevention-

toolkit/preventing-stroke-presentation

National Impact of Stroke

- Direct healthcare cost \$36.7 billion (2020)
- Direct and indirect (unemployment, missed work-days, and premature mortality) \$56.8 billion (2020)
- Projected 2035 (in 2015 dollars) direct costs \$94.3 billion, and total cost of stroke:
 - \$81.1 billion for non-Hispanic White people
 - \$32.2 billion for non-Hispanic Black people
 - \$16.0 billion for Hispanic people



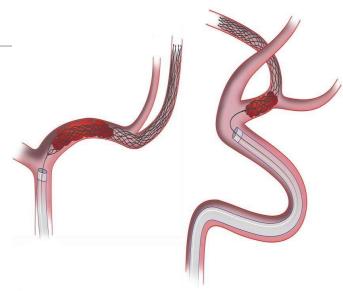
- SS Martin, et al. 2025 Heart Disease and Stroke Statistics: A Report of US and Global Data From the American Heart Association (*Circulation* 2025).
- Agency for Healthcare Research and Quality. Medical Expenditure Panel Survey (MEPS): household component summary tables: medical conditions, United States. Accessed April 2, 2023. https://meps.ahrq.gov/ mepsweb/
- RTI International. Projections of cardiovascular disease prevalence and costs: 2015–2035: technical report [report prepared for the American Heart Association]. RTI International; 2016. RTI project No. 021480.003.001.001.





Treatment Options

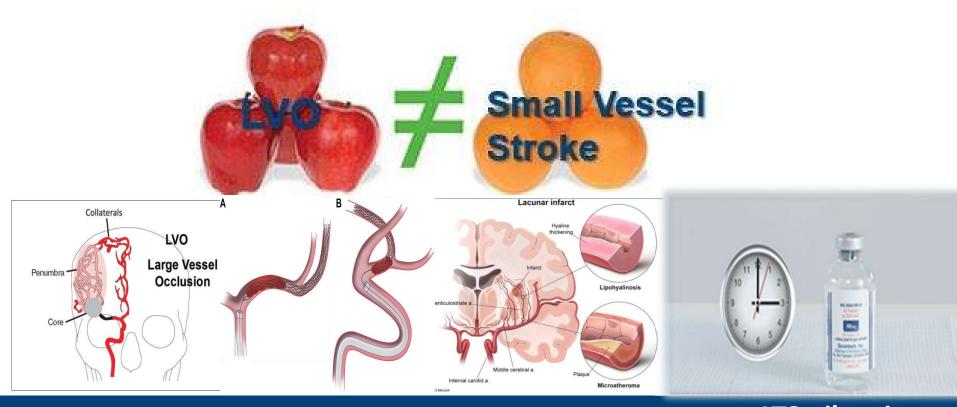
- Most effective treatment for AIS is early reperfusion therapy (opening the artery).
- Earlier treatment is associated with better recovery and outcomes.
- Factors that can delay treatment and contribute to poor outcome:
 - Limited resources
 - Pre-hospital failure to recognize stroke
 - Geographic barriers
 - Lack of expertise
- Saver, J L, et al. Dis. J. Am. Med. Assoc. Netw. 2016:13647.
- Kim, JT, et al. Circulation, 2017:135, 128–139.
- Agarwal, S, et al. Neurology. 2018: 90, e316–e322.
- Brain Attack Coalition Symposium: Inequities in Access and Delivery of Acute Stroke Care. Available online
 at: https://www.ninds.nih.gov/news-events/events/brain-attack-coalition-symposium-inequities-access-and-delivery-acute-stroke-care (accessed April 24, 2025).
- Chari, SV, et al. Am. J. Emerg. Med. 2023:63, 120–126.
- Oostema, JA, et al. J. Am. Heart Assoc. 2023:12:e026834.

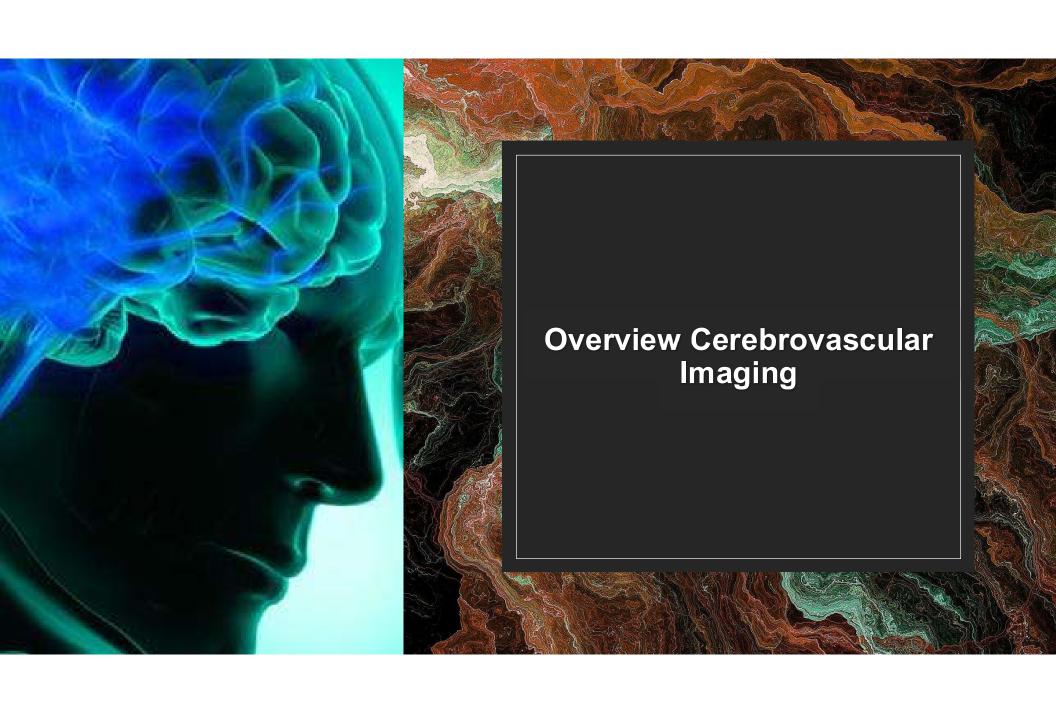




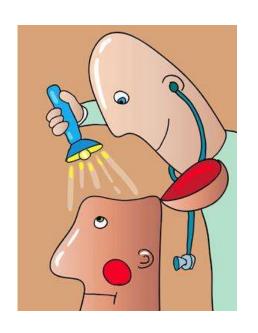
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Are All Strokes Created Equal?

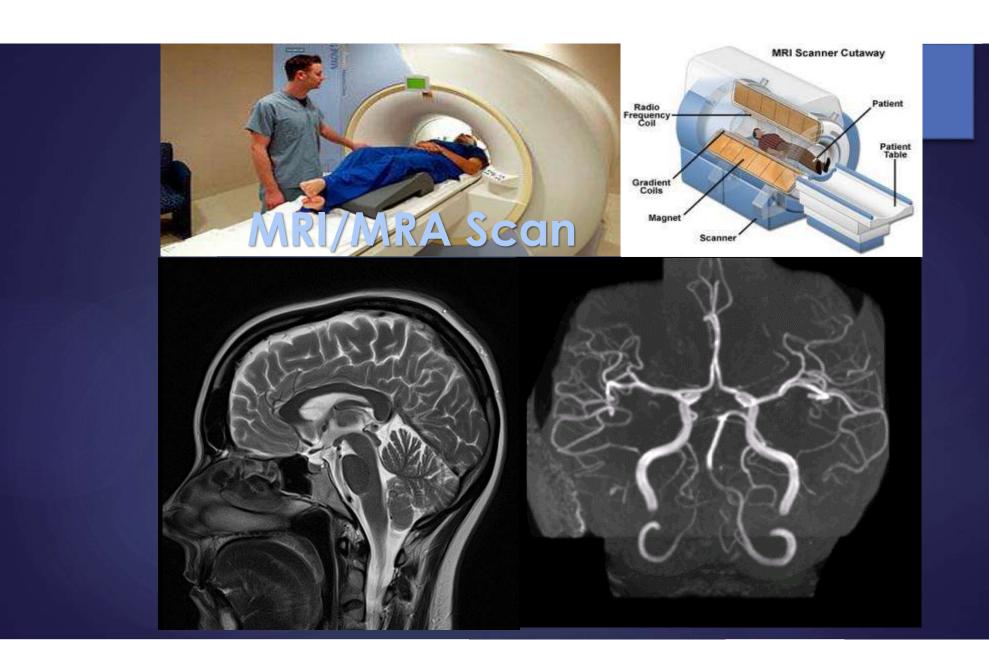








How do We Look at the Cerebrovascular anatomy?



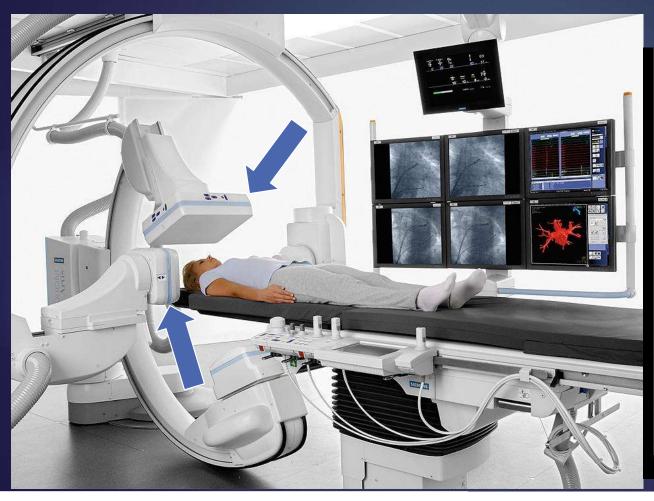
CT Scanner/ CT Angiogram

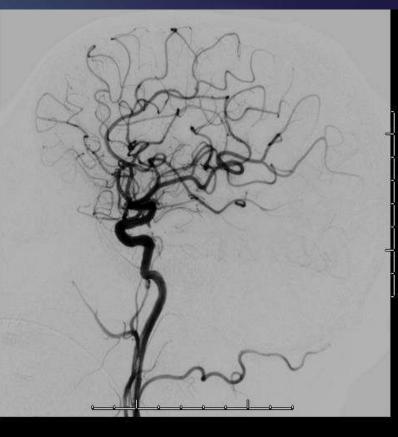


Biplane Angiography



How it Works



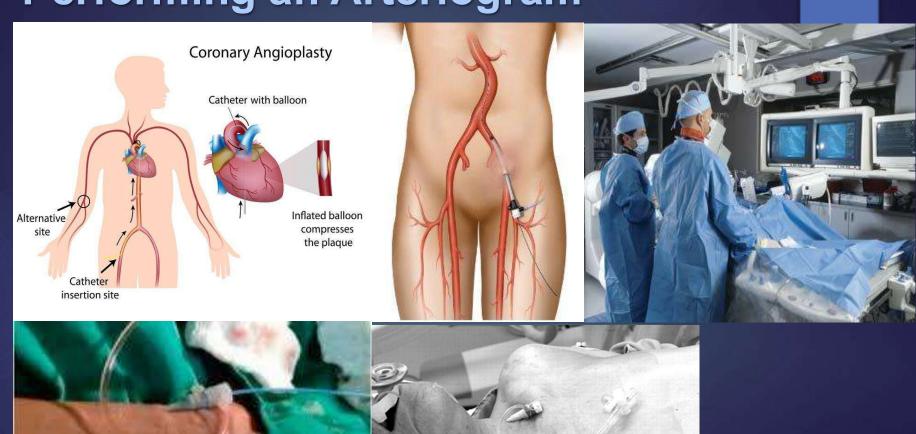


Rotational Angiography





Performing an Arteriogram



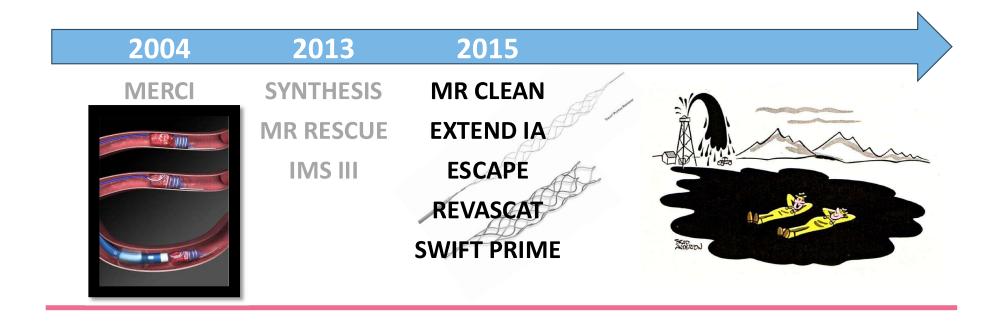




Objectives:

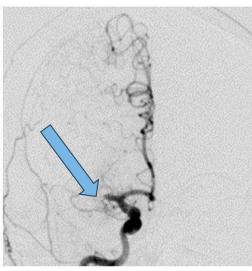
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Landmark EVT Trials – Timeline



Thrombectomy Basics – LVO



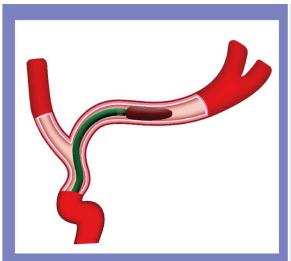


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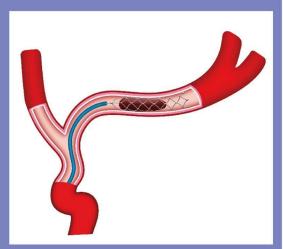
Thrombectomy Basics – LVO

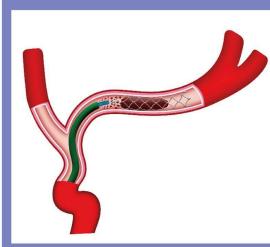
Stent Retriever

Combination (Solumbra)



Aspiration (ADAPT)

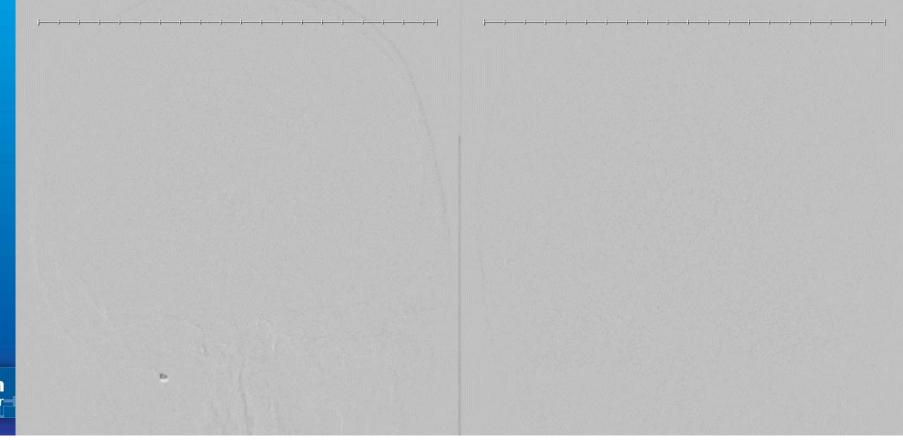




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J Am Coll Cardiol Intv. 2020 Jul, 13 (14) 1683-1696.

Endovascular Revascularization Treatment (EVT)





2015 – The Selection Era

Design of the Five Recent Randomized Controlle (12 hours)

Longest window (12 hours)

First to use CTP
Therap with RAPID

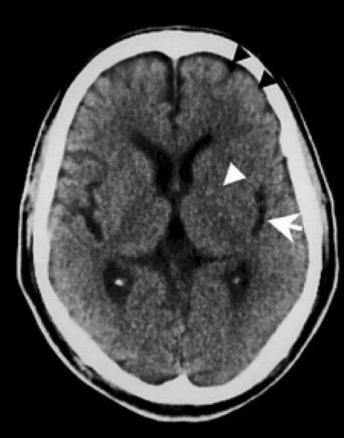
Discrepancies with ASPECTS scoring noted

	MR CLEAN	ESCAPE	EXTEND IA	SWIFT PRIME	RECAVSCAT
Design	Best medical care vs. Best medical care + EVT	Best medical care vs. Best medical care + EVT	IV rtPA vs. IV rtPA + EVT	IV rtPA vs. IV rtPA + EVT	Best medical care vs. Best medical care + EVT
Phase	Phase 3	Phase 3	Phase 2	Phase 3	Phase 3
Time	<6 hr, expected puncture tim	<12 hr, randomization	<6 hr, expected puncture time	<6 hr, expected puncture time	<8 hr, expected puncture time
EVT modality	Discretion of investigator	Recommended stent retriever	Exclusively Solitaire FR	Exclusively Solitaire FR	Exclusively Solitaire FR
Primary outcome			· Median percentage reperfusion at 24 h	90 days mRS	90 days mRS
			· NIHSS reduction >8 points or reaching 0-1 at 3 days		
Pretreatment Image	NCCT with CTA	NCCT with mCTA	NCCT with CTA and CTP	NCCT with CTA and CTP (or DWI with MRA and MRP) → NCCT with CTA (or DWI with MRA)	NCCT with CTA
Image selection criteria	Absent	Exclusion of large core (ASPECTS <6) & Exclusion of poor collateral ^a	Target mismatch ^b	Target mismatch ^c → Exclusion of large core (CT of DWI ASPECTS <6)	or Exclusion of large core (CT ASPECTS <7, DWI ASPECTS <6)
Premorbid condition	None	Barthel index >90	mRS1<1	mRS1≤1	mRS1≤1
Age	≥18	>18	≥18	18-80	18-80
NIHSS	≥2	≥6	None	8-29	≥6
Occlusive lesion	distal ICA, M1, M2, A1	distal ICA, M1, M1 equivalent	ICA, M1, M2	ICA, M1	ICA, M1
	Waited for tPA response before	sue / Clissue -	Highest % of mRS		arly EVT in patients id id () who did not respond to tPA

D Song, et al. Previous and recent evidence of endovascular therapy in acute ischemic stroke (Neurointervention 2015).

Early Ischemic Changes

ASPECT



- Loss of insular ribbon (←)
- Loss of gray-white interface (◀)
- Loss of sulci (▼)
- Acute hypodensity
- Mass effect
- Dense MCA sign

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Landmark EVT Trials – Timeline

2004	2013	2015	2016
MERCI	SYNTHESIS	MR CLEAN	THRACE
	MR RESCUE	EXTEND IA	THERAPY
	IMS III	ESCAPE	
		REVASCAT	
		SWIFT PRIME	

Landmark EVT Trials – Timeline

2004	2013	2015	2016	2017	
MERCI	SYNTHESIS	MR CLEAN	THRACE	PISTE	
	MR RESCUE	EXTEND IA	THERAPY	EASI	
	IMS III	ESCAPE			
		REVASCAT			
		SWIFT PRIME			

What Does the Evidence Show Us?

 What were the important lessons learned from multiple randomized clinical trials that showed thrombectomy was better than IV tPA in LVO cases...







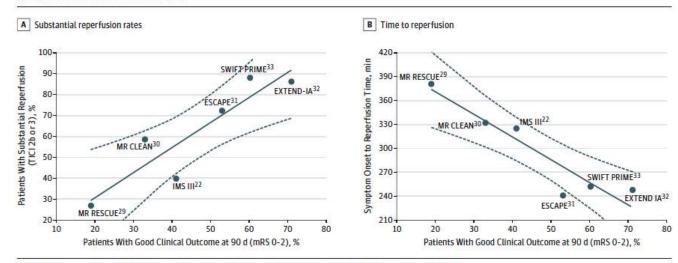
Review

Acute Stroke Intervention A Systematic Review

JAMA. 2015;313(14):1451-1462. doi:10.1001/jama.2015.3058

Shyam Prabhakaran, MD, MS; Ilana Ruff, MD; Richard A. Bernstein, MD, PhD

Figure 3. Rate of Reperfusion and Time to Reperfusion Compared With Percentage of Good Outcomes in the 6 Trials Comparing Endovascular Treatment to Medical Treatment Alone



mRS indicates modified Rankin Scale; TICI, thrombolysis in cerebral infarction. The dotted lines indicate 95% CIs.





Patients should receive endovascular therapy *with a stent retriever* if they meet all the following criteria:

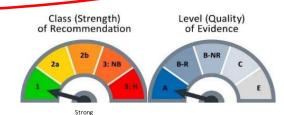
- pre-stroke modified Rankin score (mRS) 0-1
- acute ischemic stroke receiving IV rtPA within 4.5 hours of onset according to guidelines from professional medical societies,
- causative occlusion of the internal carotid artery or proximal middle cerebral artery (M1),
- age 18 years and over,
- National Institutes of Health Stroke Scale (NIHSS) score of 6 or greater,
- Alberta Stroke Program Early Computed Tomography Score (ASPECTS) of 6 or greater, and
- treatment can be initiated (groin puncture) within 6 hours of symptom onset

(Class I; Level of Evidence A).

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Recommendations

Powers WJ, Derdeyn CP, et al. Stroke. 2015;46:000-000.



European Stroke Organisation (ESO) - European Society for Minimally Invasive Neurological Therapy (ESMINT) Guidelines on Mechanical Thrombectomy in Acute Ischemic Stroke

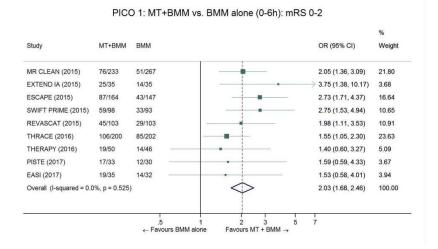


Figure 2 Pooled OR for functional independence in patients treated with MT+BMM versus BMM alone in the 0–6 hour time window. Random-effects meta-analysis. BMM, best medical management; MT, mechanical thrombectomy.

Recommendation

In adults with anterior circulation large vessel occlusion-related acute ischemic stroke presenting within 6 hours after symptom onset, we recommend mechanical thrombectomy plus best medical management—including intravenous thrombolysis whenever indicated—over best medical management alone to improve functional outcome.

Quality of evidence: **High** $\oplus \oplus \oplus \oplus$; strength of recommendation: **Strong** $\uparrow \uparrow$



Recommendations

G Turc, et al. European Stroke Organisation—European Society for Minimally Invasive Neurological Therapy expedited recommendation on indication for intravenous thrombolysis before mechanical thrombectomy in patients with acute ischaemic stroke and anterior circulation large vessel occlusion (European stroke journal 2022).

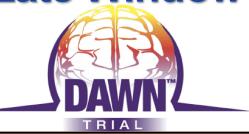


Expanding Indications

Landmark EVT Trials – Timeline

					INIAL
2004	2013	2015	2016	2017	2018
MERCI	SYNTHESIS	MR CLEAN	THRACE	PISTE	DAWN
	MR RESCUE	EXTEND IA	THERAPY	EASI	DEFUSE 3
	IMS III	ESCAPE			111
		REVASCAT			defuse 3
		SWIFT PRIME			

2018 - The Late Window Era



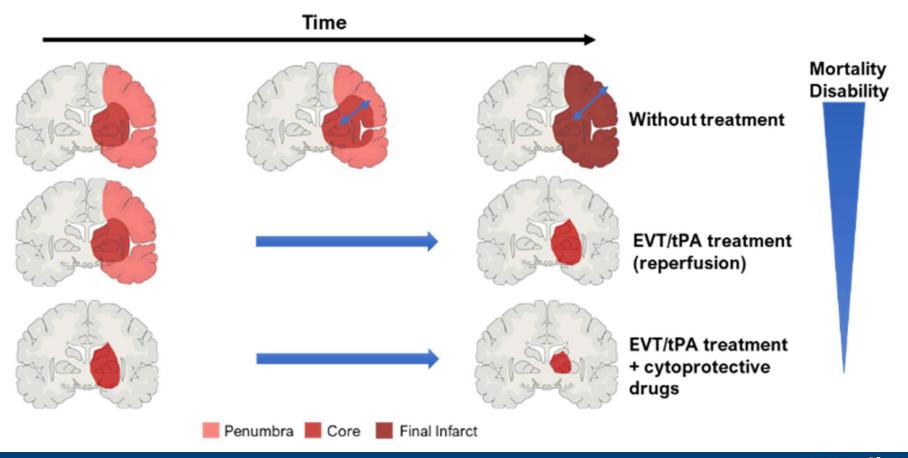


	DAWN	DEFUSE-3	
Eligibility	6-24 hours	6-16 hours	
Occlusion location	Intracranial ICA or proximal MCA	Extra- or intra-cranial ICA or proximal MCA	
Infarct volume	Age > 80 + NIHSS ≥ 10 \rightarrow < 21 mL Age < 80 + NIHSS ≥ 10 \rightarrow 31 mL Age < 80 + NIHSS ≥ 20 \rightarrow 31-51 mL	< 70 mL Mismatch ratio ≥ 1.8	
Imaging selection	CT or MRI RAPID software	CT or MRI RAPID software	
Thrombectomy device	TREVO (Stryker sponsored)	Any FDA approved device	



Nogueria, et al. (2018) N Engl J Med, 378:11-21 Albers, et al. (2018) N Engl J Med, 378:708-718

Stroke Progression



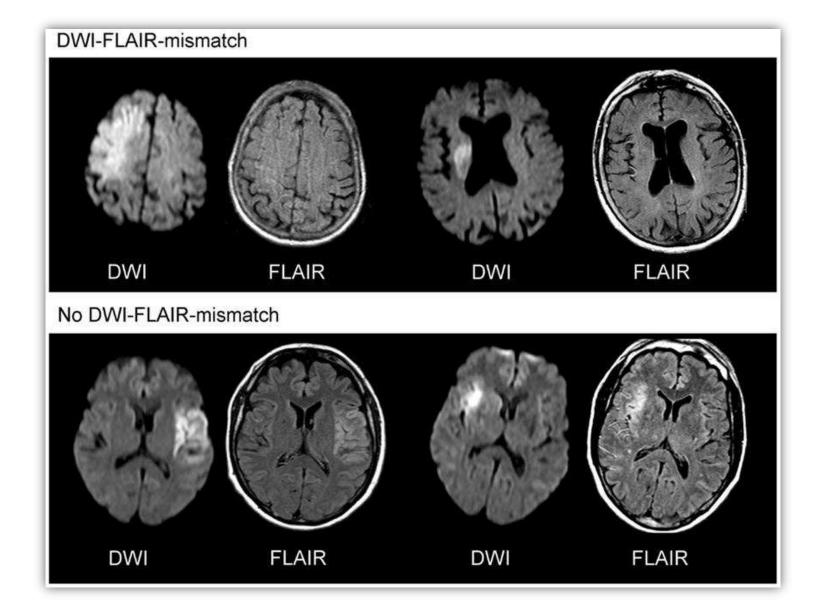
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core penumbra "CTA perfusion" **CTP MTT** NCCT CTP CBV NCCT CBF<30% volume: 67 ml Tmax>6.0s volume: 242 ml Mismatch volume: 175 ml Mismatch ratio: 3.6

CTP Penumbra Infarct Core

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DWI/FLAIR Mismatch



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AHA/ASA Guideline

Endovascular Interventions 6-24 hours

3.7 Mechanical Thrombectomy: Over 6 hours

DAWN and DEFUSE 3 Trials

- CT Perfusion, or MRI/MR perfusion to select patients with salvageable brain tissue, despite prolonged time from last normal
- Randomized to thrombectomy vs no-thrombectomy
- Both trials showed <u>large</u> benefit for thrombectomy
 - DAWN Trial: Good outcome (mRS 0-2) in 49% vs. 13%
 - DEFUSE 3 Trial: Good outcome (mRS 0-2) in 45% vs. 17%

Recommendations	COR	LOE
In selected patients with AIS onset within 6-16 hours, anterior circulation large vessel occlusion, and who meet other DAWN or DEFUSE 3 eligibility criteria, mechanical thrombectomy is recommended.	1	A
In selected patients with AIS within 6 to 24 hours of last known normal who have LVO in the anterior circulation and meet other DAWN eligibility criteria, mechanical thrombectomy is reasonable.	lla	B-R





Powers JP, Rabinstein AA, et al. Stroke. 2018;49:e46–e99. Powers JP, Rabinstein AA, et al. Stroke. 2019;50:e344–e418.

European Stroke Organisation (ESO) - European Society for Minimally Invasive Neurological Therapy (ESMINT) Guidelines on Mechanical Thrombectomy in **Acute Ischemic Stroke**

Recommendation

In adults with anterior circulation large vessel occlusion-related acute ischemic stroke presenting between 6 and 24 hours from time last known well and fulfilling the selection criteria of DEFUSE-3* or DAWN**, we recommend mechanical thrombectomy plus best medical management over best medical management alone to improve functional outcome.

Quality of evidence: Moderate $\oplus \oplus \oplus$; strength of recommendation: Strong \↑

(see below and table 3 regarding patient selection)

- **6 to 24 hours since time last known well:
- ⇒ See table 3 for details.age ≤80 years and NIHSS≥6: infarct core volume < 70 mL and penumbra volume > 15 mL and penumbra volume/core volume >1.8.
- \Rightarrow Age <80 years: infarct core ≤30 mL if NIHSS ≥10; infarct core \leq 51 mL if NIHSS \geq 20.
- \Rightarrow Age≥80 years: infarct core ≤20 mL and NIHSS ≥10.

Inclusion criteria	DEFUSE-3 ¹⁰	DAWN ⁹		
Time window	6–16 hours since time last known well	6–24 hours since time last known well		
Age	18–90 years	≥18 years		
mRS score before qualifying stroke	≤2; life expectancy ≥6 months	≤1; life expectancy ≥6 months		
NIHSS score	≥6	≥10 (see below)		
Arterial occlusion	ICA and/or M1*	ICA and/or M1		
Mismatch definition	Target mismatch profile on CT or MR perfusion imaging, as determined by an automated image postprocessing system: Infarct core volume <70 mL† AND mismatch volume >15 mL (Tmax>6 s‡) AND mismatch ratio (penumbra/core)	Clinical-imaging mismatch Age <80 years and NIHSS score ≥10 and infarct core 0-30 mL OR age <80 years and NIHSS score ≥20 and infarct core 31-51 mL OR age ≥80 years and NIHSS score ≥10 and infarct core 0-20 mL		

ICA, internal cerebral artery; MCA, middle cerebral artery; mRS, modified Rankin Scale; NIHSS, National Institutes of Health Stroke Scale,



Recommendations

G Turc, et al. European Stroke Organisation-European Society for Minimally Invasive Neurological Therapy expedited recommendation on indication for intravenous thrombolysis before mechanical thrombectomy in patients with acute ischaemic stroke and anterior circulation large vessel occlusion (European stroke journal 2022).

^{*6} to 16 hours since time last known well:

tBased on CT perfusion or MRI diffusion.

[‡]The size of the penumbra was estimated from the volume of tissue for which there was delayed arrival of an injected tracer agent (time to maximum of the residue function (Tmax) exceeding 6 s. 148

2019

SELECT

SELECT – prospective cohort

ERT vs BMM 105 total, ERT in 62 patients ASPECTS <5 or volume >50 ml

Findings: ERT **may** benefit patients with infarct-core volume of 50- 100 ml in secondary analysis.

Sarraj A, Hassan AE, et al. JAMA Neurol. 2019;76(10):1147-1156.

2019

2022

SELECT



2019 2022 2020 - 2022

SELECT

INSPIRE

BEST(2020)

BASICS(2020)

ATTENTION(2021)

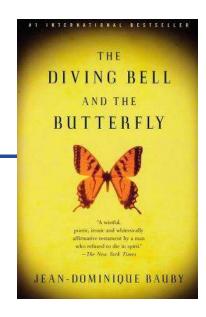
BAOCHE(2022)

BEST(2020)

Stroke Agenda

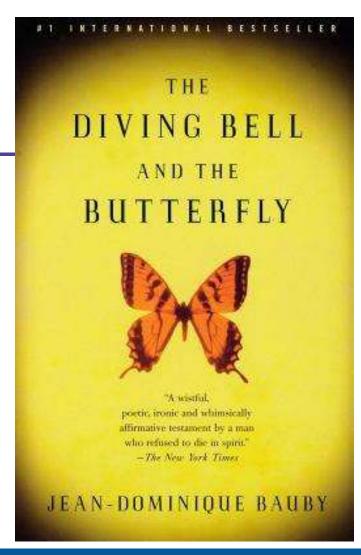
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BAO Outcomes

- Basilar artery occlusion (BAO) is a devastating stroke:
 - High mortality (85-95% untreated) and major disability.
 - Often leading to a devastating locked-in syndrome (paralysis with preserved consciousness) in survivors.
- EVT significantly improve chances of favorable outcomes in 24-40% of treated patients, emphasizing rapid treatment for better results.



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BAO Treatment

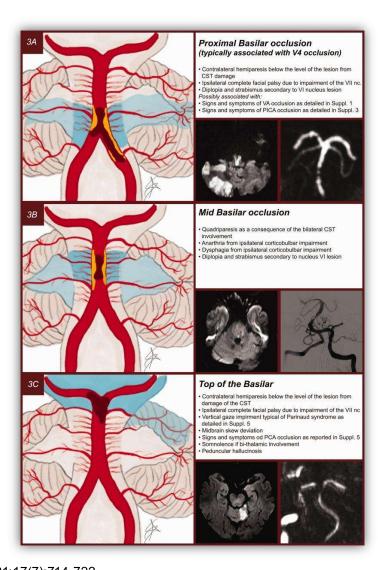




A significant percentage of posterior circulation strokes (PCS) are missed, with studies showing initial misdiagnosis rates from around **20% up to 40% or more**, often due to atypical symptoms like isolated dizziness, nausea, or headache, which are easily mistaken for less serious conditions.

BAO Etiology

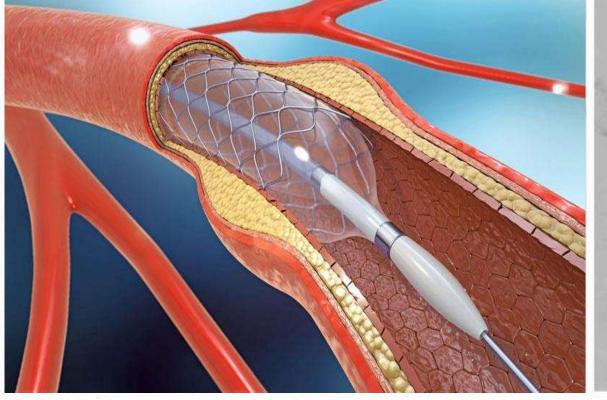


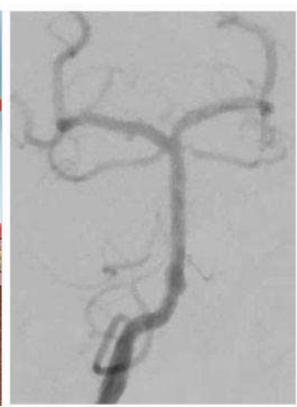




- International Journal of Stroke. 2021;17(7):714-722.
- neupsykey.com/part-iv-acute-stroke-procedures-5/

BA Stenosis







UTSouthwestern Medical Center	BEST ⁽²⁰²⁰⁾	BASICS ⁽²⁰²¹⁾	ATTENTION ⁽²⁰²¹⁾	BAOCHE ⁽²⁰²²⁾
Region	China	Europe and South America	China	China
Number of Patients (EVT, BMM)	66, 65	154, 146	226, 114	110, 107
Key Inclusion Criteri	ia			
Age (years)	≥18	≥18	≥18	≥18 and <u><</u> 80
NIHSS	Not specified	NIHSS score <u>></u> 10	<u>></u> 10	<u>></u> 6
Premorbid Disability (Baseline mRS)	0-2	0-2	0 for patients ≥80 years old 0-3 for patients <80 years old	<u><</u> 1

UTSouthwestern Medical Center	BEST (2020)	BASICS ⁽²⁰²¹⁾	ATTENTION ⁽²⁰²¹⁾	BAOCHE ⁽²⁰²²⁾
Occlusion Site	BA or functional V4 VA on CTA/MRA/DSA	BA on CTA/MRA	BA on CTA/MRA/DSA	BA or bilateral VA on CTA/MRA/DSA
Core Infarct Imaging Criteria	None None		PC-ASPECTS: ≥6 for patients <80 years ≥8 for patients ≥80 years	PC-ASPECTS score: ≥6 Pons-midbrain-index: ≤2
Time Window	8 hrs from the estimated time of occlusion	6 hrs from the estimated time of occlusion	12 hrs from the estimated time of occlusion	6-24 hrs from symptom onset
Crossover Rate	BMM to EVT: 22 % EVT to BMM: 5 %	BMM to EVT: 4.8 % EVT to BMM: 1.9 %	BMM to EVT: 2.6 % EVT to BMM: 1.3 %	BMM to EVT: 3.7% EVT to BMM: 0.9%
Etiology - Large artery atherosclerosis	EVT: 56% BMM: 49%	EVT: 36.3% BMM: 32.6%	EVT <u>:</u> 47.9% BMM <u>:</u> 36.9%	EVT: 68.2% BMM: 64.5%
Occlusion Site	BA or functional V4 VA on CTA/MRA/DSA	BA on CTA/MRA	BA on CTA/MRA/DSA	BA or bilateral VA on CTA/MRA/DSA

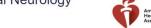
pcASPECT



Journal of Stroke 2023;25(1):55-71.

BAO EVT vs BMM

Stroke: Vascular and Interventional Neurology Volume 3, Issue 6, November 2023



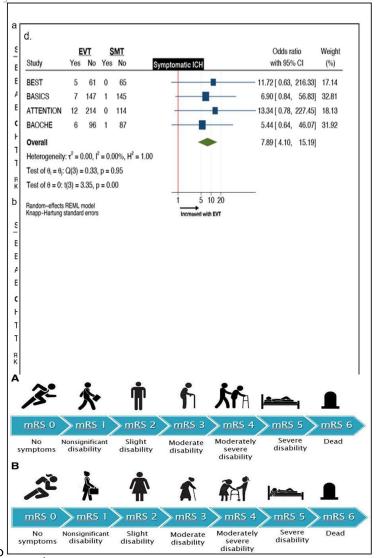
ORIGINAL RESEARCH

https://doi.org/10.1161/SVIN.123.000885

Endovascular Treatment in Acute Basilar Artery Occlusion Stroke: A Brief Practice Update From the Society of Vascular and Interventional Neurology

Kaiz S. Asif, MD, Robin Novakovic, MD, Thanh N. Nguyen, MD, Ossama Mansour, MD, Adam deHavenon, MD, Hesham E. Masoud, MD, Alicia C. Castonguay, PhD, Cynthia Kenmuir, MD, Yazan Ashouri, MD, Vallabh Janardhan, MD, David S. Liebeskind, MD, Ameer E. Hassan, MD, Amer M. Malik, MD, and Osama O. Zaidat, MD, MS for the SVIN GAPS Committee

- 90-day mRS score of ≤3 was significantly more likely in those treated with EVT (pooled OR, 1.99 [95% CI, 1.04–3.8]; P=0.042)
- Also, 90-day mRS score of 0 to 2 (pooled OR, 2.26 [95% CI, 0.78–6.57]; P=0.094).
- Mortality was significantly less likely (pooled OR, 0.64 [95% CI, 0.42–0.99]; P=0.048).
- sICH was significantly more likely with EVT (pooled OR, 7.89 [95% CI, 4.1–15.19]; P=0.002)



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K Asif, R Novakovic, et al. Stroke Vasc Interv Neuro

BAO RCTs Results

- •BAO RCTs demonstrated **EVT significantly** improved functional outcomes and reduced mortality compared to BMM when performed within 24 hours of stroke onset.
- While early trials like BEST and BASICS showed inconclusive results, more recent trials
 (ATTENTION, BAOCHE) in Chinese populations confirmed EVT's effectiveness, showing much higher rates of good functional recovery and sustained long-term benefits (e.g., 46% vs. 23% good outcomes at 90 days in ATTENTION).





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Summary of Recommendations Based on Clinical Questions

1. Should EVT be performed in patients presenting with BAO within 12 hours of symptom onset?

EVT is moderately recommended in patients presenting with BAO within 12 hours from symptom onset and meeting the following criteria: age 18 to 80 years, prestroke mRS score of 0 to 2, NIHSS score ≥10, and pc-ASPECTS ≥8 (COR 2a, LOE B-R, revised RoB analysis domain 6: bidb risk, EO for COR: EO-C)

Recommendation	COR	LOE	Expert opinion consensus on COR
EVT is moderately recommended in patients presenting with BAO within 12 hours from onset and meeting the following criteria: age 18 to 30 years, prestroke mRS score of 0 to 2, NIHSS score ≥10, and pc-ASPECTS ≥8.	2a	B-R	EO-C

2. Should EVT be performed in patients presenting with BAO between 12 and 24 hours of symptom onset?

EVT is moderately recommended in patients presenting with BAO between 12 and 24 hours from symptom onset and meeting the following criteria: age 18 to 80 years, prestroke mRS score of 0 to 1, NIHSS score ≥10, and pc-ASPECTS ≥8 (COR 2a, LOE B-R, revised RoB analysis domain 6: high risk, EO for COR: EO-C).

Recommendation	COR	LOE	Expert opinion consensus on COR
EVT is moderately recommended in patients presenting with BAO between 12 and 24 hours from onset and meeting the following criteria: age 18 to 80 years, prestroke mRS score 0 to 1, NIHSS score ≥10, and pc-ASPECTS ≥8.	2a	B-R	EO-C

3. Should EVT be performed in patients presenting with BAO attributable to large-artery atherosclerosis?

EVT is moderately recommended in patients presenting with BAO attributable to largeartery atherosclerosis presenting within 24 hours from symptom onset and meeting the following criteria: age 18 to 80 years, prestroke mRS score of 0 to 1, NIHSS score ≥10, and pc-ASPECTS ≥8 (COR 2a, LOE B-R, revised RoB analysis domain 6: high risk, EO for COR: EO.C.)

Recommendation	COR	LOE	Expert opinion consensus on COR
EVT is moderately recommended in patients presenting with BAO attributable to large-artery atherosclerosis presenting within 24 hours from onset and meeting the following criteria: age 18 to 80 years, prestroke mRS score of 0 to 1 NIHSS score ≥10, and pc-ASPECTS ≥8.	2a	B-R	EO-C

Stroke: Vascular and Interventional Neurology

Volume 3, Issue 6, November 2023 https://doi.org/10.1161/SVIN.123.000885



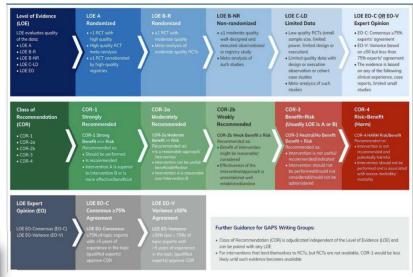
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K Asif, R Novakovic, et al. Stroke Vasc Interv Neurol. 2023;3:e000885.



4. Should EVT be performed in basilar artery occlusion with pc-ASPECTS score 6 to 7? EVT may be considered in patients with a pc-ASPECTS score of 6 to 7 presenting with BAO and meeting the following criteria: symptom onset within 24 hours, age <80 years, prestroke mRS score of 0 to 1, and NIHSS score ≥10 (COR 2b, LOE C-LD, revised RoB analysis domain 6: high risk FO for COR: FO-C)

Recommendation	COR	LOE	Expert opinion consensus on COR
EVT may be considered in patients with a pc-ASPECTS of 6 to 7 presenting with BAO and meeting the following criteria: onset within 24 hours, age <80 years, prestroke mRS score of 0 to 1, and NIHSS score ≥10.	2b	C- LD	EO-C

5. Should EVT be performed in basilar artery occlusion with patients aged >80 years? EVT may be considered in patients aged >80 years presenting with BAO and meeting the following criteria: onset within 12 hours, prestroke mRS score of 0, pc-ASPECTS ≥8, and NIHSS score ≥10 (COR 2b, LOE C-LD, revised RoB analysis domain 6: high risk, EO for COR: EO-C)

Recommendation	COR	LOE	Expert opinion consensus on COR
EVT should be considered in patients aged >80 years presenting with BAO and meeting the following criteria: onset within 12 hours, prestroke mRS score of 0, pc-ASPECTS ≥8, and NIHSS score ≥10.	2b	C- LD	EO-C



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- BAO lower-level recommendation awarded:
 - □>80 years
 - ✓ Within 12 hours
 - ✓pc-ASPECTS ≥8
 - ✓ Prestroke mRS 0
 - ✓NIHSS ≥10

K Asif, R Novakovic, et al. Stroke Vasc Interv Neurol. 2023;3:e000885.



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2019	2022	2023	
SELECT	INSPIRE	RESCUE-J LIMIT	Mantle
	BEST	SELECT2	
	BASICS	ANGEL-ASPECT	
	ATTENTION	TESLA	Large Core
	BAOCHE	TENSION	
	BEST	LASTE	Outer Molten Core

https://www.nasa.gov/solar-system/a-closer-look-at-mercurys-spin-and-gravity-reveals-the-planets-inner-solid-core/



Stroke Agenda

Objectives:

- Stroke reality and treatment options for ischemic stroke
- Once upon a thrombectomy..... a long, long, time ago!
 - ✓ Indication for thrombectomy in anterior circulation LVO
- Could there be a worse fate? Basilar artery occlusion
 - ✓ Indication for thrombectomy in BAO
- When is enough enough? Large core infarcts in anterior LVO
 - ✓ Indication for thrombectomy large core anterior LVO
- How far can we go? MeVO and DMVO
 - ✓ Indication for thrombectomy in MeVO??? Wait is there any?



	RESCUE-JAPAN- LIMIT	ANGEL- ASPECT	SELECT2	TESLA	TENSION	LASTE
	Japan	China	North America, Australia	United States	Europe, Canada	Europe, United States
Inclusion Criteria						
Age (years)	<u>≥</u> 18	18-80	18-80 18-85 18-85 <u>></u>		<u>≥</u> 18	<u>≥</u> 18
Pre mRS score	0-1	0-1	0-1	0-1 0-1		0-1
LKW/Symptom Onset	<6 hours or 6-24 hours MRI FLAIR negative	<24 hours	<24 hours	<24 hours <24 hours		<7 hours or MRI FLAIR negative
NIHSS	<u>></u> 6	6-30	<u>></u> 6	<u>></u> 6	<26	>6
Site of Occlusion	ICA or M1	Intracranial ICA or M1	ICA or M1	ICA terminus or M1	Intracranial ICA or M1	Intracranial ICA, M1, M1-M2
ASPECT Score	CT/MRI 3-5	CT 3-5	CT 3-5	CT 2-5	CT/MRI 3-5	CT/MRI 0-5 or ≥ 80 yrs 3 to 5
Additional Criteria		Or core 70-100 mL	Or core ≥50mL			

FUTSouthwestern												
Medical Center	RESCUE-JAPAN- LIMIT		ANGEL- ASPECT		SELECT2		TESLA		TENSION		LASTE	
Comparison Arms	EVT	вмм	EVT	BMM	EVT	BMM	EVT	вмм	EVT	вмм	EVT	вмм
#patients	101	102	230	225	178	174	152	148	125	128	159	165
Median Age			68	67	66	67	66	68	73	74	73	74
Mean Age	76.6	75.7										
Male sex	55	58	135	144	107	100	76	84	59	51		
Median NIHSS	22	22	16	15	19	19	19	18	19	18	21	21
Median ASPECT	3 (3-4)	4 (3-4)	3 (3-4)	3 (3-4)	4 (3-5)	4 (4-5)	4 (3-5)	4 (3-5)	0-2: 12% 3-5: 82% 6-10: 6%	18% 78% 4%	<3: 54% 3-5: 46%	58% 52%
Median Infarct Volume (mL)	94	110	61	63	82	79	166	171	206*	228*	132	137
Median (min) Symptom Onset/LKW to Randomization	229	214	453	463	545	588	653	744	120	126	271	268

EVT: endovascular thrombectomy; BMM: best medical management; ASPECT: Alberta Stroke Program Early Computed Tomographic Score; NIHSS: National Institutes of Health Stroke Scale; * at 24 hours

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Comparison Arms	EVT (N=100)	BMM (N=102)	EVT (N=230)	BMM (N=225)	EVT	вмм	EVT	вмм	EVT (N=122)	BMM (N=123)	EVT	вмм
90-Day mRS 0-3	31% (31)	12.7% (13)	47% (108)	33.3% (75)	37.9% (67/177)	18.7% (32/171)	30% (45/151)	20% (29/146)	31.5%	13.1%	33.3% (53/159)	12.7% (21/165)
90-Day Mortality	18% (18)	23.5% (24)	21.7% (50)	20% (45)	38.4% (68/177)	41.5% (71/171)	35.3% (53/150)	33.3% (49/147)	40% (49)	51% (63)	36.1% (57/158)	55.5% (91/164)
24-48 Hrs sICH	9% (9)	4.9% (5)	6.1% (14)	2.7% (6)	0.6% (1/178)	1.1% (2/174)	4% (6/151)	1.3% (2/149)	6% (7)	5% (6)	9.6% (15/157)	5.7% (9/157)

SELECT2 https://clinicaltrials.gov/study/NCT03876457?tab=results LASTE – Tudor J live presentation SLICE 2023 ANGEL-ASPECT N Engl J Med 2023;388:1272-83. RESCUE-JAPAN LIMIT N Engl J Med 2022;386:1303-13.

TENSION Lancet 2023; 402: 1753–63

TESLA https://ssrn.com/abstract=4587818 – preprint not peer reviewed

UTSouthwestern Medical Center		-JAPAN- ∕IIT	ANC ASP		SELE	ECT2	TES	SLA	TENS	SION	LAS	STE
Comparison Arms	EVT (N=100)	BMM (N=102)	EVT (N=230)	BMM (N=225)	EVT	вмм	EVT	вмм	EVT (N=122)	BMM (N=123)	EVT	вмм
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TESLA https://ssrn.com/abstract=4587818 – preprint not peer reviewed

Meta-Analysis Large Core RCTs

Systematic review

Endovascular thrombectomy for large ischemic strokes: meta-analysis of six multicenter randomized controlled trials

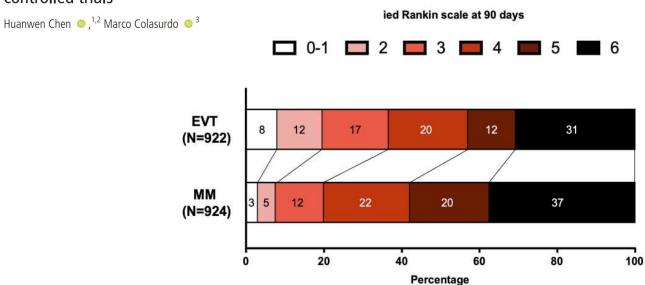


Figure 1 Pooled 90-day modified Rankin Scale outcomes from RESCUE-Japan-LIMIT, ANGEL-ASPECT, SELECT2, TESLA, TENSION, and LASTE. EVT, endovascular thrombectomy; MM, medical management.

H Chen, et al. Endovascular thrombectomy for large ischemic strokes: meta-analysis of six multicenter randomized controlled trials (*Journal of neurointerventional surgery* 2025).



Meta-Analysis Large Core RCTs

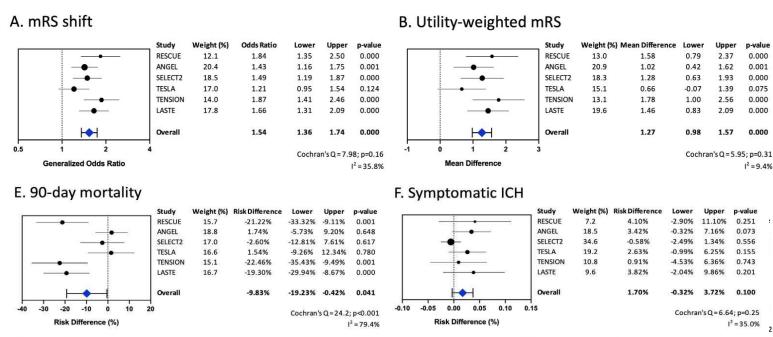


Figure 2 Pooled outcome measures across six randomized controlled trials comparing endovascular thrombectomy with medical management for patients with large ischemic strokes. ICH, intracranial hemorrhage; mRS, modified Rankin Scale.

H Chen, et al. Endovascular thrombectomy for large ischemic strokes: meta-analysis of six multicenter randomized controlled trials (*Journal of neurointerventional surgery* 2025).



Endovascular therapy for anterior circulation emergent large vessel occlusion stroke in patients with large ischemic cores: a report of the SNIS Standards and Guidelines Committee

Fawaz Al-Mufti , ¹ Franklin A Marden, ² Jan Karl Burkhardt, ³ Daniel Raper, ⁴ Clemens M Schirmer , ⁵ Amanda Baker , ⁶ Peng Roc Chen , ⁷ Ketan R Bulsara, ⁸ Kazim H Narsinh , ⁹ Matthew Robert Amans , ⁶ Jared Cooper, ¹⁰ Shadi Yaghi, ^{11,12} Mais Al-Kawaz , ^{13,14} Steven W Hetts , ¹⁵ SNIS Standards and Guidelines Committee. SNIS Board of Directors

Recommendation 1: In patients with anterior circulation ELVO who present within 24 hours of last known normal (LKN) with large infarct core (70–149 mL or ASPECTS 3–5) and meet other criteria of RESCUE-Japan LIMIT, SELECT2, ANGEL-ASPECT, TESLA, TENSION, or LASTE trials, thrombectomy is indicated (Class I, Level A)

Recommendation 2: EVT in patients with LCS aged 18–80 years (5 RCT evidence) and 80–85 years of age (4 RCT evidence) is beneficial (Class I, Level A)

Recommendation 3: EVT in patients with LCS >85 years of age (2 RCT) may be beneficial (Class I, Level B-R)

Recommendation 4: Patients with LCS and NIHSS 6–30 (5 RCT evidence) benefit from EVT in LCS (Class I, Level A)

Recommendation 5: Patients with LCS and NIHSS <6 and >30 (2 RCT evidence) may benefit from EVT in LCS (Class IIa, Level A)

Recommendation 6: Patients with LCS and low baseline mRS (0-1) (5 RCT evidence) benefit from EVT (Class I, Level A)

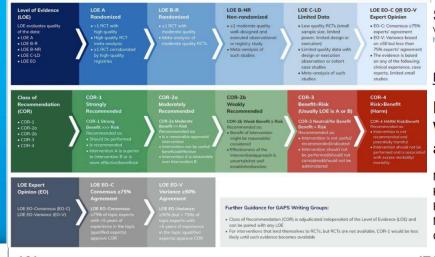
Recommendation 7: Patients with LCS and time of last known well 0–24 hours (<6 hours with 3 RCT evidence and 6–24 hours with 5 RCT evidence) benefit from EVT (Class I, Level A)

Recommendation 8: It is recommended that patients with LCS who also meet the criteria for on-label use of IV tPA receive IV tPA irrespective of whether endovascular treatments are being considered (Class I; Level B-NR)



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F Al-Mufti, et al. Endovascular therapy for anterior circulation emergent large vessel occlusion stroke in patients with large ischemic cores: a report of the SNIS Standards and Guidelines Committee (*Journal of neurointerventional surgery* 2024).



Stroke: Vascular and Interventional Neurology

Volume 5, Issue 2, March 2025

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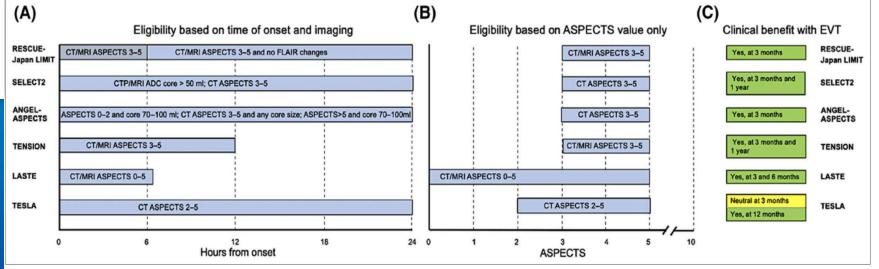


REVIEW ARTICLE

Endovascular Therapy in Patients With Acute Ischemic Stroke With Large Infarct: A Guideline From the Society of Vascular and Interventional Neurology

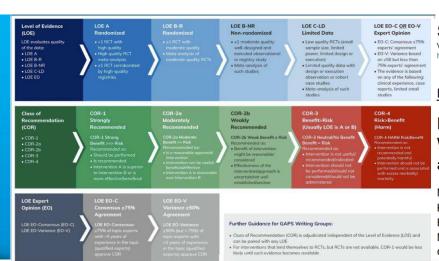
Maxim Mokin, MD, PhD (i), Tudor G. Jovin, MD, Sunil A. Sheth, MD, Thanh N. Nguyen, MD, Kaiz S. Asif, MD, Ameer E. Hassan, DO, Ashutosh P. Jadhav, MD, PhD, Cynthia Kenmuir, MD, PhD, David S. Liebeskind, MD, Ossama Mansour, MD, Raul G. Nogueira, MD, Robin Novakovic, MD, Santiago Ortega-Gutierrez, MD, MS, Albert J. Yoo, MD, PhD, Waldo R. Guerrero, MD, and Amer M. Malik, MD for the SVIN GAPS committee





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Figure 2. Key imaging characteristics of 6 large infarct randomized trails. A, The 6 trials evaluated the safety and efficacy of EVT during different time periods from symptom onset to randomization and initiation of EVT. Key imaging metrics for each trial are shown. B, ASPECTS were used in all 6 trials. Upper and lower ASPECTS thresholds are shown. C, Overall clinical benefit was shown in 5 trials. One trial (TESLA) did not meet its primary end point at 3 months but showed clinical benefit at the 1-year mark. ADC indicates apparent diffusion coefficient; ANGEL-ASPECT, Endovascular Therapy in Acute Anterior Circulation Large Vessel Occlusive Patients With a Large Infarct Core; ASPECTS, Alberta Stroke Program Early CT score; CT, computed tomography, CTP, computed tomography perfusion; EVT, endovascular therapy; FLAIR, fluid-attenuated inversion recovery; LASTE, Large Stroke Therapy Evaluation; MRI, magnetic resonance imaging; RESCUE-Japan LIMIT, Recovery by Endovascular Stroke United Stroke; TENSION, Efficacy and Safety of Thrombectomy in Stroke With Extended Lesion and Extended Time Window; and TESLA, Thrombectomy for Emergent Salvage of Large Anterior Circulation Ischemic Stroke.



Stroke: Vascular and Interventional Neurology

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REVIEW ARTICLE

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EVT within 0–6 hours of stroke onset	Class of recommendation	Level of evidence
In patients with anterior circulation stroke presenting within 0–6 h from symptom onset, baseline mRS score 0–1, age 18–80 y old, occlusion of the ICA or MCA M1 segment, and ASPECTS of 0–5 on noncontrast CT or MRI, EVT is recommended. (Evidence: LASTE trial; meta-analyses AlMajali et al, Roman et al, Winkelmeier et al)	1	A

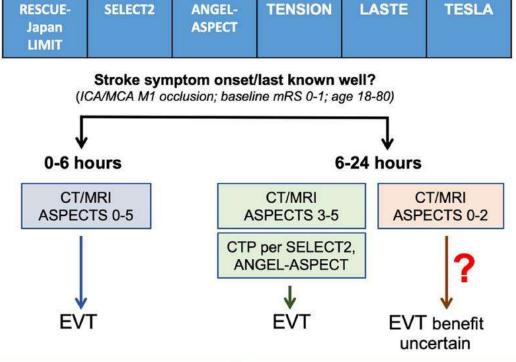
EVT within 6–24 hours of stroke onset	Class of recommendation	Level of evidence
In patients with anterior circulation LVO presenting within 6–24 h from symptom onset, baseline mRS score 0–1, age 18–80 years old, occlusion of the ICA or MCA M1 segment, and ASPECTS of 3–5 on noncontrast CT or MRI, EVT is recommended. (Evidence: ANGEL-ASPECT, RESCUE-Japan LIMIT; meta-analyses Palaiodimou et al, Wei et al, and Li et al)	1	A
In patients with anterior circulation LVO presenting within 6–24 h from symptom onset, baseline mRS score 0–1, age 18–80 years old, occlusion of the ICA or MCA M1 segment with CTP imaging according to the SELECT2 and ANGEL-ASPECT eligibility criteria, EVT is recommended. (Evidence: ANGEL-ASPECT, SELECT2)	1	A
In patients with anterior circulation LVO presenting within 6–24 h from symptom onset, baseline mRS score 0–1, age 18–80 years old, occlusion of the ICA or MCA M1 segment, and ASPECTS of 0–2 on noncontrast CT or MRI, the benefit of EVT is uncertain.	2b	B-R



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Endovascular therapy in acute ischemic stroke patients with large infarct: a guideline from the Society of Vascular and Interventional Neurology (SVIN)



- Despite increased hemorrhage rates
- Benefit of thrombectomy for large core infarcts, up to
 24hrs



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M Mokin, et al. Endovascular therapy in patients with acute ischemic stroke with large infarct: A guideline from the Society of vascular and Interventional Neurology (*Stroke: vascular and interventional neurology* 2025).

2019	2022	2023	2024	2025
SELECT	INSPIRE	RESCUE-J LIMIT		
	BEST	SELECT2	DISTALS	2026
	BASICS	ANGEL-ASPECT	The F	uture
	ATTENTION	TESLA	NEAT	
	BAOCHE	TENSION		
	BEST	LASTE		

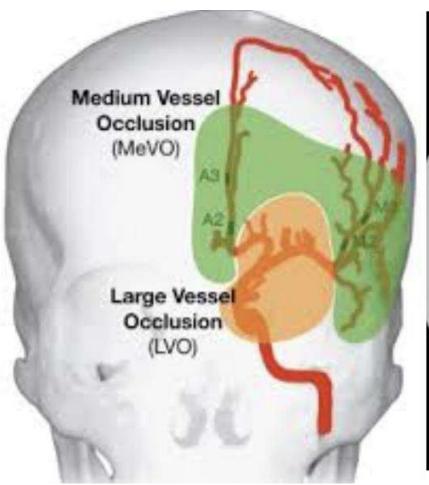


Stroke Agenda

Objectives:

- Stroke reality and treatment options for ischemic stroke
- Once upon a thrombectomy..... a long, long, time ago!
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- Could there be a worse fate? Basilar artery occlusion
 - ✓ Indication for thrombectomy in BAO
- When is enough enough? Large core infarcts in anterior LVO
 - ✓ Indication for thrombectomy large core anterior LVO
- How far can we go? MeVO and DMVO
 - ✓ Indication for thrombectomy in MeVO??? Wait is there any?

DMVO vs LVO





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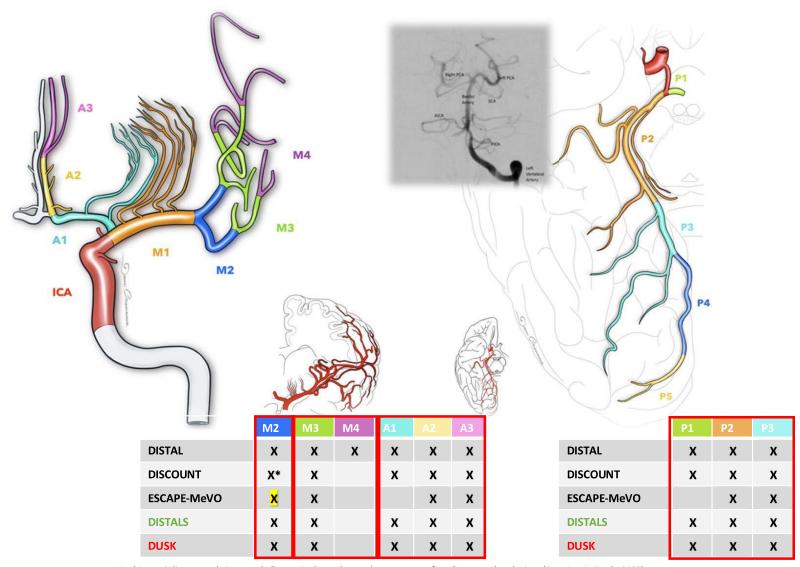
Distal Medium vs Medium Vessel DMVO/MeVO





DMVO Clinical Trials





A Rodriguez-Calienes, et al. Current challenges in the endovascular treatment of medium vessel occlusions (Frontiers in Stroke 2023).

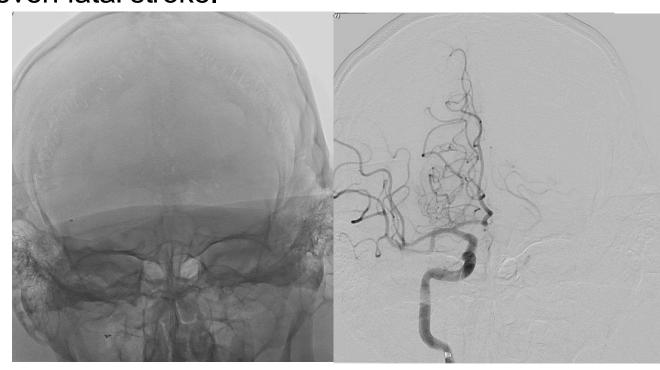
DMVO

- •DMVOs account for 25% to 40% of all AIS.
- Less deficits and disability caused by DMVO(median NIHSS 7) compared to LVO.
- But still, observational studies showed 1/3 do not achieve functional independence and excellent functional outcome in only 50% with best medical management.
- Mortality 9% at 90 days.
- Recanalization rates <50% at 2 to 6 hours with IV alteplase.
 - Duloquin G, et al. Stroke. 2020;51:2122–2130.
 - Saver JL, et al; Stroke. 2020;51:2872–2884.
 - Ospel JM, et al. Stroke. 2020;51:3232–3240.



Clinical Presentation

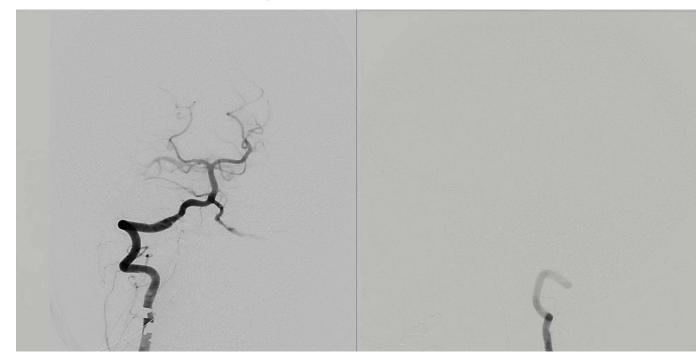
- MCA territory DMVOs most common (70%–85%).
 - oClinical manifestations highly variable, from clinically silent, to fleeting symptoms, to severely disabling or even fatal stroke.





Clinical Presentation

- PCA second most common site (~10% to 20%).
 - oContralateral homonymous hemianopia or (usually superior) quadrantanopia most common symptoms.
 - oLess often visual agnosias, alexia, achromatopsia.





Clinical Presentation

- ACA third most common site (~5%).
 - Contralateral leg weakness and sensory loss most common.

Frontal lobe infarcts present with behavioral

syndrome like abulia.



- JM Ospel, et al. Stroke. 2020;51:3232–3240.
- M Goyal et al. ESCAPE-MeVO investigators. N Engl J Med. 2025;392:1385–1395.
- M Psychogios, et al; DISTAL Investigators. N Engl J Med. 2025;392:1374–1384.
- WK Diprose, et al. Stroke. 2026;57:00–00. DOI: 10.1161/STROKEAHA.125.05040



3 RCTs Criteria

Table 1. Trial Protocols

Trial Name	DISCOUNT	DISTAL	ESCAPE-MeVO
Primary outcome	90-d mRS score 0-2	90-d mRS score (shift analysis)	90-d mRS score 0-1
Included occlusion locations	Distal M2/M3 MCA, A1/A2/A3 ACA, P1/P2/P3 PCA	Nondominant or codominant M2/M3/M4 MCA, A1/A2/A3 ACA, P1/P2/P3 PCA	M2/M3 MCA, A2/A3 ACA, P2/P3 PCA
Other imaging criteria	Absence of carotid tandem occlusion	0–6 h: No criteria; 6–24 h: Hypoperfusion-hypodensity mismatch or FLAIR-DWI mismatch	Evidence of salvageable tissue required (moderate collaterals, core–penumbra mismatch, or no extensive ischemic changes)
Included NIHSS score range	≥5 or significant aphasia	≥4 or disabling symptoms	>5 or 3-5 with disabling symptoms
Time frame (max from LKW to randomization)	≤8 h	≤24 h	≤12 h
Devices allowed	Trevo, pReset, CatchView Mini, 3MAX, 4MAX, 5MAX, Q3, Q5	Any commercially available device	Solitaire X as first-line device
Exclusion criteria	Large core infarct, spontaneous recanalization, prestroke mRS score > 1	Dominant M2 occlusion, nondisabling deficits, evidence of large infarcts, poor life expectancy	ASPECTS ≤ 8, living in nursing home or requiring daily nursing care or assistance with ADLs, or major comorbid illness

ADLs indicates activities of daily living; ASPECTS, Alberta Stroke Program Early Computed Tomography Score; DISCOUNT, Evaluation of Mechanical Thrombectomy in Acute Ischemic Stroke Related to a Distal Arterial Occlusion; DISTAL; Endovascular Therapy Plus Best Medical Treatment [BMT] Versus BMT Alone for Medium Vessel Occlusion Stroke; ESCAPE-MeVO, Endovascular Treatment to Improve Outcomes for Medium Vessel Occlusions; FLAIR DWI, diffusion-weighted imaging-fluid-attenuated inversion recovery; LKW, last known well; MCA, middle cerebral artery; mRS, modified Rankin Scale; and NIHSS, National Institutes of Health Stroke Scale.



3 RCTs

- •Studies found that the benefits of thrombectomy may have limits, particularly for medium and distal vessel occlusions.
- •3 trials found no clinical benefit EVT compared to BMM.

Table 3. Trial Outcome Measures

Outcome	DISCOUNT OR (95% CI)	P value	DISTAL OR (95% CI)	P value	ESCAPE-MeVO OR (95% CI)	P value
Primary outcome						
mRS score ≤ 2 at 90 d		0.024	0.90 (0.67-1.22)	1-2	0.92 (0.79-1.07)	1 1 - 1
mRS score ≤ 1 at 90 d	0.42 (0.2-0.88)					
Per protocol analysis (DISCOUNT)	0.30 (0.12–0.74)	0.009	-,	-	=	3-
Secondary outcomes						
mRS score 0-1 at 90 d	1,1-2,1	1775	0.88 (0.61-1.25)	<u>0</u> =2	0.97 (0.79-1.18)	1
Death at 90 d	-	_	1.17 (0.71-1.90)	1 - 2	1.82 (1.06-3.12)	0.03

Cl indicates confidence interval; DISCOUNT, Evaluation of Mechanical Thrombectomy in Acute Ischemic Stroke Related to a Distal Arterial Occlusion; DISTAL; Endovascular Therapy Plus Best Medical Treatment [BMT] Versus BMT Alone for Medium Vessel Occlusion Stroke; ESCAPE-MeVO, Endovascular Treatment to Improve Outcomes for Medium Vessel Occlusions; MeVO, medium vessel occlusions; mRS, modified Rankin Scale; and OR, odds ratio.



3 RCTs

- ESCAPE-MeVO higher mortality in the EVT group (13.3%) vs. BMM group (8.4%) (adjusted hazard ratio (adj HR) of 1.82 (95% Cl, 1.06–3.12).
- **DISTAL no significant difference** in mortality between groups (15.5% EVT group vs. 14.0% BMT group) (odds ratio [OR] 1.17 [95% CI, 0.71–1.90]).
- **DISCOUNT lower mortality rate in the EVT group** (3%) vs. BMT arm (7%).

Table 3. Trial Outcome Measures

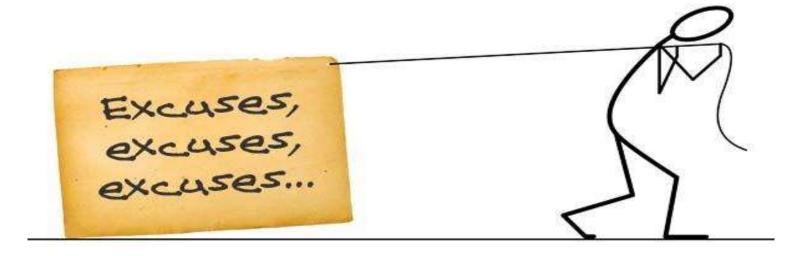
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Primary outcome						
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Secondary outcomes					į.	
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The Defense

•Any time trials fail to show expected benefit or negative you are going to hear why it failed.





Median age significantly older than LVO RCTs (67-71 yo)

Factors that might impact outcomes

Characteristic	DISCOUNT EVT + BMT (N=64)	DISCOUNT BMT alone (N=97)	DISTAL EVT + BMT (N=271)	DISTAL BMT alone (N=272)	ESCAPE-MeVO EVT + usual care (N=257)	ESCAPE-MeVO usual care (N=272)
Age, y (median, IQR)	75 (11)	72 (13)	77 (68–83)	77.5 (68–84)	74 (63–82)	76 (65–83)
Female sex (%)	37 (46%)	38 (48%)	116 (42.8%)	123 (45.2%)	118 (46.3%)	127 (46.4%)
NIHSS (median, IQR)	8 (6–11)	8 (6–12)	6 (5–9)	6 (5–9)	8 (6–11)	7 (5–11)
Intravenous thrombolysis (%)	57 (70%)	57 (71%)	168 (62.0%)	187 (68.8%)	144 (56.5%)	165 (60.2%)
Serious adverse events (%)	25 (39%)	30 (31%)	114 (42.1%)	88 (32.3%)	87 (33.9%)	70 (25.7%)
Symptomatic intracranial hemorrhage (%)	8 (12%)	6 (6%)	16 (5.9%)	7 (2.6%)	14 (5.4%)	6 (2.2%)
Death at 90 days (%)	2 (3%)	7 (7%)	42 (15.5%)	38 (14.0%)	34 (13.3%)	23 (8.4%)
modified Rankin Scale score 0-2 at 90 days (%)	45 (60%)	59 (77%)	153 (56.5%)	145 (54.7%)	138 (54.1%)	161 (58.8%)
modified Rankin Scale score 0-1 at 90 days (%)	a a		94 (34.7%)	101 (37.5%)	106 (41.6%)	118 (43.1%)
Time from symptom onset to final recanalization (min [IQR])		-		-	358 (233–530)	-
Time from last known well to puncture (median [IQR])	240 [192-312]	(3)	292.0 [177.7–640.5)		-	
TICI 2b-3 (%)	49 (77%)	8.5	194 (71.7%		190 (75.1%)	1 85
Procedural technique		(98)			8	(*)
Combined approach			148 (64.6%)			
Aspiration only	22 (34.4%)		36 (15.7%)			
Stent retriever only	39 (61.0%)		36 (15.7%)		257 (100.0%)	
Anesthesia modality	-	SH2		-		(**
General	Ö		129 (49.6%)		105 (41.3%)	
Sedation			1975)		98 (38.6%)	
Local anesthesia			(/#./		51 (20.1%)	
Occlusion site						
MCA: distal M2 or M3	62 (76%)	60 (75%)	-	-	-	
MCA: proximal M2	=	N=	S#1	-	64 (25.3%)	58 (21.6%)



Higher rates IV thrombolysis than seen in real world registries (40.8 to 44.6%)

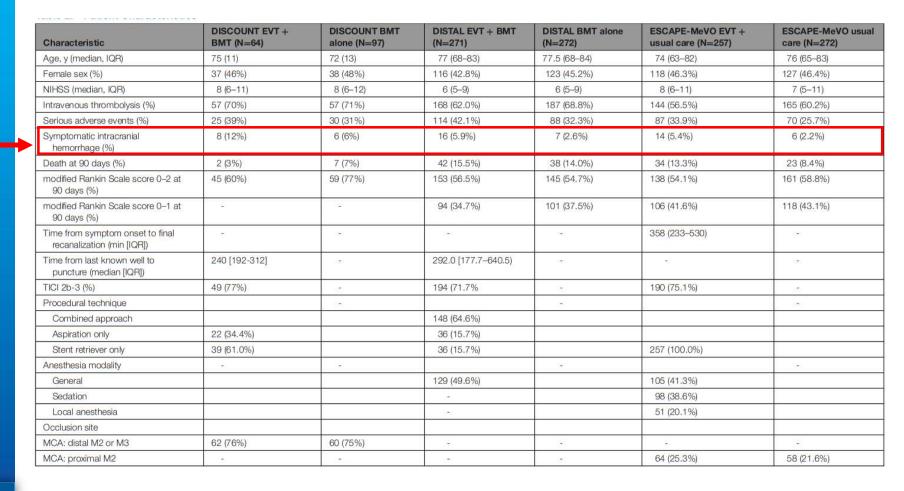
IV TNK/tPA

Characteristic	DISCOUNT EVT + BMT (N=64)	DISCOUNT BMT alone (N=97)	DISTAL EVT + BMT (N=271)	DISTAL BMT alone (N=272)	ESCAPE-MeVO EVT + usual care (N=257)	ESCAPE-MeVO usual care (N=272)
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modified Rankin Scale score 0-1 at 90 days (%)	El El		94 (34.7%)	101 (37.5%)	106 (41.6%)	118 (43.1%)
Time from symptom onset to final recanalization (min [IQR])	-1	-	-		358 (233–530)	-
Time from last known well to puncture (median [IQR])	240 [192-312]	(8)	292.0 [177.7–640.5)		-	(e
TICI 2b-3 (%)	49 (77%)	85	194 (71.7%		190 (75.1%)	
Procedural technique		(s e)			8	(*)
Combined approach			148 (64.6%)			
Aspiration only	22 (34.4%)		36 (15.7%)			
Stent retriever only	39 (61.0%)		36 (15.7%)		257 (100.0%)	
Anesthesia modality		(*)		-		
General	in the second se		129 (49.6%)		105 (41.3%)	
Sedation	2		973		98 (38.6%)	
Local anesthesia			(6)		51 (20.1%)	
Occlusion site						
MCA: distal M2 or M3	62 (76%)	60 (75%)	-	-	8	-
MCA: proximal M2			S = 1		64 (25.3%)	58 (21.6%)



Higher rates of sICH

sICH





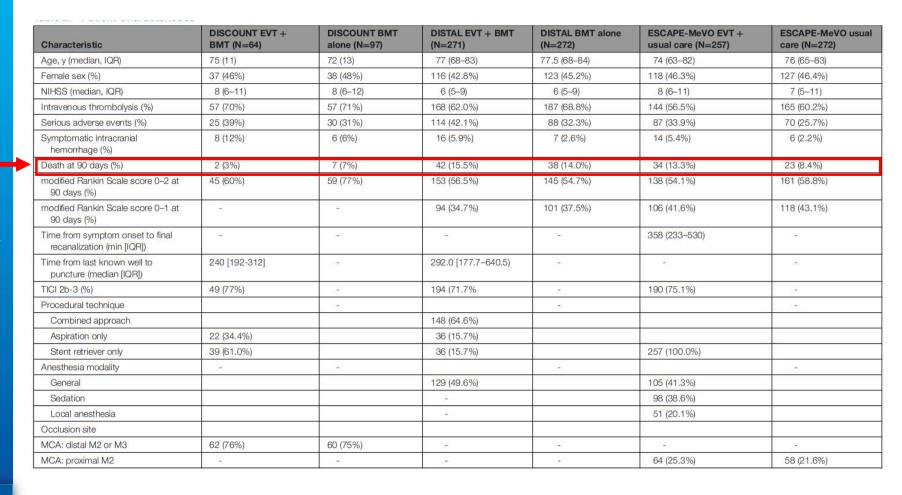
Despite this finding overall mortality did not significantly differ Mortality

Characteristic	DISCOUNT EVT + BMT (N=64)	DISCOUNT BMT alone (N=97)	DISTAL EVT + BMT (N=271)	DISTAL BMT alone (N=272)	ESCAPE-MeVO EVT + usual care (N=257)	ESCAPE-MeVO usual care (N=272)
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modified Rankin Scale score 0-1 at 90 days (%)	El El		94 (34.7%)	101 (37.5%)	106 (41.6%)	118 (43.1%)
Time from symptom onset to final recanalization (min [IQR])	-9	-	-	-	358 (233–530)	-
Time from last known well to puncture (median [IQR])	240 [192-312]	((#)	292.0 [177.7–640.5)		5	(E)
TICI 2b-3 (%)	49 (77%)	85	194 (71.7%		190 (75.1%)	1 85
Procedural technique		(se)			8	(0.5)
Combined approach			148 (64.6%)			
Aspiration only	22 (34.4%)		36 (15.7%)			
Stent retriever only	39 (61.0%)		36 (15.7%)		257 (100.0%)	
Anesthesia modality	-	(14)		-	, c	(Se)
General	Ġ.		129 (49.6%)		105 (41.3%)	
Sedation			0.73		98 (38.6%)	
Local anesthesia			((*)		51 (20.1%)	
Occlusion site						
MCA: distal M2 or M3	62 (76%)	60 (75%)	-		-	
MCA: proximal M2	5) * :) *		64 (25.3%)	58 (21.6%)



Suggesting not the sole driver of morbidity and mortality

Mortality





Proximal M2

Only ESCAPE-MeVO included proximal M2 =

UTSouthwestern Medical Center

Characteristic	DISCOUNT EVT + BMT (N=64)	DISCOUNT BMT alone (N=97)	DISTAL EVT + BMT (N=271)	DISTAL BMT alone (N=272)	ESCAPE-MeVO EVT + usual care (N=257)	ESCAPE-MeVO usua care (N=272)
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Time from symptom onset to final recanalization (min [IQR])	-9		-	-	358 (233–530)	
Time from last known well to puncture (median [IQR])	240 [192-312]	(2)	292.0 [177.7–640.5)	-	-1) e
TICI 2b-3 (%)	49 (77%)	85	194 (71.7%		190 (75.1%)	i sa
Procedural technique		(18)				(m)
Combined approach			148 (64.6%)			
Aspiration only	22 (34.4%)		36 (15.7%)			
Stent retriever only	39 (61.0%)		36 (15.7%)		257 (100.0%)	
Anesthesia modality	-	(*)		-		
General	ii		129 (49.6%)		105 (41.3%)	
Sedation			1975		98 (38.6%)	
Local anesthesia			(5 11)		51 (20.1%)	
Occlusion site						
MCA: distal M2 or M3	62 (76%)	60 (75%)		•	8	-
MCA: proximal M2)(#)		64 (25.3%)	58 (21.6%)

DISTAL only included patients with co-dominant or nondominant M2

DMVO

Table 2. (Continued)

Characteristic	DISCOUNT EVT + BMT (N=64)	DISCOUNT BMT alone (N=97)	DISTAL EVT + BMT (N=271)	DISTAL BMT alone (N=272)	ESCAPE-MeVO EVT + usual care (N=257)	ESCAPE-MeVO usual care (N=272)
MCA: distal M2	(*)	-			63 (24.9%)	41 (15.2)
MCA: M2 segment	100		129 (47.6%)	110 (40.4%)	*	
MCA: M3 segment	-		62 (22.9%)	84 (30.9%)	90 (35.6%)	126 (46.8%)
MCA: M4 segment			3 (1.1%)	0 (0.0%)	e e	
PCA: P1, P2, P3	13 (16%)	14 (18%)		34	÷	a
PCA: P1 segment	-		17 (6.3%)	13 (4.8%)	*	
PCA: P2 segment	-		32 (11.8%)	41 (15.1%)	24 (9.5%)	26 (9.7%)
PCA: P3 segment	-		6 (2.2%)	11 (4.0%)	0 (0.0%)	6 (2.2%)
ACA: A1, A2, A3	6 (7%)	6 (8%)	-	<u>.</u>	*	
ACA: A1 segment	-		0 (0.0%)	1 (0.4%)	*	-
ACA: A2 segment		-	11 (4.1%)	5 (1.8%)	4 (1.6%)	9 (3.3%)
ACA: A3 segment		7-	9 (3.3%)	5 (1.8%)	8 (3.2%)	3 (1.1%

Values in this table were extracted from published data in New England Journal of Medicine articles and presentations from the International Stroke Conference. Missing data were left blank. Where categories did not converge exactly, they were included as individual rows. For instance, when occlusion site definitions differed across studies, the most specific category available was used in separate rows to preserve granularity. For procedural technique, if a combination strategy was not explicitly reported, the first-line device was assumed to be used in isolation. The most commonly reported serious adverse events in ESCAPE-MeVO were pneumonia, recurrent stroke, worsening of index stroke, and symptomatic intracranial hemorrhage. In DISTAL, serious adverse events most commonly included pneumonia, recurrent ischemic stroke, any intracranial hemorrhage, and a broad "other" category. The "other" category encompassed various medical complications largely unrelated to EVT, such as sepsis, endocarditis, post-stroke depression, and gastric ulcer. Detailed characterization of serious adverse events in the DISCOUNT trial has not yet been published.

ACA indicates anterior cerebral artery; BMT, best medical treatment; DISCOUNT, Evaluation of Mechanical Thrombectomy in Acute Ischemic Stroke Related to a Distal Arterial Occlusion; DISTAL; Endovascular Therapy Plus Best Medical Treatment [BMT] Versus BMT Alone for Medium Vessel Occlusion Stroke; ESCAPE-MeVO, Endovascular Treatment to Improve Outcomes for Medium Vessel Occlusions; EVT, endovascular thrombectomy; IQR, interquartile range; MCA, middle cerebral artery; MeVO, medium distal vessel occlusions; NIHSS, National Institutes of Health Stroke Scale; PCA, posterior cerebral artery; and TICI, thrombolysis in cerebral infarction.





Proximal M2

- Although no dedicated RCT confirmed benefit of EVT for proximal or dominant M2, existing data suggest benefit.
- There may have been lack of equipoise during patient selection and patients may have not been enrolled in the trials (suggested by overall low NIHSS in the trials).
- Thus, the trial results for M2 occlusions should be interpreted with caution.
- Data from the 3 RCTs may be more relevant to nondominant, codominant, and distal M2 occlusions.



TICI score

Lower rates of successful reperfusion compared to post-hoc analysis of LVC trials



Characteristic	DISCOUNT EVT + BMT (N=64)	DISCOUNT BMT alone (N=97)	DISTAL EVT + BMT (N=271)	DISTAL BMT alone (N=272)	ESCAPE-MeVO EVT + usual care (N=257)	ESCAPE-MeVO usual care (N=272)
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Intravenous thrombolysis (%)	57 (70%)	57 (71%)	168 (62.0%)	187 (68.8%)	144 (56.5%)	165 (60.2%)
Serious adverse events (%)	25 (39%)	30 (31%)	114 (42.1%)	88 (32.3%)	87 (33.9%)	70 (25.7%)
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modified Rankin Scale score 0-1 at 90 days (%)	21	~	94 (34.7%)	101 (37.5%)	106 (41.6%)	118 (43.1%)
Time from symptom onset to final recanalization (min [IQR])			-	-	358 (233–530)	-
Time from last known well to puncture (median [IQR])	240 [192-312]	(2)	292.0 [177.7–640.5)	100		(=)
TICI 2b-3 (%)	49 (77%)	85	194 (71.7%		190 (75.1%)	-
Procedural technique		(s =)				(#)
Combined approach			148 (64.6%)			
Aspiration only	22 (34.4%)		36 (15.7%)			
Stent retriever only	39 (61.0%)		36 (15.7%)		257 (100.0%)	
Anesthesia modality	-	(*)		-		
General	ä		129 (49.6%)		105 (41.3%)	
Sedation	2		075		98 (38.6%)	
Local anesthesia			(96)		51 (20.1%)	
Occlusion site						
MCA: distal M2 or M3	62 (76%)	60 (75%)		-	-	•
MCA: proximal M2)(#)		64 (25.3%)	58 (21.6%)

3 RCTs

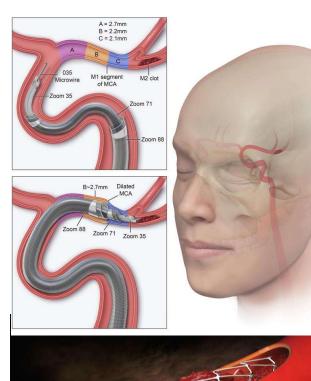
Unclear if due to technical limitations of devices, anatomy or operator bias

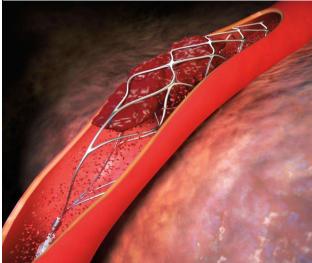


Characteristic	DISCOUNT EVT + BMT (N=64)	DISCOUNT BMT alone (N=97)	DISTAL EVT + BMT (N=271)	DISTAL BMT alone (N=272)	ESCAPE-MeVO EVT + usual care (N=257)	ESCAPE-MeVO usual care (N=272)
Age, y (median, IQR)	75 (11)	72 (13)	77 (68–83)	77.5 (68–84)	74 (63–82)	76 (65–83)
Female sex (%)	37 (46%)	38 (48%)	116 (42.8%)	123 (45.2%)	118 (46.3%)	127 (46.4%)
NIHSS (median, IQR)	8 (6–11)	8 (6–12)	6 (5–9)	6 (5–9)	8 (6–11)	7 (5–11)
Intravenous thrombolysis (%)	57 (70%)	57 (71%)	168 (62.0%)	187 (68.8%)	144 (56.5%)	165 (60.2%)
Serious adverse events (%)	25 (39%)	30 (31%)	114 (42.1%)	88 (32.3%)	87 (33.9%)	70 (25.7%)
Symptomatic intracranial hemorrhage (%)	8 (12%)	6 (6%)	16 (5.9%)	7 (2.6%)	14 (5.4%)	6 (2.2%)
Death at 90 days (%)	2 (3%)	7 (7%)	42 (15.5%)	38 (14.0%)	34 (13.3%)	23 (8.4%)
modified Rankin Scale score 0-2 at 90 days (%)	45 (60%)	59 (77%)	153 (56.5%)	145 (54.7%)	138 (54.1%)	161 (58.8%)
modified Rankin Scale score 0-1 at 90 days (%)	21	~	94 (34.7%)	101 (37.5%)	106 (41.6%)	118 (43.1%)
Time from symptom onset to final recanalization (min [IQR])			-	-	358 (233–530)	-
Time from last known well to puncture (median [IQR])	240 [192-312]	(2)	292.0 [177.7–640.5)	100		(=)
TICI 2b-3 (%)	49 (77%)	85	194 (71.7%		190 (75.1%)	-
Procedural technique		(s =)				(#)
Combined approach			148 (64.6%)			
Aspiration only	22 (34.4%)		36 (15.7%)			
Stent retriever only	39 (61.0%)		36 (15.7%)		257 (100.0%)	
Anesthesia modality	-	(*)		-		
General	ä		129 (49.6%)		105 (41.3%)	
Sedation	2		075		98 (38.6%)	
Local anesthesia			(96)		51 (20.1%)	
Occlusion site						
MCA: distal M2 or M3	62 (76%)	60 (75%)		-	-	•
MCA: proximal M2)(#)		64 (25.3%)	58 (21.6%)

EVT Devices

- **ESCAPE-MeVO** and **DISTAL** trials, stent retrievers were used as the first-line treatment.
- Stents highly effective for LVOs, but may carry increased procedural risks in smaller, more tortuous arteries, such as those involved in DMVOs.
- Direct aspiration as a first-line approach was permitted in **DISTAL** (15.7% of enrolled) and **DISCOUNT** (34% of enrolled) but underrepresented in all 3 RCTs.
- Maybe we need better devices to go distal!!!







Recommendations Coming

- These trials were subject to selection bias and other methodological limitations, and there remains a suggestion that certain subgroups of patients may still benefit from EVT.
- Until further data emerge, physicians should take pause when considering offering EVT for patients who are older with less severe deficits and higher baseline disability.
- Further research should concentrate on procedural techniques, specific anatomical location of occlusion, rates of thrombolysis, and differences between real-world populations and trial populations, as these factors affect outcomes.
- M Goyal, et al. Endovascular Treatment of Stroke Due to Medium-Vessel Occlusion. N Engl J Med. 2025;392:1385-1395.
- M Psychogios, et al. Endovascular Treatment for Stroke Due to Occlusion of Medium or Distal Vessels. N Engl J Med. 2025;392:1374-1384.
- F Clarencon, et al. Evaluation of mechanical thrombectomy in acute ischemic stroke related to a distal arterial occlusion: A randomized controlled trial. *Int J Stroke*. 2024;19:367-372.



Future Trials

- Additional RCTs and registries underway on DMVO.
- Including: DISTALS, STEP, ORIENTAL MeVO and DUSK.
- Methodologies vary:
 - ODISTALS to evaluate the safety and effectiveness of the Tigertriever 13 Revascularization, includes a distinct endpoint of successful reperfusion (assessed on 24-hour imaging) without sICH.
 - Excluded patients who received intravenous thrombolysis and differentiated M2s, only including nondominant or codominant M2 occlusions.
 - **STEP trial**, a randomized, multifactorial, adaptive platform registry-based trial.
 - Includes a unique end point of utility-weighted mRS and neurologic deficit (NIHSS) at 24 hours (±12 hours).
- Distal Ischemic Stroke Treatment With Adjustable Low-profile Stentriever (DISTALS). [cited 2025 February 11]; Available from: https://clinicaltrials.gov/study/NCT05152524
- StrokeNet Thrombectomy Endovascular Platform (STEP). [cited 2025 February 11]; Available from: https://clinicaltrials.gov/study/ NCT06289985
- Evaluation of Endovascular Treatment in Acute Intracranial Distal Medium Vessel Occlusion Stroke. [cited 2025 February 11];
 Available from: https://clinicaltrials.gov/study/NCT06146790
- MH Mohammaden, et al. Endovascular versus medical management in distal medium vessel occlusion stroke: the DUSK Study. Stroke. 2024;55(6):1489–1497.





3.7.2. 0 to 6 Hours From Onset (Continued)

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Endovascular Interventions MeVo and DMVO

AHA/ASA Guideline

Guidelines for the Early Management of Patients With Acute Ischemic Stroke: 2019 Update to the 2018 Guidelines for the Early Management of Acute Ischemic Stroke

A Guideline for Healthcare Professionals From the American Heart Association/American Stroke Association

LOE

COR

New, Revised, or Unchanged

3. Although the benefits are uncertain, the use of mechanical thrombectomy with stent retrievers may be reasonable for carefully selected patients with AIS in whom treatment can be initiated (groin puncture) within 6 hours of symptom onset and who have causative occlusion of the MCA segment 2 (M2)	llb	B-R	Recommendation reworded for clarity from 2015 Endovascular. COR unchanged. LOE revised. See Table XCV in online Data
or MCA segment 3 (M3) portion of the MCAs.			Supplement 1 for original wording.
In pooled patient-level data from 5 trials (HERMES, which included the 5 trials MR CLEA SWIFT PRIME, and EXTEND-IA), the direction of treatment effect for mechanical thromber was favorable in M2 occlusions, but the adjusted cOR was not significant (1.28 [95% Cl level data pooled from trials in which the Solitaire was the only or the predominant devia meta-analysis (SEER Collaboration: SWIFT PRIME, ESCAPE, EXTEND-IA, and REVASCAT) of treatment effect was favorable for mechanical thrombectomy over standard care in MOR and 95% Cl were not significant. ¹⁹⁰ In an analysis of pooled data from SWIFT (Solitain Thrombectomy), STAR (Solitaire Flow Restoration Thrombectomy for Acute Revasculariz among patients with M2 occlusions, reperfusion was associated with excellent functions OR, 2.2 [95% CI, 1.0–4.7]). ¹⁹⁵ Therefore, the recommendation for mechanical thrombect does not change substantively from the 2015 AHA/ASA focused update.	ectomy over sta , 0.51–3.21]). ¹⁸ ce used, a pres showed that th 12 occlusions, b re With the Inte tation), DEFUSE al outcomes (m	Indard care In patient- pecified In edirection But the Intion for	See Tables XVII and XLV in online Data Supplement 1.

Powers JP, Rabinstein AA, et al. Stroke. 2019;50:e344-e418.

Standards

European Stroke Organisation (ESO) - European Society for Minimally Invasive Neurological Therapy (ESMINT) Guidelines on Mechanical Thrombectomy in Acute Ischemic Stroke

Expert opinion on mechanical thrombectomy for M2 occlusion

There is a consensus among the guideline group (11/11 votes) that patients with M2 occlusion fulfilled the inclusion criteria in most randomized trials and therefore mechanical thrombectomy is reasonable in this situation.

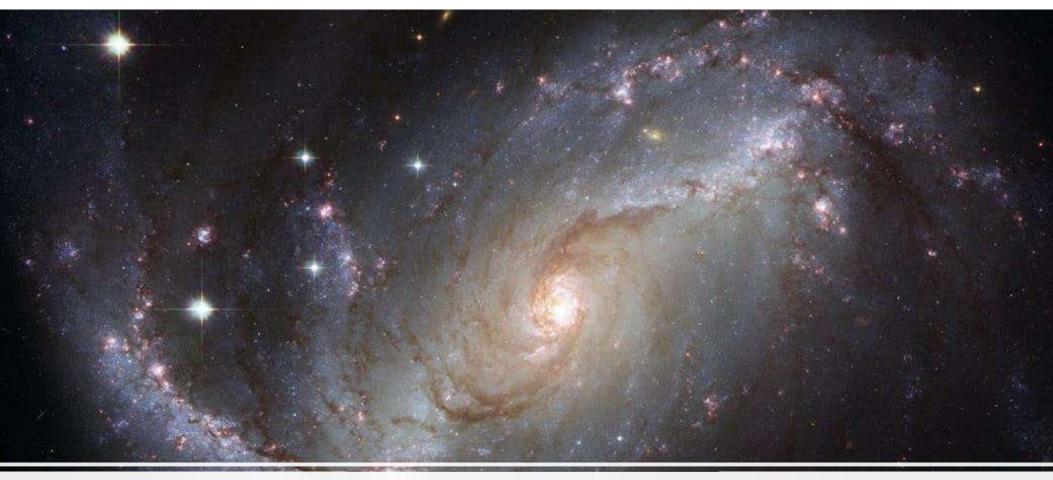


G Turc, et al. European Stroke Organisation—European Society for Minimally Invasive Neurological Therapy expedited recommendation on indication for intravenous thrombolysis before mechanical thrombectomy in patients with acute ischaemic stroke and anterior circulation large vessel occlusion (*European stroke journal* 2022).

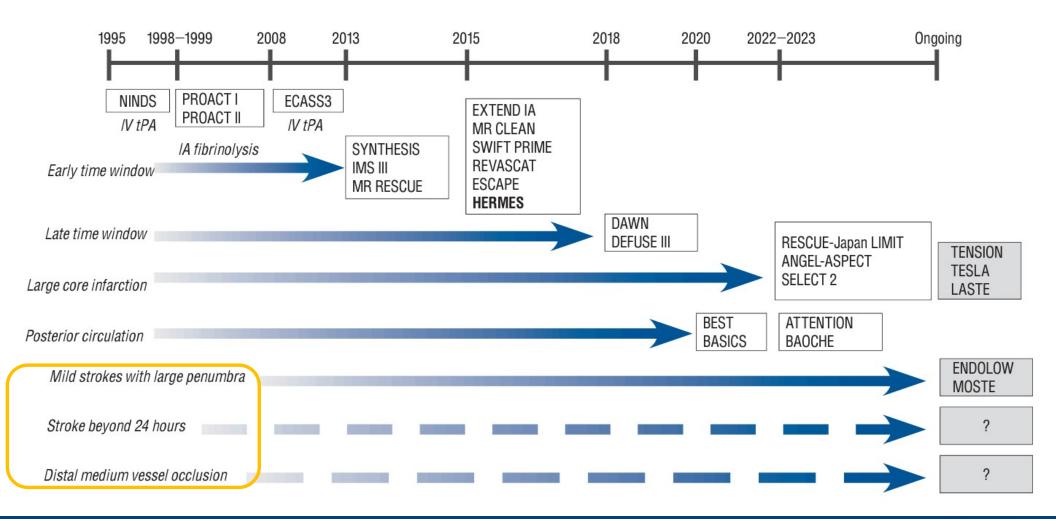


Vascular Territory	Recommendation	COR	LOE
M2 (MCA Division)	EVT is reasonable for disabling M2 occlusions—especially dominant or proximal branches—with favorable imaging with CT, CTP or MRI. Routine EVT for nondominant or low-severity M2 occlusions is not recommended. Additional randomized trial data is needed.		B-NR/A
M3/M4 (Distal MCA)	Routine EVT with mechanical devices is not recommended for distal M3/M4 occlusions due to lack of benefit and potential harm. EVT may be considered in patients with severe disabling deficits, favorable imaging, and poor response or contraindication to IV thrombolysis. Additional randomized trial data are needed.	III (no benefit) / IIb (select	B-R / C- LD
ACA (A2/A3)	Routine EVT is not recommended for isolated distal ACA (A2/A3) occlusions due to lack of demonstrated benefit. EVT may be considered for patients with disabling deficits (e.g., profound leg weakness, abulia, bilateral involvement), large territory at risk, favorable imaging, and good baseline functional status. Additional randomized trial data is needed.	III (no benefit) /	B-NR
PCA (P2/P3)	Routine EVT for isolated PCA (P1/P2/P3) occlusions is not recommended due to lack of demonstrated clinical benefit and possible increased mortality risk. EVT may be considered for patients with severe, disabling neurological deficits significantly impacting quality of life or those unable to receive IV thrombolysis. Additional randomized trial data is needed.	III (no benefit) / IIb (select cases)	B-NR

UTSouthwestern Medical Center



Expanding Access



Brain Institute

EUROPEAN STROKE JOURNAL

Tenecteplase in central retinal artery occlusion study (TenCRAOS): protocol for a randomized-controlled trial

Can tenecteplase (TNK) improve outcomes of central retinal artery occlusion (CRAO)?

Study design



Prospective RCT, doubledummy, double-blind phase 3 multi-centre trial of TNK 0.25 mg/kg + placebo vs. ASA + placebo



Population: CRAO within 4.5 hours from onset



Randomization: 2 arms , 1:1 block randomisation



Sample size: 78

Diagnosis of CRAO performed by ophthalmologist Patient screening by ophthalmologist and stroke specialist

Eligibility criteria checked + informed consent

Randomization (1:1)

TNK 0.25 mg/kg + placebo vs. ASA + placebo Monitoring in stroke unit >24 hours

Post-treatment visit at 24 (18–36) hours
In the stroke unit

30-day Primary endpoint 90-day follow-up Primary endpoint

Best-corrected visual acuity ≤ 0.7 logMAR (decimal best-corrected visual acuity ≥ 0.2) at 30-day



JAMA Neurology

RCT: Intra-Arterial Tenecteplase After Successful Reperfusion in Large Vessel Occlusion Stroke

POPULATION

113 Males, 92 Females



Adults with large vessel occlusion stroke and successful reperfusion within 24 h

INTERVENTION

205 Patients randomized and analyzed



48 Tenecteplase, 0.0313 mg/kg

Intra-arterial tenecteplase at a dose of 0.0313 mg/kg after successful reperfusion

48 Tenecteplase, 0.0625 mg/kg

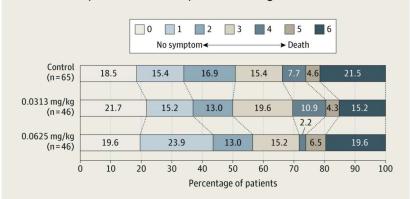
Intra-arterial tenecteplase at a dose of 0.0625 mg/kg after successful reperfusion

65 Control

Ended the procedure without further intra-arterial thrombolysis

FINDINGS

Adjunctive intra-arterial tenecteplase in doses of 0.0313 mg/kg or 0.0625 mg/kg showed adequate safety to advance to larger trials to determine the potential benefits in patients with large vessel occlusion stroke



SETTINGS/LOCATIONS



30 Stroke centers in China

PRIMARY OUTCOME

The primary outcome was the proportion of patients with a nondisabled outcome at 90 d (as indicated by modified Rankin Scale [mRS] score O-1; range, O [no symptoms] to 6 [death])

Phase 2a, mRS score 0-1

Tenecteplase, 0.0313 mg/kg, group:

Adjusted risk ratio [RR] vs control, 0.85; 95% CI, 0.54-1.35; P = .50

Tenecteplase, 0.0625 mg/kg, group:

Adjusted RR vs control, 1.15; 95% CI, 0.73-1.80; P = .55

Hou X, Huang J, Wang L, et al. Intra-arterial tenecteplase after successful reperfusion in large vessel occlusion stroke: a randomized clinical trial. *JAMA Neurol*. Published online July 5, 2025. doi: 10.1001/jamaneurol.2025.2036

