



OPEN ACCESS

EDITED BY

Susanna Bacigaluppi,
University of Genoa, Italy

REVIEWED BY

Mikito Hayakawa,
University of Tsukuba, Japan
Benjamin Y. Q. Tan,
National University Hospital, Singapore

*CORRESPONDENCE

Philipp Taussky
✉ ptaussky@bidmc.harvard.edu

RECEIVED 08 March 2025

ACCEPTED 04 July 2025

PUBLISHED 29 July 2025

CORRECTED 30 July 2025

CITATION

Mensah EO, Fong Y-W, Muram S, Ogilvy CS
and Taussky P (2025) The evolution of
personalized stroke thrombectomy.
Front. Surg. 12:1590146.
doi: 10.3389/fsurg.2025.1590146

COPYRIGHT

© 2025 Mensah, Fong, Muram, Ogilvy and
Taussky. This is an open-access article
distributed under the terms of the [Creative
Commons Attribution License \(CC BY\)](#). The
use, distribution or reproduction in other
forums is permitted, provided the original
author(s) and the copyright owner(s) are
credited and that the original publication in
this journal is cited, in accordance with
accepted academic practice. No use,
distribution or reproduction is permitted
which does not comply with these terms.

The evolution of personalized stroke thrombectomy

Emmanuel O. Mensah¹, Yew-Weng Fong^{1,2}, Sandeep Muram¹,
Christopher S. Ogilvy¹ and Philipp Taussky^{1*}

¹Neurosurgical Service, Beth Israel Deaconess Medical Center and Harvard Medical School, Boston, MA, United States, ²Department of Neurosurgery, Cathay General Hospital, Taipei, Taiwan

Stroke is a leading cause of disability and death worldwide, with acute ischemic stroke accounting for most cases. Mechanical thrombectomy is a widely accepted treatment modality in appropriately selected patients, demonstrating improved functional outcomes through safe and effective recanalization. However, traditional trials have focused on a narrow subset of patients, limiting its applicability to diverse populations who would otherwise benefit from thrombectomy. Advances in neurovascular imaging, device innovation, and procedural techniques are driving a paradigm shift toward personalized stroke thrombectomy. This review explores personalization strategies across various domains, including lesion-specific considerations such as medium vessel occlusions (MeVOs), basilar artery occlusions (BAOs), and tandem lesions, as well as patient-specific factors like infarct size, low NIHSS scores, advanced age, and unique biomarkers. Additionally, we discuss procedural innovations, such as tailored device use and alternative access strategies to address anatomical and clinical complexities. While substantial progress has been made, challenges remain in refining patient selection criteria, mitigating procedural risks, and ensuring equitable access to thrombectomy. Future directions include taking full advantage of advanced imaging modalities, incorporating biomarkers for personalized care, and optimizing thrombectomy devices to support the use of thrombectomy in underrepresented populations. Precision thrombectomy has the potential to be adapted to a broader spectrum of patients, improving outcomes and ultimately reducing the global burden of stroke.

KEYWORDS

stroke thrombectomy, personalized medicine, large vessel occlusion, endovascular treatment, medium vessel occlusion

1 Introduction

Stroke is a leading cause of permanent disability and death worldwide, representing a global health concern (1). Accounting for approximately 87% of all strokes (2), acute ischemic stroke (AIS) occurs secondarily to thrombosis, embolism, or systemic hypoperfusion, of which approximately 40% are large vessel occlusions (LVOs) (3). Mechanical thrombectomy has emerged as the gold standard for AIS treatment, restoring cerebral perfusion and improving functional outcomes through complete recanalization of occluded intracranial arteries. While landmark studies validated the safety and efficacy of thrombectomy in acute ischemic stroke, these studies selected patients according to stringent inclusion criteria, such as early presentation and small core infarcts (4–8). However, many patients affected by stroke deviate from this phenotype, presenting new challenges and opportunities for personalized approaches.

Advances in neurovascular imaging and iterative thrombectomy device development have improved patient selection and procedural safety. The evolution of stroke thrombectomy is shifting towards precision medicine, optimizing treatment strategies to patient- and lesion-specific factors, including vessel occlusion type, infarct size, collateralization status, patient age, comorbidities, and individual biomarkers. This review explores this paradigm with the aim to guide clinicians and researchers in advancing personalized approaches to thrombectomy to expand its eligibility to traditionally excluded patient groups.

2 Personalization by occluded vessel type

2.1 Large vessel occlusion

Personalized stroke thrombectomy in large vessel occlusion (LVO) focuses on tailoring treatment strategies based on patient-specific and stroke-specific factors to maximize outcomes, with initial trials focusing on patients in early time windows (≤ 6 – ≤ 12 h from last known well, LKW) (Table 1). The MR CLEAN trial first demonstrated the superiority of thrombectomy using second-generation stent retrievers, such as Solitaire[®] and Trevo[®], over standard medical management, including intravenous thrombolysis (IVT) (4). Patients treated with thrombectomy showed significantly higher rates of functional independence (mRS 0–2 at 90 days: 32.6% vs. 19.1%) and smaller infarct volumes. These findings were corroborated by

subsequent trials, which included LKW times of ≤ 6 h (SWIFT PRIME and EXTEND-IA) (5, 6), extending to ≤ 8 h (REVASCAT) (7), and finally ≤ 12 h (ESCAPE) (8), collectively leading to the HERMES meta-analysis (9). Time from LKW status was 6 h in SWIFT. This analysis confirmed that thrombectomy nearly doubled the odds of achieving good functional outcomes (adjusted OR 2.49), emphasizing the importance of rapid intervention in a specific subgroup of patients. Following these results, thrombectomy was recommended as the preferred treatment for LVOs in the anterior circulation. The American Heart Association/American Stroke Association (AHA/ASA) provided a Class IA recommendation for patient inclusion criteria for thrombectomy, including baseline functional status (mRS 0–1), high ASPECTS (≥ 6), and NIHSS ≥ 6 for strokes in the anterior circulation (internal carotid artery and M1 segment of the middle cerebral artery) who present within 6 h of onset (10).

Personalized thrombectomy in the late time windows (6–24 h from LKW) represents a paradigm shift from time-based to tissue-based patient selection, driven by advanced imaging techniques (Table 2). Landmark trials such as DAWN and DEFUSE-3 demonstrated the efficacy of thrombectomy in patients with anterior circulation LVOs who had a small ischemic core but significant salvageable penumbra. The DAWN trial, which included patients up to 24 h post-stroke onset, showed a nearly fourfold increase in functional independence at 90 days (mRS 0–2 in 49% vs. 13% of the control group) (11), while DEFUSE-3 reported similar benefits for patients treated within 6–16 h (12). The results were transformative, with a number needed to treat (NNT) of just 3 to achieve improved

TABLE 1 Characteristics of RCTs investigating mechanical thrombectomy in large vessel occlusion in the early window.

Trial	MR CLEAN (4)	SWIFT PRIME (5)	EXTEND-IA (6)	REVASCAT (7)	ESCAPE (8)
Study characteristics					
Publication year	2015	2015	2015	2015	2015
Enrolment years	2010–2014	2012–2014	2012–2014	2012–2014	2013–2014
Country	Netherlands	Multi-national	Australia, New Zealand	Spain	Multi-national
Sample	500	196	70	206	315
Inclusion criteria					
Age	≥ 18	18–85, later changed to 18–80	≥ 18	18–80, later changed to 81–85 if ASPECTS >8	≥ 18
Time	0–6 h	0–6	0–6	0–8	0–12
Vessels	ICA, M1, M2, A1, A2	ICA, M1	ICA, M1, M2	ICA, M1	ICA, M1, M2
ASPECTS/Lesion volume	0–10	ASPECTS ≥ 6 ; volume <50 ml with mismatch	Volume <70 ml with mismatch	ASPECTS ≥ 7 (≥ 8 if age 81–85)	ASPECTS ≥ 6
NIHSS	≥ 2	8–29	–	≥ 6	≥ 5
Baseline mRS	–	≤ 1	≤ 1	≤ 1	–
TPA eligibility	Not required	Required	Required	Not required	Not required
Outcomes					
mTICI 2b, 3	75.4%	88.0%	86.2%	66.0%	72.4%
mRS 0–2	32.6% vs. 19.1%	60.2% vs. 35.5%	71% vs. 40%	43.7% vs. 28.2%	53% vs. 29.3%
Mortality	21% vs. 22%	12% vs. 9%	9% vs. 20%	18% vs. 15%	10% vs. 19%
sICH	7.7% vs. 6.4%	0% vs. 3%	0% vs. 6%	1.9% vs. 1.9%	3.6% vs. 2.7%
NNT	7.1	4	3.2	6.5	4.2

A1, A1 segment of the anterior cerebral artery (ACA); A2, A2 segment of the ACA; ASPECTS, Alberta Stroke Program Early CT Score; ICA, internal carotid artery; M1, M1 segment of the middle cerebral artery (MCA); M2, M2 segment of the MCA; mRS, modified Rankin score; mTICI, modified thrombolysis in cerebral infarction score; NIHSS, National Institutes of Stroke Scale; NNT, number needed to treat; sICH, symptomatic intracranial bleeding; TPA, tissue plasminogen activator.

TABLE 2 Characteristics of RCTs investigating mechanical thrombectomy in large vessel occlusion in the late window.

Trial	DAWN (11)	DEFUSE-3 (12)	MR CLEAN-LATE (17)
Study characteristics			
Publication year	2017	2018	2023
Enrolment years	2014–2017	2016–2017	2018–2022
Sample	206	182	502
Inclusion criteria			
Age	≥18	18–90	≥18
Time	6–24 h	6–16 h	6–24 h
Vessels	ICA, M1	ICA, M1	ICA, M1, proximal M2
ASPECTS	≥10	≥6	–
Lesion/Infarct volume	<21 ml for age ≥80 and NIHSS ≥10 <31 ml for age <80 and NIHSS ≥10 31–51 ml for age <80 and NIHSS ≥20	<70 ml, mismatch penumbra/core ratio ≥1.8 and absolute penumbra vol ≥15 ml	–
NIHSS	≥10 and ≥20	≥6	≥2
Baseline mRS	≤1	≤2	≤2
TPA eligibility	Ineligible for, or failed, IV tPA	Not required	Not required
Outcomes			
mTICI 2b or 3	72% (90%)	76%	73%
mRS 0–2	49% vs. 13%	45% vs. 17%	39% vs. 34%
Mortality	19% vs. 18%	14% vs. 26%	24% vs. 30%
sICH	6% vs. 3%	7% vs. 4%	7% vs. 2%
NNT	2.8	3.6	–

ASPECTS, Alberta Stroke Program Early CT Score; ICA, internal carotid artery; M1, M1 segment of the middle cerebral artery (MCA); M2, M2 segment of the MCA; mRS, modified Rankin score; mTICI, modified thrombolysis in cerebral infarction score; NIHSS, National Institutes of Stroke Scale; NNT, number needed to treat; sICH, symptomatic intracranial bleeding; TPA, tissue plasminogen activator.

functional outcomes. These findings led to revised guidelines supporting thrombectomy for selected late-window patients, emphasizing the importance of advanced imaging, such as CT perfusion or MRI, to identify ischemic core-penumbra mismatch. Recent meta-analyses, including the Analysis of Pooled Data from Randomized Studies of Thrombectomy More Than 6 Hours After Last Known Well (AURORA) collaboration, further solidified the benefit of thrombectomy across diverse subgroups, including older patients (age ≥80 years) and those with wake-up strokes (13). An illustrative case of an 89-year-old male presenting with National Institutes of Stroke Scale (NIHSS) score of 16 due to a left distal M1 occlusion, successfully recanalized using aspiration thrombectomy, is shown in Figure 1.

Collateralization status is the underlying phenomenon for this tissue-based approach, as it influences infarct progression and treatment outcomes. Collateral vessels provide alternative pathways for blood flow, sustaining ischemic penumbra and delaying core infarct expansion (14). Studies have shown that

patients with robust collaterals, termed “slow progressors,” lose neurons at significantly lower rates compared to those with poor collaterals, or “fast progressors,” who experience rapid infarct growth (15, 16). The MR CLEAN-LATE trial demonstrated that thrombectomy for anterior circulation ischemic stroke is effective in the late window (6–24 h from LKW) when patients are selected based on collateral flow assessment via CTA. At 90 days, thrombectomy was associated with significantly better functional outcomes compared to best medical management alone, with a shift towards lower modified Rankin Scale (mRS) scores [adjusted common odds ratio (OR) 1.67; 95% CI 1.20–2.32] (17). These benefits persisted at 2 years, with the thrombectomy group maintaining improved functional outcomes (adjusted OR 1.41; 95% CI 1.00–1.99) (18). Importantly, all patients included had confirmed collateral flow, emphasizing its pivotal role in identifying viable ischemic penumbra and guiding late-window thrombectomy. While symptomatic intracranial hemorrhage (sICH) was more frequent in the thrombectomy group, all-cause mortality was not significantly different, and thrombectomy proved safe and effective for improving long-term outcomes.

2.2 Medium vessel occlusions

Medium vessel occlusion refers to distal intracranial arterial occlusions, typically 0.75–2 mm in diameter (19). These occlusions occur in M2, M3, or M4 branches of the MCA, A2 or A3 branches of the ACA, and P2 or P3 branches of the PCA (20). Additionally, some authors consider occlusions of the A1 and P1 segments as MeVOs (21). Occlusions of the M2 segment of the MCA may also be considered an LVO or MeVO depending on the dominance of the M2 branch, with dominant M2 branch occlusions categorized as LVOs and co-dominant or non-dominant M2 branch occlusions categorized as MeVOs (22). MeVOs, arising as either primary occlusions or secondarily to previous angiographic or thrombectomy procedures, are responsible for 25%–40% of all acute ischemic strokes (19, 23). Given the high risk of long-term disability associated with MeVOs in up to 50% of patients (24), mechanical thrombectomy should be offered to appropriately selected patients. However, several challenges to the use of mechanical thrombectomy for these lesions exist. By categorization, MeVOs represent a heterogeneous group of conditions as characterized by the variability in occlusion sites and neurological deficits. These occluded vessels are also smaller, with thinner and more fragile walls, and are more prone to dissection, rupture, and vasospasm. The tortuosity of these distal vessels also impede navigation during mechanical thrombectomy (25–27). Despite the challenges, development of smaller mechanical thrombectomy devices, along with the effectiveness of intra-arterial fibrinolysis in just a third of these cases (19, 28), and reported efficacy of thrombectomy for MeVOs in retrospective studies (29), thrombectomy represents a valuable treatment modality for patients with MeVOs. As such, the 2019 AHA/ASA and Society for Neuro-interventional Surgery guidelines recommend

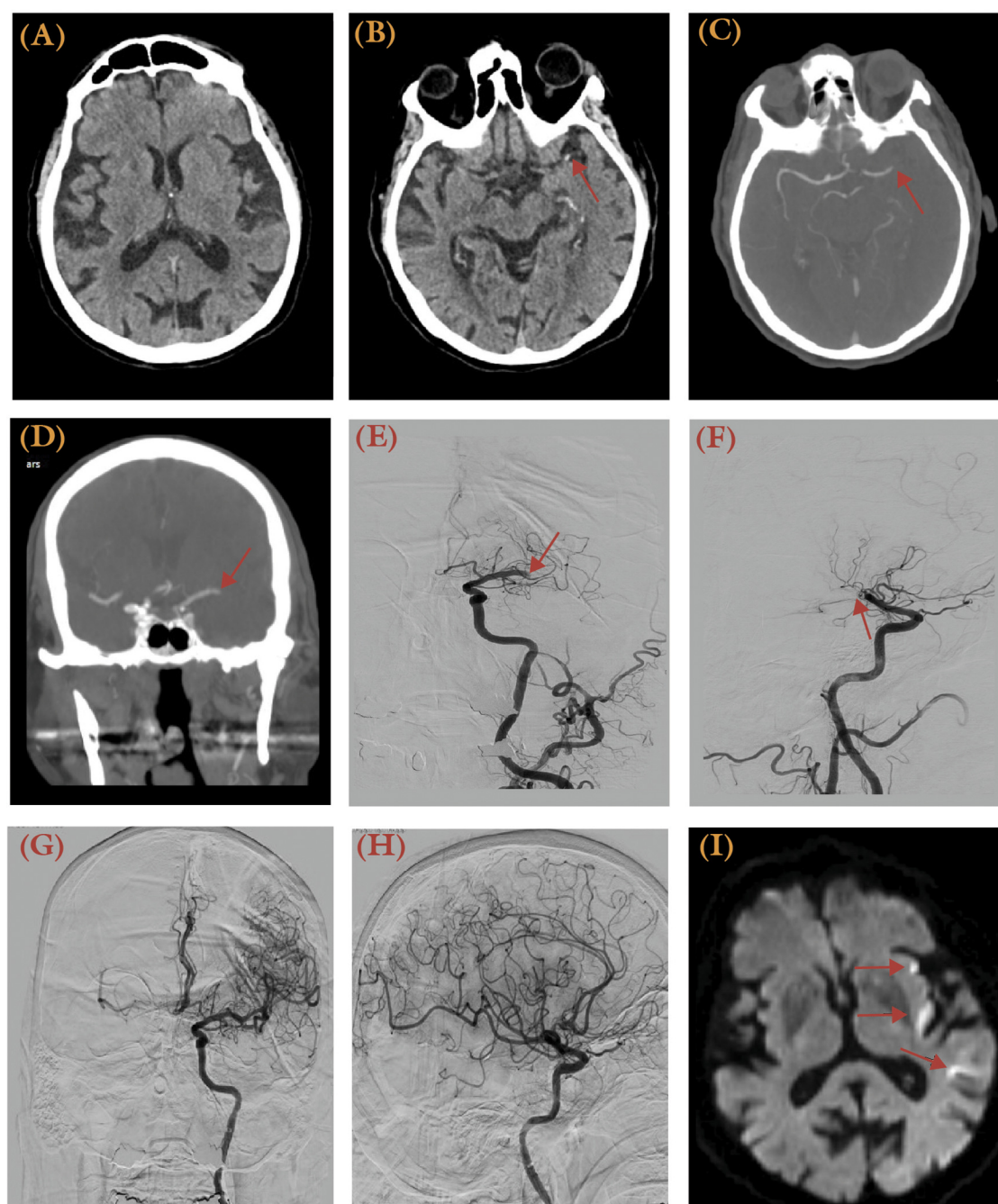


FIGURE 1

An 89-year-old male with a history of paroxysmal atrial fibrillation treated with 15 mg daily rivaroxaban, hypertension, and hyperlipidemia presented with right-sided weakness and aphasia, with the last known well 12 h prior. He had an NIH Stroke Scale (NIHSS) score of 16 and an ASPECTS score of 10. (A) No hypodense lesion was apparent from initial CT at presentation. (B) MCA dot sign was visible (arrow). (C,D) A computed tomography angiography (CTA) revealed a left distal M1 occlusion (arrow) with no distal perfusion. (E,F) Anteroposterior (E) and lateral (F) views of the digital subtraction angiography (DSA) illustrates the occlusion site at the left distal M1 (Arrow). (G,H) Post-thrombectomy, a TICI 3 recanalization was achieved using the ADAPT technique in anteroposterior (G) and lateral (H) views of the angiogram. (I) A post-thrombectomy MRI diffusion-weighted imaging (DWI) series revealed multiple punctate areas of restricted diffusion in the left insular and temporal regions, representing acute infarcts.

thrombectomy for patients with M2 or M3 occlusions (class IIB) and M2 occlusions, respectively (10, 30).

The safety and efficacy of EVT for MeVOs in the MCA circulation has been documented. A single center study of 62%

primary and 33% secondary heterogenous MeVO lesions reported significant reperfusion (mTICI 2b-3 in 83%) with observed 30% 90-day functional independence and 20% mortality rates (31). A patient level meta-analysis of M2 MCA

occlusions from data in the HERMES Collaboration demonstrated superior outcomes in thrombectomy-treated patients (mRS 0–2 in 58.2%) compared to controls (39.7%), with maximal benefit in proximal and dominant M2 occlusions (32). A case of a 32-year-old female with NIHSS 6 due to a left M2 occlusion, treated successfully with the ADAPT technique achieving TICI 2C recanalization, is shown in Figure 2. For non-M2 MCA MeVOs, a recent meta-analysis showed limited benefit for thrombectomy over control in A2, M3, P2, P3, and P4 lesions, with good outcomes in 54.7% vs. 54.5% respectively, although not statistically significant (33).

In the posterior circulation, the TOPMOST trial demonstrated efficacy and safety of thrombectomy in P2 and P3 occlusions among IVT-ineligible patients with admission NIHSS ≥ 10 (34). Similarly, thrombectomy was shown to be feasible for A2 to A4 occlusions, primarily due to large artery atherosclerosis, with a low sICH rate of 2% (35). The PLATO study, which focused on posterior circulation MeVOs, observed greater NIHSS improvement and earlier clinical benefit in the thrombectomy group, despite no significant shift in 90-day mRS. However, this group also experienced higher rates of sICH and mortality,

emphasizing the need to balance functional benefit with procedural risk. Stratified analysis suggested thrombectomy benefit among patients with baseline NIHSS >6 and no increase in adverse events. The limited value of mRS and NIHSS scores in posterior circulation strokes has also been reported, necessitating holistic assessments in functional status following treatment of these MeVOs (36).

Notably, two recent RCTs provide further clarity. The ESCAPE-MeVO trial, enrolling 530 patients with acute MeVOs (predominantly MCA branch occlusions), found no significant difference in 90-day functional independence (mRS 0–1 in 41.6% EVT vs. 43.1% control, $p=0.61$) and reported higher mortality and sICH rates in the EVT group (37). Likewise, the DISTAL trial, evaluating EVT in distal ACA, MCA, and PCA occlusions, showed no significant improvement in mRS at 90 days with EVT over best medical therapy (common OR 0.9, $p=0.50$), with comparable mortality and sICH rates between groups (38). As future trials continue to refine treatment indications, especially for patients with mild symptoms or distal occlusions, personalization will remain central to expanding the therapeutic frontier for MeVOs.

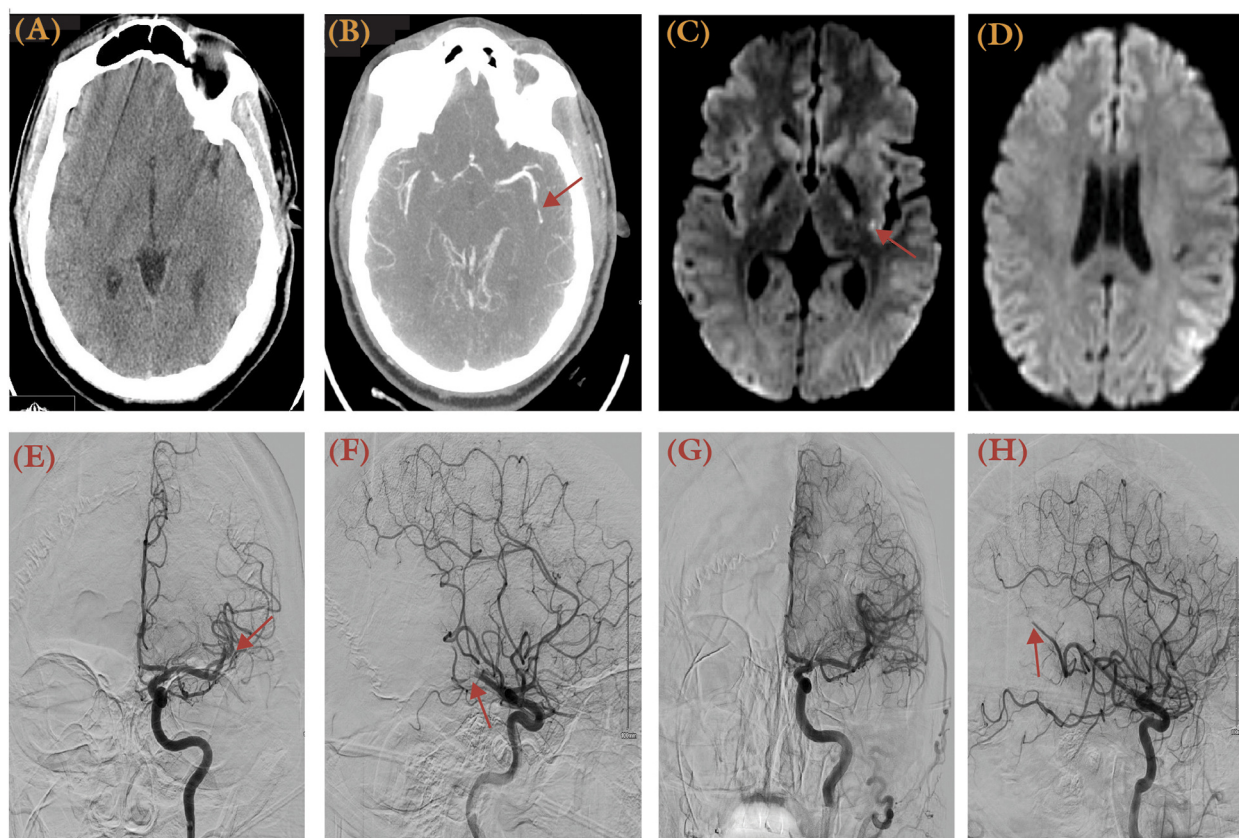


FIGURE 2

A 32-year-old obese male with a history of obesity and heart failure presented with aphasia, dysarthria, and right arm numbness, having been last known normal 1 h prior to presentation. His NIHSS score was 6. (A) Non-contrast brain CT showed an ASPECTS score of 10. (B) Brain CTA revealed an occluded left M2 vessel (arrow). (C,D) MRI DWI series revealed focal restrictions at left insular and parietal regions. Anterior-posterior (E) and lateral views (F) of DSA (left ICA injection) illustrated the site of occlusion as the M2 segment (arrow). (G,H) DSA after thrombectomy using the ADAPT technique demonstrated a TICI 2C recanalization. Distal occlusion at M3 segment was not pursued further.

2.3 Basilar artery occlusions

Basilar artery occlusions are associated with high mortality rates despite treatment (39). Patients with these strokes have a varied presentation, with clinical symptoms including cranial nerve defects, dysarthria, dysphagia, gait ataxia, and hemiplegia that progresses to locked-in syndrome (39). These signs and symptoms are typically not accurately captured with the NIHSS score, requiring more appropriate endpoints for assessing patient functional status. To mitigate this, scoring systems like the POST-NIHSS score have been proposed. Imaging-based markers, like the pons-midbrain index, have also been explored, though they are not validated clinical severity scores (40, 41). Factors associated with favorable outcomes post-thrombectomy for BAOs include a low pretreatment NIHSS score, duration of symptom (<6 h), good collateralization status, the absence of an early pontine infarct, and thrombectomy technique. Proper identification of appropriate patients with BAO for thrombectomy is strongly supported by current guidelines (36, 42).

Data from the BASILAR registry, including 1,254 patients, was the first to support the efficacy of thrombectomy within 24 h for BAO, demonstrating significantly better treatment safety and efficacy (43). Two subsequent trials, the BASICS (BASilar artery International Cooperation Study) and the BEST (Endovascular Treatment vs. Standard Medical Treatment for Vertebrobasilar

Artery Occlusion) investigated thrombectomy for BAOs in patients presenting within 6 h and 8 h of stroke, respectively (Table 3) (44, 45). BASICS randomized patients into the thrombectomy and best medical therapy arms, with slightly better functional outcomes seen in the thrombectomy group (44.2% vs. 37.7%) and lower mortality rates (38.4% vs. 43.2%), but with higher sICH rates (4.5% vs. 0.7%). Patients who demonstrated higher rates of actual or potential favorable 90-day mRS scores (mRS 0–3) after thrombectomy were those who presented with moderate (NIHSS 10–19) and severe (NIHSS ≥ 20) neurological deficit. A case of a 78-year-old female with NIHSS score of 28 due to an upper basilar occlusion, successfully treated with the SOLUMBRA technique achieving TICI 3 recanalization, is presented in Figure 3. Similarly, BEST compared thrombectomy to best medical therapy but was terminated early due to poor recruitment and high treatment crossover rates (22% from best medical therapy to thrombectomy). 47% of patients treated with thrombectomy reported higher favorable 90-day mRS scores (mRS 0–3), compared to 24% of control. Overall, BASICS and BEST did not demonstrate superiority of thrombectomy over best medical treatment, which were addressed by subsequent trials.

ATTENTION (Endovascular Treatment for Acute Basilar Artery Occlusion) is a Chinese trial that randomized thrombectomy and medical treatment for patients with BAO presenting within 12 h of stroke, with baseline NIHSS score > 10,

TABLE 3 Characteristics of RCTs investigating mechanical thrombectomy in basilar artery occlusion.

Trial	BASICS (42)	BEST (43)	ATTENTION (44)	BAOCHE (45)
Study characteristics				
Publication year	2021	2020	2022	2022
Enrolment period	2011–2019	2015–2017	2021–2022	2016–2022
Country	Multi-national (7 countries)	China	China	China
Patients	300	131	340	217
Inclusion criteria				
Age	Initially 18–85, later expanded to ≥ 18	≥ 18	≥ 18	18–80
Time	0–6 h	0–8 h	0–12 h	6–24
Vessels	BA (proximal, middle, and distal)	VA (V4), BA	VA (V4), BA (proximal, middle, and distal)	BA (Proximal, middle, and distal)
pc-ASPECTS	≥ 10 , later expanded to <10	–	≥ 6 for age <80 ≥ 8 for age ≥ 80	≥ 10 , later expanded to ≥ 6 and pons-midbrain index ≤ 2
NIHSS	Initially ≥ 10 but removed after 91 patients enrolled	None	≥ 10	≥ 10 initially, changed to ≥ 6 after 84 patients enrolled
Baseline mRS	0–2	0–2	0–2 (age ≤ 80); 0 (mRS >80)	0–1
IV tPA eligibility	Not required	Not required	Not required	Not required
Outcomes				
mTICI 2b, 3	72%	71%	93.3%	88.1%
mRS 0–2	35% vs. 30%	33% vs. 28%	33.2% vs. 10.5%	39% vs. 14%
mRS 0–3	44% vs. 37.7%	47% vs. 24%	46 vs. 22.8%	46.4% vs. 24.3%
Mortality	38.3% vs. 43.2%	33% vs. 38%	36.7% vs. 55.3%	30.9% vs. 42.1%
sICH	3.9% vs. 0.7%	8% vs. 0%	5.3% vs. 0%	5.9% vs. 1.1%
Procedure-related complications	3%	–	14%	11%

ASPECTS, Alberta Stroke Program Early CT Score; BA, basilar artery; mRS, modified Rankin score; mTICI, modified thrombolysis in cerebral infarction score; NIHSS, National Institutes of Stroke Scale; NNT, number needed to treat; sICH, symptomatic intracranial bleeding; TPA, tissue plasminogen activator; V4, V4 segment of the vertebral artery.

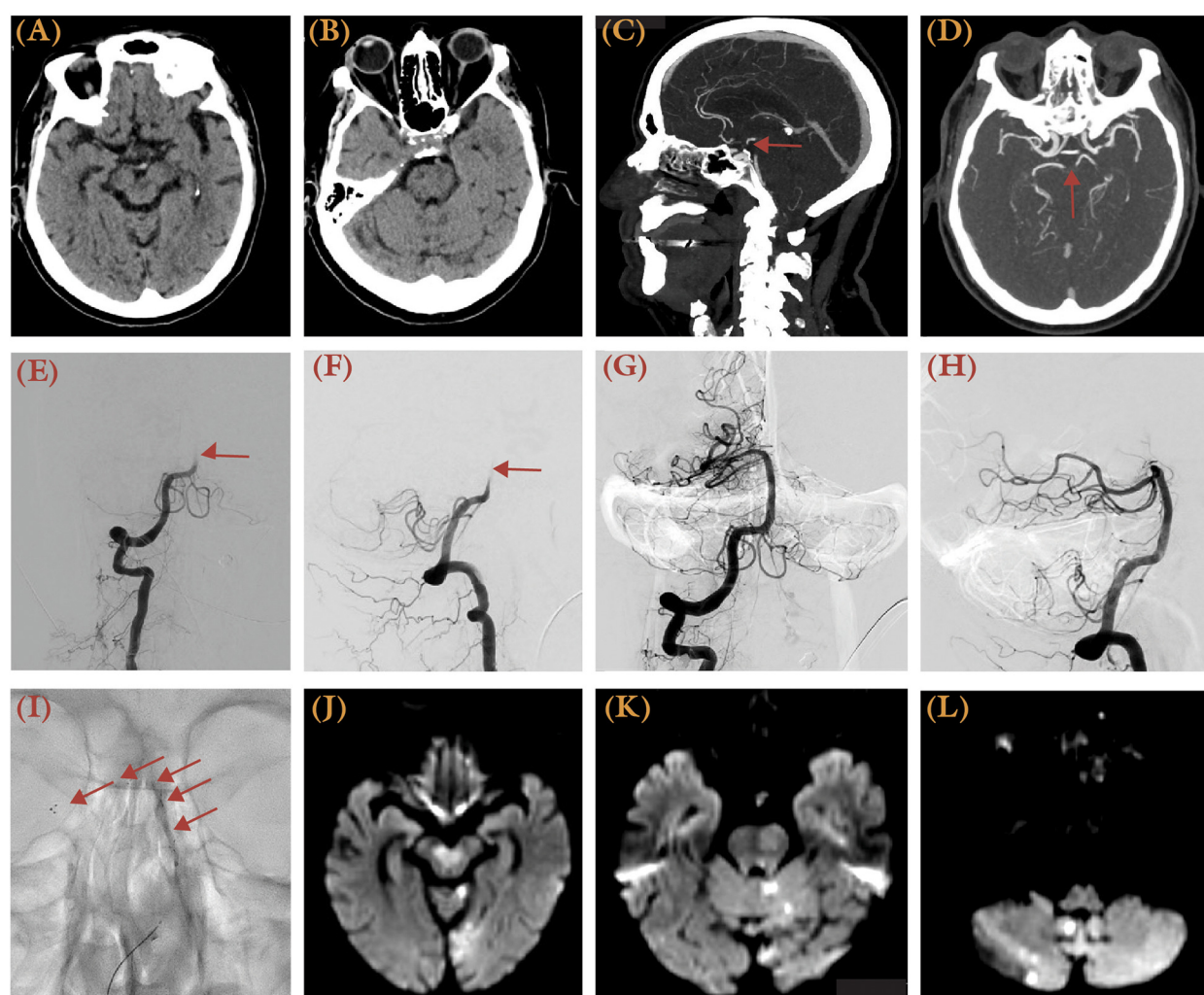


FIGURE 3

This 78-year-old female with a history of coronary heart disease, atrial fibrillation, hypertension, and hyperlipidemia experienced acute dizziness, gaze palsy, aphasia, and right-sided weakness, presenting with an NIHSS score of 28 and last known well 3.5 h prior to presentation. (A,B) Brain CT demonstrated possible hypodense over the bilateral midbrain and pons, with <50% involvement and the left cerebellum. Sagittal (C) and axial (D) views of brain CTA MIP images demonstrated an upper basilar occlusion (arrow). (E,F) Right vertebral artery injection DSA showed restriction of contrast flow at the vertebrobasilar junction (arrow) in both anterior-posterior (E) and lateral views (F) before mechanical thrombectomy. (G,H) A TICI 3 recanalization was achieved after two passes using the SOLUMBRA technique. (I) Unsubtracted x-ray illustrates the stent retriever, as delineated by arrows. (J–L) A follow-up MRI showed areas of restricted diffusion over the cerebellum, left occipital lobe, and brainstem.

and age-adjusted pc-ASPECTS (>6 points for patients aged <80; > 8 points for those aged > 80) (46). Additionally, 40% of patients in the thrombectomy group underwent angioplasty and stenting. A significantly higher proportion of patients in the thrombectomy demonstrated good functional outcome (mRS 0–3) compared to control (46% vs. 23%). The Basilar Artery Chinese Endovascular Trial (BAOCHE) is another RCT that investigated the safety and efficacy of thrombectomy in the late time window (6–24 h), with NIHSS ≥ 6 , and pc-ASPECTS <6, and pons-midbrain index ≤ 2 (47). Like ATTENTION, additional angioplasty or stenting was performed in 55% of thrombectomy-treated patients. The thrombectomy group demonstrated higher rates of favorable functional outcomes (46.4% mRS 0–3 vs.

24.3%), lower mortality rates (31% vs. 42%), but with increased rates of sICH (6% vs. 1%). The etiology of BAO is also important given the Chinese population recruited in both studies, who are likely to have a significant intracranial atherosclerosis burden (26).

2.4 Tandem occlusions

Tandem occlusions are defined as concurrent intracranial LVOs and extracranial internal carotid artery (ICA) occlusion or severe stenosis. With an incidence of 10%–20% of anterior circulation LVO strokes (48), endovascular treatment of tandem

occlusions is clinically and technically challenging, requiring strict management of blood pressure, balancing thrombotic and bleeding risks, and treatment of both extracranial and intracranial lesions. Tandem occlusions are primarily caused by carotid artery atherosclerosis or, less commonly, acute dissection (49). The lack of efficacy of IVT in tandem occlusions has shifted treatment focus to thrombectomy to achieve sufficient recanalization in these patients (50).

Various thrombectomy strategies have been proposed for tandem occlusions, including antegrade approaches (treating the extracranial lesion first), retrograde approaches (treating the intracranial LVO first), balloon angioplasty, and conservative thrombectomy without extracranial intervention. Meta-analyses and registry data indicate no significant differences in outcomes between antegrade and retrograde approaches, though the retrograde method may offer faster recanalization (51, 52). Acute carotid stenting, while effective in restoring blood flow and reducing stroke recurrence, requires immediate initiation of antiplatelet therapy, increasing the risk of hemorrhagic complications. Despite these risks, stenting has demonstrated improved odds of functional independence and successful reperfusion in multiple studies, and patient selection should consider factors like underlying etiology, core infarct volume, and patient anatomy (53). A case of a 75-year-old male with NIHSS score of 14 and severe stenosis within left ICA bifurcation, treated with angioplasty and aspiration achieving TIC1 3 reperfusion, is illustrated in Figure 4.

Emerging evidence supports the overall benefit of thrombectomy for tandem occlusions. A meta-analysis of 1,373 patients demonstrated improved functional outcomes at 90 days (OR 1.43) (54), while pooled data from the ETIS and TITAN registries highlighted better reperfusion and favorable outcomes with stenting, although with higher rates of non-symptomatic intracranial hemorrhage (55). However, clinical benefit was less pronounced in tandem occlusions secondary to dissection, emphasizing the need for personalized decision-making based on etiology. Ongoing randomized trials, such as DST-TANDEM, EASI-TOC, and TITAN, aim to provide more definitive guidance on optimal management strategies for tandem occlusions (56–58). These studies are expected to clarify the benefits of acute stenting and refine patient selection criteria.

3 Personalization by lesion core and infarct size

3.1 Large core infarcts

Large core infarcts are strokes identified through neuroimaging as those with extensive areas of brain tissue injury. The Alberta Stroke Program Early CT (ASPECTS) score characterizes stroke infarct burden, with earlier stroke trials focusing on small preprocedural infarct sizes, defined as ASPECTS 6–10 (9, 11). The stroke core volume on CT perfusion (CTP) or MRI also defines infarct burden, with the DAWN and DIFUSE-3 trials selecting patients with median core volumes of 7.6 and 9.4 ml,

respectively (11, 59). Infarct burden is associated with outcomes after thrombectomy, with a favorable likelihood significantly decreasing with each ASPECT score (60). Accordingly, upper limits of 50–70 ml core volume or ASPECTS of 6 were set in earlier trials to reflect this (28).

The RESCUE-Japan LIMIT was the first trial to randomize patients with large core infarcts for mechanical thrombectomy and best medical treatment (Table 4) (61). Selecting patients with ASPECTS 3–5, thrombectomy significantly improved functional independence, with 31% of patients achieving 90-day mRS 0–3 scores compared to 12.7% in the best medical treatment group. Similarly, the SELECT 2 trial defined large infarcts as ASPECTS 3–5 and stroke volume ≥ 50 ml, with 20.7% of thrombectomy-treated patients achieving an mRS score of 0–2 compared to 7% in the medical treatment group (62). The ANGEL-ASPECT trial also supported these findings, with large infarcts defined as ASPECTS 3–5 and ASPECTS 0–2 or ≥ 5 with infarct volumes 70–100 ml (63). This trial showed that 30% of thrombectomy patients achieved mRS 0–2 compared to 11.6% in the medical group. National stroke guidelines now support thrombectomy for select patients with ASPECTS 3–5, recognizing the potential for improved outcomes and cost-effectiveness in this population (64). While these trials confirmed the benefits of thrombectomy, the challenges in treating large infarct strokes with thrombectomy are well recognized. These patients are at increased risk for hemorrhagic transformation, cerebral edema, prolonged hospitalization, and mortality (53). In contrast, thrombectomy may reduce the need for decompressive hemicraniectomy by preventing malignant cerebral edema, potentially improving both survival and quality of life (65).

The Thrombectomy in Stroke with Extended Lesion and Extended Time Window (TENSION) trial, randomizing patients with ASPECTS 3–5 within 12 h of onset, was halted early due to significantly better mRS scores and lower mortality rates in the thrombectomy group, with comparable sICH rates between the two treatment arms (66). Recently, the TESLA and LASTE trials have demonstrated a holistic approach in thrombectomy that considers salvageable penumbra, infarct location, and patient-specific factors, in addition to lesion volume. TESLA evaluated thrombectomy in patients with ASPECTS 2–5 within 24 h of stroke onset (67). While the 90-day mean-utility mRS score favored thrombectomy (2.93 vs. 2.27 in the control group), the adjusted difference did not meet statistical significance. Mortality rates were similar between groups (35.3% vs. 33.35%), but higher rates of sICH (4.0% vs. 1.3%) and other hemorrhagic complications were noted in the thrombectomy group. LASTE demonstrated that thrombectomy benefits patients with large infarcts regardless of size, including patients with ASPECTS ≤ 5 and no minimum ASPECTS limit (68). Among patients with a median ASPECTS of 2, thrombectomy improved 90-day functional outcomes (OR 1.63; 95% CI 1.29–2.06; NNT = 4) and reduced all-cause mortality (36.1% vs. 55.5%). Benefits extended to quality of life at 90 and 180 days and reduced the rate of decompressive craniectomy. Like TESTLA, thrombectomy was associated with higher rates of sICH (9.6% vs. 5.7%) and procedural complications (6.9%).

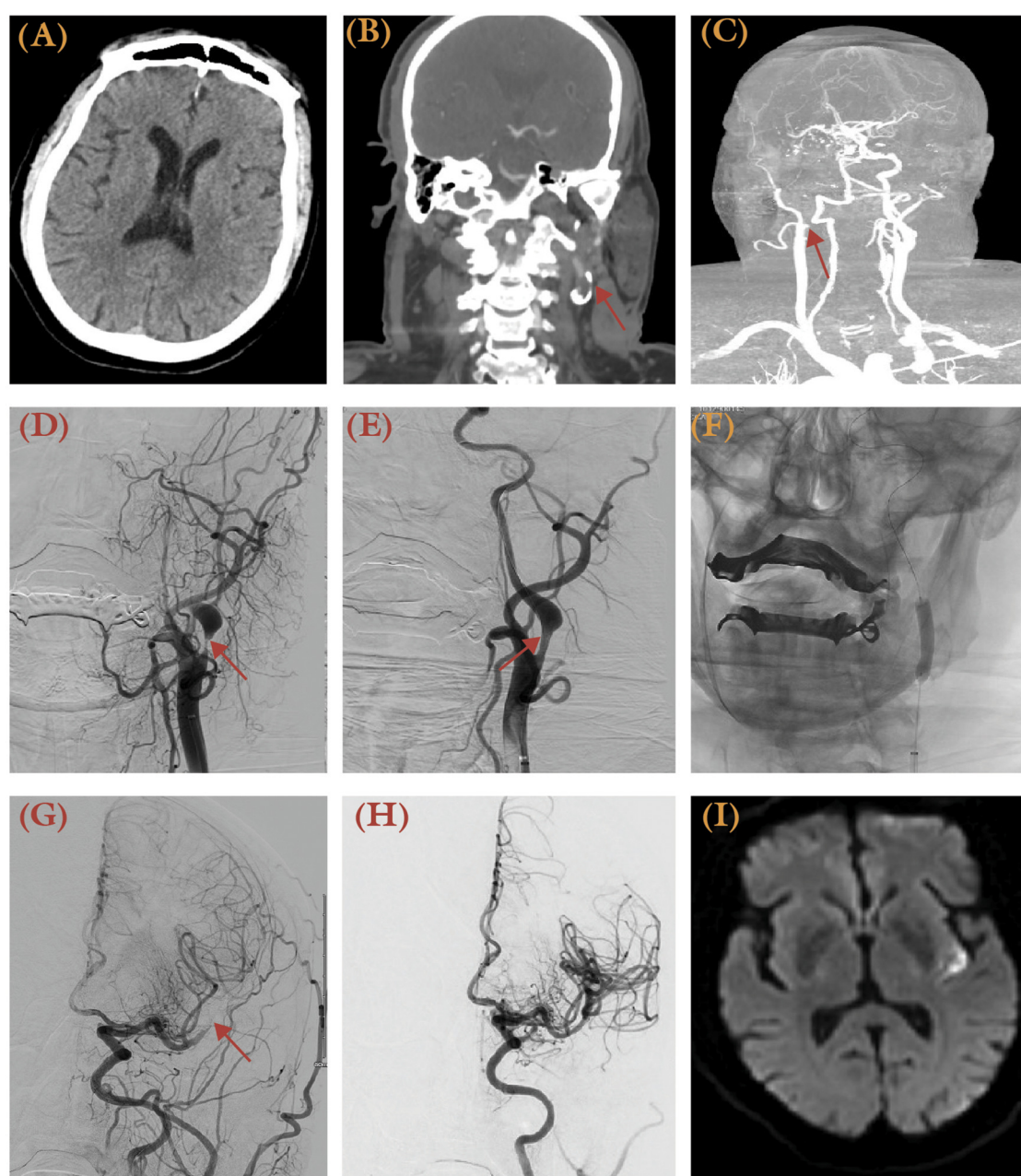


FIGURE 4

A 75-year-old male with a history of hypertension presented with altered mental status and aphasia, with NIHSS 14 and last known well 2 h prior to presentation. (A) Brain CT revealed no apparent hypodensity of the brain parenchymal. (B,C) CTA revealed atherosclerotic plaque resulting in severe stenosis within the left carotid bifurcation. AP view of digital subtraction angiography (DSA) revealed severe stenosis at the origin of the left ICA following a left CCA injection before (D) and after angioplasty (E), with improved caliber of the ICA (arrow). (F) AP view of unsubtracted x-ray illustrates the balloon used for angioplasty (arrows). (G) AP view angiography of the left CCA injection demonstrated an occlusion over the left M2 (arrow). (H) Following mechanical thrombectomy with aspiration technique, a TIC1 3 reperfusion was achieved. (I) Despite successful recanalization, post-procedural MRI confirmed a small left posterior insular cortex infarct.

With some patients with large infarcts achieving favorable recovery, while others with smaller infarcts may have poor prognoses, careful patient selection based on holistic considerations rather than infarct size alone is pertinent to guide therapy (69). The ratio of salvageable penumbra to infarct core size is critical in determining patient eligibility

and potential benefit. Infarct location also plays a role, with the involvement of eloquent brain tissue potentially impacting outcomes significantly (70). Younger patients often also show better recovery potential, making them more likely to benefit from thrombectomy even with larger infarct burdens.

TABLE 4 Characteristics of RCTs investigating mechanical thrombectomy in large core infarcts.

Trial	RESCUE-JAPAN LIMIT (59)	SELECT 2 (60)	ANGEL-ASPECT (61)	TENSION (64)	TESLA (65)	LASTE (66)
Study characteristics						
Publication year	2022	2023	2023	2023	2024	2024
Enrolment year	2018–2020	2019–2022	2020–2022	2018–2023	2019–2022	2019–2022
Country	Japan	Multi-national	China	Europe and Canada	USA	France and Spain
Patients	203	352	456	253	300	333
Inclusion criteria						
Age	≥18	18–85	18–80	≥18	18–85	≥18
Time	0–24 h	0–24 h	0–24 h	0–11 h	0–24 h	0–6.5 h
Vessels	ICA, M1	ICA, M1	ICA, M1	ICA, M1	ICA, M1	ICA, M1
ASPECTS/ Lesion volume	ASPECTS 3–5	ASPECTS 3–5; infarct core volume ≥50 ml	ASPECTS 3–5; ASPECTS <3 with infarct core 70–100 ml; ASPECTS >5 with volume 70–100 ml	3–5	2–5	0–5 (4–5 for patients ≥80 years)
NIHSS	≥6	≥6	6–30	<26	≥6	≥6
Baseline mRS	0–1	0–1	0–1	0–2	0–1	0–1
TPA eligibility	Not required	Not required	Not required	Not required	Not required	Not required
Outcomes						
mRS 0–2	14% vs. 7.8%	20% vs. 7%	30% vs. 12%	17% vs. 2%	14.6% vs. 8.9%	13.3% vs. 4.9
mRS 0–3	31% vs. 12.7%	38% vs. 19%	47% vs. 33%	31% vs. 13%	29.8% vs. 19.9%	–
Mortality	18% vs. 23.5%	38% vs. 42%	22% vs. 20%	40% vs. 51%	35% vs. 33%	36.1% vs. 55.5%
sICH	9% vs. 4.9%	0.6% vs. 1.1%	6.1% vs. 2.6%	5% vs. 5%	4% vs 1.3%	9.6% vs. 5.7%

ASPECTS, Alberta Stroke Program Early CT Score; ICA, internal carotid artery; M1, M1 segment of the middle cerebral artery (MCA); mRS, modified Rankin score; mTICI, modified thrombolysis in cerebral infarction score; NIHSS, National Institutes of Stroke Scale; NNT, number needed to treat; sICH, symptomatic intracranial bleeding; TPA, tissue plasminogen activator.

3.2 Small core infarcts

Small core infarcts, characterized by minimal established ischemic damage, offer a larger penumbral region for potential salvage, making them prime candidates for thrombectomy (71). The ESCAPE, REVASCAT, SWIFT PRIME, and EXTEND-IA trials, combined in the HERMES meta-analysis, demonstrated the significant benefit of thrombectomy in patients with small core infarcts, as identified by high ASPECTS scores on non-contrast CT or favorable perfusion imaging using software like RAPID (5–9). Across these studies, thrombectomy significantly improved functional outcomes, with a strikingly low NNT of 2.6 for one-point mRS improvement and 5 for achieving functional independence (mRS 0–2). In extended time windows (>6 h), trials like DAWN and DEFUSE 3 reaffirmed the efficacy of thrombectomy in late presenters with small core infarcts and favorable penumbral patterns, selected through advanced neuroimaging (11, 12). Remarkably, these trials showed no significant increase in sICH between thrombectomy and control groups, reinforcing the safety of thrombectomy when patient selection is rigorously guided by imaging. Retrospective studies have also shown that simplified imaging strategies, such as multiphase CTA, can reliably identify patients with good collaterals and small infarct volumes eligible for thrombectomy beyond 6 h, with comparable outcomes to more advanced imaging modalities (72, 73).

4 Personalization by patient-specific characteristics

4.1 Patients with low NIHSS scores

The National Institutes of Health Stroke Scale (NIHSS) is an 11-category tool designed to quantify neurological deficits in stroke patients (74). While a NIHSS score ≥6 is typically associated with more severe strokes and is often an inclusion criterion for mechanical thrombectomy trials, patients with NIHSS scores 0–5 comprise a unique subset with distinct challenges and opportunities for treatment personalization (75). Low NIHSS scores are observed in 4%–8% of patients with minor neurological deficits despite harboring an LVO, with these patients making up 10% of all LVO strokes undergoing thrombectomy (76, 77). Importantly, ≥50% of these patients present beyond six hours from symptom onset and up to 35% of these patients may have unfavorable outcomes at 90 days, complicating treatment decisions (28). A key challenge in managing low NIHSS stroke patients is the risk of neurologic deterioration, defined as an NIHSS increase in ≥2 points (78). Proximal occlusions, larger thrombus burdens, and failure of collateral blood flow are predictors of this decline, which is associated with poor patient outcomes (79).

As low NIHSS patients have been historically excluded from most thrombectomy trials, evidence for mechanical thrombectomy in this group is limited to observational studies

and case series, with no RCTs definitively establishing efficacy. Retrospective analyses and meta-analyses have yielded mixed results. While some studies suggest thrombectomy may confer benefits in select low NIHSS patients, others report no significant differences compared to best medical therapy, with thrombectomy associated with higher risks of sICH in some cases (80–83). Personalization of treatment for low NIHSS patients involve appropriate patients, which may comprise higher-risk subgroups like those with proximal occlusions, larger perfusion deficits, or symptoms affecting motor or language function (28).

While some retrospective studies suggest that immediate thrombectomy may lead to better outcomes than rescue therapy following neurological deterioration in this subgroup of patients, the heterogeneity of low NIHSS stroke patients necessitates further research to define optimal selection criteria. Current guidelines recommend enrolling low NIHSS patients with LVO into RCTs (25), which ongoing trials like ENDOLOW (Endovascular Therapy for Low NIHSS Ischemic Strokes) and MOSTE (Minor Stroke Therapy Evaluation) aim to address the knowledge gaps in this area (84).

4.2 Older patients—octogenarians and nonagenarians

The role of patient age in stroke thrombectomy is a critical area for personalization, particularly in older populations such as octogenarians and nonagenarians. These groups often present unique challenges due to a higher prevalence of comorbidities, disabilities, and frailty, as well as lower brain reserve, which can influence both the risks and benefits of thrombectomy (85).

While most thrombectomy trials have not excluded older adults, the average age of participants included is 68 years, with only 15% over the age of 80 (9). Despite these limitations, growing data suggest that thrombectomy can be both technically successful and clinically beneficial for select older patients (86). Furthermore, studies have shown that older patients can achieve comparable success rates to younger patients, provided that procedural challenges such as increased vessel tortuosity, which can complicate arterial access, are effectively managed (87).

An important association with thrombectomy outcomes for older patients is their baseline functional status. Studies indicate that nonagenarians with a baseline mRS score ≤ 2 are more likely to achieve favorable outcomes after thrombectomy compared to those with higher mRS scores. For instance, premorbidly independent nonagenarians have demonstrated favorable outcomes in up to 39% of cases, with some studies reporting similar or even better results compared to octogenarians (85, 88). However, patients with significant pre-stroke disabilities (e.g., mRS > 2) often experience poorer outcomes, though exceptions may exist for otherwise healthy individuals with orthopedic or non-cognitive impairments (89). While overall outcomes for older patients are generally poorer compared to younger populations, thrombectomy has been shown to provide meaningful benefits (90). Nonagenarians who achieve successful

reperfusion demonstrate greater early neurological recovery, reduced mortality, and improved 90-day functional outcomes compared to those managed with thrombolytic therapy alone (89). However, the range of favorable outcomes is wide, with rates reported between 12% and 31% (91, 92).

The decision to pursue thrombectomy in older patients should be individualized, considering baseline functional independence, comorbidities, and patient-specific goals of care. Currently, the European Stroke Association guidelines recommend thrombectomy for octogenarians with appropriate imaging characteristics, although the case for nonagenarians is not specifically addressed (10, 93). This highlights the importance of a holistic approach to decision-making, including the evaluating imaging findings, the risk of procedural complications, and potential benefits in restoring independence.

4.3 Patient-specific biomarkers

The paradigm of personalization of stroke thrombectomy has increasingly shifted to understanding the predictive potential of patient-specific biochemical, inflammatory, and physiological markers in optimizing outcomes and minimizing risks. High blood glucose levels, markers of renal health (e.g., low serum uric acid to creatinine ratio), and high C-reactive protein have been found to be predictors of hemorrhagic transformation and poor outcomes post-thrombectomy (94–96). The TICI-ASPECTS-glucose (TAG) score, representing the combined effects of moderate to low recanalization, low ASPECTS, and hyperglycemia has been validated for predicting sICH post-thrombectomy, with a 50% increased sICH likelihood per-point increase in TAG score (97). Baseline glucose, in addition to increasing age, baseline NIHSS, onset to puncture > 6 h, sICH, and pneumonia were also found to be independent predictors of poor functional outcomes post-thrombectomy in a Chinese population (98). The role of systemic inflammation in stroke recovery and thrombectomy outcomes is also well documented. Inflammatory markers like the blood cell ratios and indices of systemic inflammation have been linked to functional outcomes, sICH, stroke-associated pneumonia, and mortality rates post-thrombectomy. For example, higher platelet-to-neutrophil (PNR) and lower platelet-to-lymphocyte (PLR) ratios were associated with favorable 3-month outcomes post-thrombectomy, with optimal cutoff values of PNR > 41 and PLR < 145 (99). Similarly, patients with a high systemic immune-inflammatory index (SII) were more likely to develop malignant cerebral edema, which accounted for 40.3% of the poorer prognosis among these individuals (100), while systemic inflammatory response index (SIRI) has demonstrated predictive value for stroke-associated pneumonia post-thrombectomy (101). Blood pressure variability (BPV) and other hemodynamic factors also impact thrombectomy outcomes. Increased pulse pressure variability (PPV), a parameter of BPV, in the first 24 h post-thrombectomy strongly predicts 90-day functional outcomes, with adjusted ORs exceeding 40 for the highest PPV ranges. Similarly, deviations from a linear systolic blood pressure course, such as failing to

maintain a gradual decline from 130 to 123 mmHg over 24 h, were associated with a 47% reduced likelihood of functional independence (102). High-resolution BP monitoring also suggests that successive variation in systolic BP over 5-minute intervals better predicts outcomes compared to static BP measures (AUC 0.74, $p = 0.031$) (103). The DETERMINE trial is exploring whether personalized BP management, defined as maintaining MAP within 10% of baseline, leads to better functional outcomes compared to the standard target range of 140–180 mmHg (104). Preliminary observational studies support this approach, as patients who spent more time outside their personalized autoregulatory BP limits showed worse 90-day outcomes, with odds of poor outcomes increasing by 1.84 for every 10% of time spent above the upper autoregulatory limit. Notably, hemorrhagic transformation was more frequent when mean arterial pressure exceeded these limits (16.0% vs. 10.9%, $p = 0.042$) (105).

Patient clinical, demographic, and biomarker data can be integrated into predictive algorithms to enhance personalized thrombectomy. Machine learning (ML) algorithms like random forest (RF), extreme gradient boosting (XGBoost), and neural networks have demonstrated improved stroke detection, thrombectomy candidacy, and prognosis. An RF model, using 35 clinical and demographic variables, was able to predict the presence of LVOs and appropriately thrombectomy candidates with AUCs of 0.91 and 0.93, respectively, outperforming conventional stroke scales (106). By integrating preoperative and postoperative variables, such as NIHSS scores, an XGBoost model was able to improve predictive ability for unfavorable 3-month outcomes, enabling timely therapeutic adjustments (107). Additionally, a Back propagation neural network was able to predict 3-month mRS post-thrombectomy using patient clinical data, achieving prediction accuracy, sensitivity, and specificity of 96.1%, 98.3%, and 87.5%, respectively (108). The utility of ML applications in personalized stroke thrombectomy has been confirmed by a recent meta-analysis, which pooled five studies in predicting favorable 90-day functional outcomes (mRS 0–2) (109). With overall AUC of 0.85, sensitivity of 0.80, specificity of 0.78, and diagnostic odds of 12.6, the efficacy of ML-based predictive value holds great potential for optimizing patient identification for thrombectomy.

4.4 Special patient populations

Special populations refer to patient groups who are often excluded from major clinical trials, including children, patients with hematologic abnormalities, active cancer, recent surgery, collagen vascular diseases, or pregnant women. Although case series have demonstrated the safety and feasibility of thrombectomy in many of these populations, these cases are rare, and more studies are necessary to strengthen the evidence base of precision thrombectomy care (28). Guidelines support personalized thrombectomy in neonates, infants, children, or adolescents, recommending standard radiation dose-minimization techniques for diagnostic and therapeutic procedures, and non-ionizing imaging for initial stroke evaluation in children under

10 years of age. A thrombocytopenic status should not preclude thrombectomy, although hematologic consultation and, in some cases, platelet transfusion may be required for extremely thrombocytopenic patients ($<20,000/\text{mm}^3$ platelet count). Female patients should be offered thrombectomy regardless of pregnancy status, with proper focus on appropriate patient selection, radiation safety protocols, and multidisciplinary team input where necessary. Personalized thrombectomy should be also offered in patients at high risk of diagnostic and therapeutic complications, including those with active endocarditis, collagen vascular disease, and recent surgical history. These patient groups will benefit from prioritization of non-invasive imaging where appropriate, careful procedural planning, and multidisciplinary input to mitigate risks of procedural planning or hemorrhagic transformation (110).

5 Personalization by devices and approaches

5.1 Personalization through thrombectomy devices

Mechanical thrombectomy restores cerebral blood flow in patients with acute ischemic stroke using stent retrievers, aspiration catheters, or a combination of these devices to remove the thrombus under fluoroscopic guidance. Device selection and procedural techniques have evolved significantly, driven by innovations in catheter systems, guidewires, and thrombectomy devices, which have enhanced safety, efficacy, and procedural speed.

Stent retrievers are among the earliest and most established devices for thrombectomy. These devices are introduced through a microcatheter, deployed at the site of occlusion, and used to capture and remove the thrombus by retracting it along the vessel wall. Stent retrievers can be personalized according to occlusion site and clot characteristics. More recent innovations of stent retrievers are optimized for distal vessel occlusions with vessel diameters 0.5–2 mm (111, 112). Stent retrievers are also preferred for soft, red-blood-cell-rich thrombi, demonstrating higher recanalization in these lesions (113). While stent retrievers are effective, challenges such as vessel tortuosity and clot stiffness can reduce their efficacy. Additionally, dragging the device across the endothelium may cause vascular damage (114), a limitation addressed by emerging techniques combining stent retrievers with aspiration catheters.

Aspiration catheters remove thrombi by creating negative pressure through a syringe or pump (115). This approach, typified by the ADAPT technique, is simpler than stent retriever thrombectomy and often results in less endothelial damage (116). Larger bore catheters, such as the Trac Star 088 and Millipede 088, have improved the first-pass effect (FPE) and revascularization rates by maximizing suction force and clot capture. Small diameter aspiration catheters have also been developed for distal vessel occlusion (117). Like stent retrievers, the efficacy of aspiration catheters also depends heavily on clot

characteristics, demonstrating efficacy in soft, fibrin-rich clots (118). Aspiration catheters also favor lesions where the angle between the catheter and the clot is greater than 125.5° (119). Aspiration is particularly advantageous in reducing procedural time and complications like embolization to new territories. It is also associated with less endothelial damage (116).

Combined techniques utilizing both stent retrievers and aspiration catheters have emerged to address the limitations of individual devices. Techniques like Solumbra and SAVE (Stent Retriever Assisted Vacuum-locked Extraction) capture the thrombus between the stent and catheter, enhancing recanalization rates (120, 121). The ASTER (Contact Aspiration vs. Stent Retriever for Successful Revascularization) and COMPASS (Comparison of Direct Aspiration vs. Stent Retriever as a First Approach) trials found no significant difference in outcomes between stent retrievers and aspiration used as standalone strategies (122, 123). Similarly, the ASTER2 trial did not show a significant improvement in final near-complete or complete reperfusion (eTICI 2c/3) with the combined approach combined to stent retriever alone, though the combination did result in higher early successful reperfusion after the first device pass (124). The VECTOR trial further found no significant advantage of the combined technique over contact aspiration alone in patients with susceptibility vessel sign-positive occlusions (125). Finally, the Separator-3D trial demonstrated noninferiority of a 3D stent retriever with aspiration compared to aspiration alone, with no significant differences in clinical or angiographic outcomes (126).

Combined techniques may still offer practical benefits in certain clinical contexts, such as clot heterogeneity and vessel tortuosity (127). The addition of balloon guide catheters (BGCs) to these techniques has traditionally been thought to improve procedural outcomes and reduce embolic complications (128). However, recent evidence from the PROTECT-MT trial raises concerns about their use (129). In this large multicenter RCT, BGC use was associated with significantly worse 90-day functional outcomes and a higher, though not statistically significant, rate of mortality compared to conventional guide catheters. These results challenge previous assumptions and highlight the need for further studies to clarify the role of BGCs in modern thrombectomy workflows.

5.2 Personalization through thrombectomy approaches

While transfemoral arterial access (TFA) is the standard approach, transradial access (TRA) or direct carotid puncture can be employed in specific cases. TRA is becoming increasingly popular among neurointerventionalists due to its reduced rate of access site complications, improved patient satisfaction, earlier ambulation, and shorter hospitalization (130, 131). However, this approach is technically difficult, requiring a steeper learning curve and specific patient anatomical patterns to ensure procedural success (132, 133). Early case series have demonstrated the safety and feasibility of TRA for anterior circulation LVOs, with results comparable to conventional

transfemoral systems (134). A comprehensive meta-analysis involving 13 studies and 4,759 patients has also confirmed safety and efficacy of TRA in acute ischemic stroke, revealing comparable recanalization rates, first pass reperfusion rates, mean number of passes, mortality and sICH rates, and post-treatment mRS and NIHSS scores to TFA (135). TRA is particularly suited for patients with challenging vascular anatomy, including those with a high B.A.D. (Bovine arch variant, Aortic arch type, and ICA Dolichoarteriopathy) score (136, 137). The B.A.D. is a composite score that assigns points based on the presence of a bovine arch (1 point), aortic arch type II (1 point) or III (2 points), and ipsilateral ICA tortuosity or coiling (1 point), with higher scores (≥ 2) indicating more difficult access (137). TRA may also benefit patients with complex comorbidities, prior vascular procedures, and older age (131, 138). While the growing evidence supports TRA as a viable alternative in appropriate patient groups, more studies and integrative radial-specific catheter development are necessary to establish broader consensus and widespread adoption.

When TFA or TRA fail due to anatomical constraints, direct carotid puncture serves as a rescue strategy in improving angiographic and functional outcomes (139). Direct carotid puncture allows direct access to the occlusion site, bypassing tortuous or diseased vessels. This technique allows direct access to the occlusion site, favoring vessels with high B.A.D. scores and patients with co-existing vascular pathologies (137, 140). Although this technique can result in complications of the punctured vessel, thromboembolic events, and airway compromise, only 1%–2% of affected patients will require re-intervention (141).

6 Conclusions

Through advances in neurovascular imaging and developments in thrombectomy devices and approaches, thrombectomy has revolutionized AIS treatment, with current evidence demonstrating a shift to personalized approaches to optimize outcomes. Through careful patient selection that incorporates patient-specific, lesion-specific, and biomarker-guided strategies, the paradigm shift in stroke thrombectomy has expanded patient eligibility criteria, allowing safe and effective interventions for previously excluded groups. The increase in patient load requires appropriate distribution of the neurovascular workforce to ensure equitable access to stroke thrombectomy. However, challenges persist in refining patient selection, with ongoing trials aiming to inform treatment decisions whilst balancing treatment-related risks. Further studies are warranted to refine the role of patient-specific biomarkers, incorporating emerging tools like artificial intelligence to enhance stroke management.

Author contributions

EM: Conceptualization, Data curation, Investigation, Methodology, Writing – original draft, Writing – review & editing.

Y-WF: Data curation, Investigation, Supervision, Validation, Writing – review & editing. SM: Data curation, Project administration, Supervision, Writing – review & editing. CO: Project administration, Supervision, Validation, Writing – review & editing. PT: Conceptualization, Methodology, Project administration, Supervision, Validation, Writing – review & editing.

Funding

The author(s) declare that no financial support was received for the research and/or publication of this article.

Conflict of interest

The authors declare that the research was conducted in the absence of any commercial or financial relationships that could be construed as a potential conflict of interest.

The author(s) declared that they were an editorial board member of Frontiers, at the time of submission. This had no impact on the peer review process and the final decision.

References

- 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) 20:795–820. doi: 10.1016/S1474-4422(21)00252-0
- Virani SS, Alonso A, Benjamin EJ, Bittencourt MS, Callaway CW, Carson AP, et al. Heart disease and stroke statistics-2020 update: a report from the American Heart Association. *Circulation.* (2020) 141:e139–596. doi: 10.1161/CIR.0000000000000757
- Rennett RC, Wali AR, Steinberg JA, Santiago-Dieppa DR, Olson SE, Pannell JS, et al. Epidemiology, natural history, and clinical presentation of large vessel ischemic stroke. *Neurosurgery.* (2019) 85:S4–8. doi: 10.1093/neuros/nyz042
- Berkhemer OA, Fransen PS, Beumer D, van den Berg LA, Lingsma HF, Yoo AJ, et al. A randomized trial of intraarterial treatment for acute ischemic stroke. *N Engl J Med.* (2015) 372:11–20. doi: 10.1056/NEJMoa1411587
- Saver JL, Goyal M, Bonafe A, Diener HC, Levy EI, Pereira VM, et al. Stent-retriever thrombectomy after intravenous t-PA vs. t-PA alone in stroke. *N Engl J Med.* (2015) 372:2285–95. doi: 10.1056/NEJMoa1415061
- Campbell BCV, Mitchell PJ, Kleinig TJ, Dewey HM, Churilov L, Yassi N, et al. Endovascular therapy for ischemic stroke with perfusion-imaging selection. *N Engl J Med.* (2015) 372:1009–18. doi: 10.1056/NEJMoa1414792
- Jovin TG, Chamorro A, Cobo E, de Miquel MA, Molina CA, Rovira A, et al. Thrombectomy within 8 h after symptom onset in ischemic stroke. *N Engl J Med.* (2015) 372:2296–306. doi: 10.1056/NEJMoa1503780
- Goyal M, Demchuk AM, Menon BK, Eesa M, Rempel JL, Thornton J, et al. Randomized assessment of rapid endovascular treatment of ischemic stroke. *N Engl J Med.* (2015) 372:1019–30. doi: 10.1056/NEJMoa1414905
- Goyal M, Menon BK, van Zwam WH, Dippel DW, Mitchell PJ, Demchuk AM, et al. Endovascular thrombectomy after large-vessel ischaemic stroke: a meta-analysis of individual patient data from five randomised trials. *Lancet.* (2016) 387:1723–31. doi: 10.1016/S0140-6736(16)00163-X
- Powers WJ, Rabinstein AA, Ackerson T, Adeoye OM, Bambakidis NC, Becker K, et al. 2018 guidelines for the early management of patients with acute ischemic stroke: a guideline for healthcare professionals from the American Heart Association/American stroke association. *Stroke.* (2018) 49:e46–110. doi: 10.1161/STR.0000000000000158
- Nogueira RG, Jadhav AP, Haussen DC, Bonafe A, Budzik RF, Bhuva P, et al. Thrombectomy 6–24 h after stroke with a mismatch between deficit and infarct. *N Engl J Med.* (2018) 378:11–21. doi: 10.1056/NEJMoa1706442
- Albers GW, Marks MP, Kemp S, Christensen S, Tsai JP, Ortega-Gutierrez S, et al. Thrombectomy for stroke at 6–16 h with selection by perfusion imaging. *N Engl J Med.* (2018) 378:708–18. doi: 10.1056/NEJMoa1713973
- Jovin TG, Nogueira RG, Lansberg MG, Demchuk AM, Martins SO, Mocco J, et al. Thrombectomy for anterior circulation stroke beyond 6 h from time last known well (AURORA): a systematic review and individual patient data meta-analysis. *Lancet.* (2022) 399:249–58. doi: 10.1016/S0140-6736(21)01341-6
- Bang OY, Goyal M, Liebeskind DS. Collateral circulation in ischemic stroke: assessment tools and therapeutic strategies. *Stroke.* (2015) 46:3302–9. doi: 10.1161/STROKEAHA.115.010508
- Regenhardt RW, González RG, He J, Lev MH, Singhal AB. Symmetric CTA collaterals identify patients with slow-progressing stroke likely to benefit from late thrombectomy. *Radiology.* (2022) 302:400–7. doi: 10.1148/radiol.2021210455
- Vagal A, Aviv R, Sucharew H, Reddy M, Hou Q, Michel P, et al. Collateral clock is more important than time clock for tissue fate. *Stroke.* (2018) 49:2102–7. doi: 10.1161/STROKEAHA.118.021484
- Olthuis SGH, Pirson FAV, Pinckaers FME, Hinsenvelde WH, Nieboer D, Ceulemans A, et al. Endovascular treatment versus no endovascular treatment after 6–24 h in patients with ischaemic stroke and collateral flow on CT angiography (MR CLEAN-LATE) in The Netherlands: a multicentre, open-label, blinded-endpoint, randomised, controlled, phase 3 trial. *Lancet.* (2023) 401:1371–80. doi: 10.1016/S0140-6736(23)00575-5
- Huijberts I, Pinckaers FME, Olthuis SGH, van Kuijk SMJ, Postma AA, Boogaarts HD, et al. Collateral-based selection for endovascular treatment of acute ischaemic stroke in the late window (MR CLEAN-LATE): 2-year follow-up of a phase 3, multicentre, open-label, randomised controlled trial in The Netherlands. *Lancet Neurol.* (2024) 23:893–900. doi: 10.1016/S1474-4422(24)00228-X
- Saver JL, Chapot R, Agid R, Hassan A, Jadhav AP, Liebeskind DS, et al. Thrombectomy for distal, medium vessel occlusions: a consensus statement on present knowledge and promising directions. *Stroke.* (2020) 51:2872–84. doi: 10.1161/STROKEAHA.120.028956
- Goyal M, Ospel JM, Menon BK, Hill MD. MeVO: the next frontier? *J Neurointerv Surg.* (2020) 12:545–7. doi: 10.1136/neurintsurg-2020-015807
- Mistry EA, Dumont AS. Medium vessel occlusion and acute ischemic stroke: a call for treatment paradigm reappraisal. *Stroke.* (2020) 51:3200–2. doi: 10.1161/STROKEAHA.120.032357
- Nguyen TN, Abdalkader M, Fischer U, Qiu Z, Nagel S, Chen HS, et al. Endovascular management of acute stroke. *Lancet.* (2024) 404:1265–78. doi: 10.1016/S0140-6736(24)01410-7
- Ospel JM, Goyal M. A review of endovascular treatment for medium vessel occlusion stroke. *J Neurointerv Surg.* (2021) 13:623–30. doi: 10.1136/neurintsurg-2021-017321
- Ospel JM, Menon BK, Demchuk AM, Almekhlafi MA, Kashani N, Mayank A, et al. Clinical course of acute ischemic stroke due to medium vessel occlusion with

Correction Note

This article has been corrected with minor changes. These changes do not impact the scientific content of the article.

Generative AI statement

The author(s) declare that no Generative AI was used in the creation of this manuscript.

Publisher's note

All claims expressed in this article are solely those of the authors and do not necessarily represent those of their affiliated organizations, or those of the publisher, the editors and the reviewers. Any product that may be evaluated in this article, or claim that may be made by its manufacturer, is not guaranteed or endorsed by the publisher.

- and without intravenous alteplase treatment. *Stroke*. (2020) 51:3232–40. doi: 10.1161/STROKEAHA.120.030227
25. Fiehler J, Nawka MT, Meyer L. Persistent challenges in endovascular treatment decision-making for acute ischaemic stroke. *Curr Opin Neurol*. (2022) 35:18–23. doi: 10.1097/WCO.0000000000001006
26. Molad J, Honig A. Current advances in endovascular treatment. *Curr Opin Neurol*. (2023) 36:125–30. doi: 10.1097/WCO.0000000000001142
27. Sahoo A, Abdalkader M, Yamagami H, Huo X, Sun D, Jia B, et al. Endovascular therapy for acute stroke: new evidence and indications. *J Neuroendovasc Ther*. (2023) 17:232–42. doi: 10.5797/jnet.ra.2023-0047
28. Jadhav AP, Desai SM, Jovin TG. Indications for mechanical thrombectomy for acute ischemic stroke: evidence and beyond. *Neurology*. (2021) 97:S126–36. doi: 10.1212/WNL.00000000000012801
29. Al Saiegh F, Munoz A, Velagapudi L, Theofanis T, Suryadevara N, Patel P, et al. Patient and procedure selection for mechanical thrombectomy: toward personalized medicine and the role of artificial intelligence. *J Neuroimaging*. (2022) 32:798–807. doi: 10.1111/jon.13003
30. Mokin M, Ansari SA, McTaggart RA, Bulsara KR, Goyal M, Chen M, et al. Indications for thrombectomy in acute ischemic stroke from emergent large vessel occlusion (ELVO): report of the SNIS standards and guidelines committee. *J Neurointerv Surg*. (2019) 11:215–20. doi: 10.1136/neurintsurg-2018-014640
31. Grossberg JA, Rebello LC, Haussen DC, Bousslama M, Bowen M, Barreira CM, et al. Beyond large vessel occlusion strokes: distal occlusion thrombectomy. *Stroke*. (2018) 49:1662–8. doi: 10.1161/STROKEAHA.118.020567
32. Menon BK, Hill MD, Davalos A, Roos YBWEM, Campbell BCV, Dippel DWJ, et al. Efficacy of endovascular thrombectomy in patients with M2 segment middle cerebral artery occlusions: meta-analysis of data from the HERMES collaboration. *J Neurointerv Surg*. (2019) 11:1065–9. doi: 10.1136/neurintsurg-2018-014678
33. Waqas M, Kuo CC, Dossani RH, Monteiro A, Baig AA, Alkhalidi M, et al. Mechanical thrombectomy versus intravenous thrombolysis for distal large-vessel occlusion: a systematic review and meta-analysis of observational studies. *Neurosurg Focus*. (2021) 51:E5. doi: 10.3171/2021.4.FOCUS21139
34. Meyer L, Stracke CP, Jungi N, Wallocha M, Brooks G, Sporns PB, et al. Thrombectomy for primary distal posterior cerebral artery occlusion stroke: the TOPMOST study. *JAMA Neurol*. (2021) 78:434–44. doi: 10.1001/jamaneurol.2021.0001
35. Meyer L, Stracke P, Brooks G, Elsharkawy M, Sporns P, Piechowiak EI, et al. Thrombectomy versus medical management for isolated anterior cerebral artery stroke: an international multicenter registry study. *Radiology*. (2023) 307:e220229. doi: 10.1148/radiol.220229
36. Bücke P, Cohen JE, Horvath T, Cimpoia A, Bhogal P, Bärner H, et al. What you always wanted to know about endovascular therapy in acute ischemic stroke but never dared to ask: A comprehensive review. *Rev Cardiovasc Med*. (2022) 23:340. doi: 10.31083/j.rcm.2310340
37. Goyal M, Ospel JM, Ganesh A, Dowlatabadi D, Volders D, Möhlenbruch MA, et al. Endovascular treatment of stroke due to medium-vessel occlusion. *N Engl J Med*. (2025) 392:1385–95. doi: 10.1056/NEJMoa2411668
38. Psychogios M, Brehm A, Ribo M, Rizzo F, Strbian D, Rätty S, et al. Endovascular treatment for stroke due to occlusion of medium or distal vessels. *N Engl J Med*. (2025) 392:1374–84. doi: 10.1056/NEJMoa2408954
39. Schonewille WJ, Wijman CA, Michel P, Rueckert CM, Weimar C, Mattle HP, et al. Treatment and outcomes of acute basilar artery occlusion in the basilar artery international cooperation study (BASICS): a prospective registry study. *Lancet Neurol*. (2009) 8:724–30. doi: 10.1016/S1474-4422(09)70173-5
40. Schaefer PW, Yoo AJ, Bell D, Barak ER, Romero JM, Nogueira RG, et al. CT angiography-source image hypoattenuation predicts clinical outcome in posterior circulation strokes treated with intra-arterial therapy. *Stroke*. (2008) 39:3107–9. doi: 10.1161/STROKEAHA.108.517680
41. Alemseged F, Rocco A, Arba F, Schwabova JP, Wu T, Cavicchia L, et al. Posterior national institutes of health stroke scale improves prognostic accuracy in posterior circulation stroke. *Stroke*. (2022) 53:1247–55. doi: 10.1161/STROKEAHA.120.034019
42. Lavie J, Vellimana AK, Chatterjee AR. Endovascular thrombectomy treatment: beyond early time windows and small core. *Top Magn Reson Imaging*. (2021) 30:173–80. doi: 10.1097/RMR.0000000000000291
43. Zi W, Qiu Z, Wu D, Li F, Liu H, Liu W, et al. Assessment of endovascular treatment for acute basilar artery occlusion via a nationwide prospective registry. *JAMA Neurol*. (2020) 77:561–73. doi: 10.1001/jamaneurol.2020.0156
44. Liu X, Dai Q, Ye R, Zi W, Liu Y, Wang H, et al. Endovascular treatment versus standard medical treatment for vertebrobasilar artery occlusion (BEST): an open-label, randomised controlled trial. *Lancet Neurol*. (2020) 19:115–22. doi: 10.1016/S1474-4422(19)30395-3
45. Langezaal LCM, van der Hoeven EJRI, Mont'Alverne FJA, de Carvalho JFF, Lima FO, Dippel DW, et al. Endovascular therapy for stroke due to basilar-artery occlusion. *N Engl J Med*. (2021) 384:1910–20. doi: 10.1056/NEJMoa2030297
46. Tao C, Nogueira RG, Zhu Y, Sun J, Han H, Yuan G, et al. Trial of endovascular treatment of acute basilar-artery occlusion. *N Engl J Med*. (2022) 387:1361–72. doi: 10.1056/NEJMoa2206317
47. Jovin TG, Li C, Wu L, Wu C, Chen J, Jiang C, et al. Trial of thrombectomy 6–24 h after stroke due to basilar-artery occlusion. *N Engl J Med*. (2022) 387:1373–84. doi: 10.1056/NEJMoa2207576
48. Rubiera M, Ribo M, Delgado-Mederos R, Santamarina E, Delgado P, Montaner J, et al. Tandem internal carotid artery/middle cerebral artery occlusion: an independent predictor of poor outcome after systemic thrombolysis. *Stroke*. (2006) 37:2301–5. doi: 10.1161/01.STR.0000237070.80133.1d
49. Jacquin G, Poppe AY, Labrie M, Daneault N, Deschaintre Y, Gioia LC, et al. Lack of consensus among stroke experts on the optimal management of patients with acute tandem occlusion. *Stroke*. (2019) 50:1254–6. doi: 10.1161/STROKEAHA.118.023758
50. Tsvigoulis G, Katsanos AH, Schellinger PD, Köhrmann M, Varelas P, Magoufis G, et al. Successful reperfusion with intravenous thrombolysis preceding mechanical thrombectomy in large-vessel occlusions. *Stroke*. (2018) 49:232–5. doi: 10.1161/STROKEAHA.117.019261
51. Wilson MP, Murad MH, Krings T, Pereira VM, O'Kelly C, Rempel J, et al. Management of tandem occlusions in acute ischemic stroke—intracranial versus extracranial first and extracranial stenting versus angioplasty alone: a systematic review and meta-analysis. *J Neurointerv Surg*. (2018) 10:721–8. doi: 10.1136/neurintsurg-2017-013707
52. Haussen DC, Turjman F, Piotin M, Labreuche J, Steglich-Arnholm H, Holtmannspötter M, et al. Head or neck first? Speed and rates of reperfusion in thrombectomy for tandem large vessel occlusion strokes. *Interv Neurol*. (2020) 8:92–100. doi: 10.1159/000496292
53. Raha O, Hall C, Malik A, D'Anna L, Lobotesis K, Kwan J, et al. Advances in mechanical thrombectomy for acute ischaemic stroke. *BMJ Med*. (2023) 2:e000407. doi: 10.1136/bmjmed-2022-00040754
54. Dufort G, Chen BY, Jacquin G, Keezer M, Labrie M, Rioux B, et al. Acute carotid stenting in patients undergoing thrombectomy: a systematic review and meta-analysis. *J Neurointerv Surg*. (2021) 13:141–5. doi: 10.1136/neurintsurg-2020-015817
55. Anadani M, Marnat G, Consoli A, Papanagiotou P, Nogueira RG, Siddiqui A, et al. Endovascular therapy of anterior circulation tandem occlusions: pooled analysis from the TITAN and ETIS registries. *Stroke*. (2021) 52:3097–105. doi: 10.1161/STROKEAHA.120.033032
56. ClinicalTrials.gov. Stent thrombosis after tandem lesions thrombectomy (DST-TANDEM) (2018). Available online at: <https://clinicaltrials.gov/study/NCT03588780> (Accessed January 21, 2025).
57. Zhu F, Hossu G, Soudant M, Richard S, Achit H, Beguin M, et al. Effect of emergent carotid stenting during endovascular therapy for acute anterior circulation stroke patients with tandem occlusion: a multicenter, randomized, clinical trial (TITAN) protocol. *Int J Stroke*. (2021) 16:342–8. doi: 10.1177/1747493020929948
58. ClinicalTrials.gov. Endovascular Acute Stroke Intervention—Tandem Occlusion Trial (EASI-TOC) (2023). Available online at: <https://clinicaltrials.gov/study/NCT04261478> (Accessed January 21, 2025).
59. Campbell D, Butler E, Barber PA. End the confusion: general anaesthesia improves patient outcomes in endovascular thrombectomy. *Br J Anaesth*. (2022) 129:461–4. doi: 10.1016/j.bja.2022.06.018
60. Wollenweber FA, Tiedt S, Alegiani A, Alber B, Bangard C, Berrouschot J, et al. Functional outcome following stroke thrombectomy in clinical practice. *Stroke*. (2019) 50:2500–6. doi: 10.1161/STROKEAHA.119.026005
61. Siegler JE, Qureshi MM, Nogueira RG, Tanaka K, Nagel S, Michel P, et al. Endovascular vs medical management for late anterior large vessel occlusion with prestroke disability: analysis of CLEAR and RESCUE-Japan. *Neurology*. (2023) 100:e751–63. doi: 10.1212/WNL.000000000000201543
62. Sarraj A, Hassan AE, Savitz S, Sitton C, Grotta J, Chen P, et al. Outcomes of endovascular thrombectomy vs medical management alone in patients with large ischemic cores: a secondary analysis of the optimizing patient's selection for endovascular treatment in acute ischemic stroke (SELECT) study. *JAMA Neurol*. (2019) 76:1147–56. doi: 10.1001/jamaneurol.2019.2109
63. Huo X, Ma G, Tong X, Zhang X, Pan Y, Nguyen TN, et al. Trial of endovascular therapy for acute ischemic stroke with large infarct. *N Engl J Med*. (2023) 388:1272–83. doi: 10.1056/NEJMoa2213379
64. Sanmartin MX, Katz JM, Wang J, Malhotra A, Sangha K, Bastani M, et al. Cost-effectiveness of endovascular thrombectomy in acute stroke patients with large ischemic core. *J Neurointerv Surg*. (2023) 15:E166–71. doi: 10.1136/jnis-2022-019460
65. Rumalla K, Ottenhausen M, Kan P, Burkhardt J-K. Recent nationwide impact of mechanical thrombectomy on decompressive hemicraniectomy for acute ischemic stroke. *Stroke*. (2019) 50:2133–9. doi: 10.1161/STROKEAHA.119.025063
66. Bendszus M, Fiehler J, Subtil F, Bonekamp S, Aamodt AH, Fuentes B, et al. Endovascular thrombectomy for acute ischaemic stroke with established large infarct: multicentre, open-label, randomised trial. *Lancet*. (2023) 402:1753–63. doi: 10.1016/S0140-6736(23)02032-9
67. Writing Committee for the TESLA Investigators, Yoo AJ, Zaidat OO, Sheth SA, Rai AT, Ortega-Gutierrez S, et al. Thrombectomy for stroke with large infarct on noncontrast CT: the TESLA randomized clinical trial. *JAMA*. (2024) 332:1355–66. doi: 10.1001/jama.2024.13933

68. Costalat V, Jovin TG, Albuher JF, Cognard C, Henon H, Nouri N, et al. Trial of thrombectomy for stroke with a large infarct of unrestricted size. *N Engl J Med.* (2024) 390:1677–89. doi: 10.1056/NEJMoa2314063
69. Boers AMM, Jansen IGH, Beenen LFM, Devlin TG, San Roman L, Heo JH, et al. Association of follow-up infarct volume with functional outcome in acute ischemic stroke: a pooled analysis of seven randomized trials. *J Neurointerv Surg.* (2018) 10:1137–42. doi: 10.1136/neurintsurg-2017-013724
70. Zhou Y, Zhang R, Yan S, Zhang M, Chen Z, Hu H, et al. Identification of corticospinal tract lesion for predicting outcome in small perfusion stroke. *Stroke.* (2018) 49:2683–91. doi: 10.1161/STROKEAHA.118.021426
71. Baron J-C. The core/penumbra model: implications for acute stroke treatment and patient selection in 2021. *Eur J Neurol.* (2021) 28:2794–803. doi: 10.1111/ene.14916
72. Motyer R, Thornton J, Power S, Brennan P, O'Hare A, Looby S, et al. Endovascular thrombectomy beyond 12 h of stroke onset: a stroke network's experience of late intervention. *J Neurointerv Surg.* (2018) 10:1043–6. doi: 10.1136/neurintsurg-2017-013575
73. DiBiasio EL, Jayaraman MV, Goyal M, Yaghi S, Tung E, Hidlay DT, et al. Dismantling the ability of CT and MRI to identify the target mismatch profile in patients with anterior circulation large vessel occlusion beyond six hours from symptom onset. *Emerg Radiol.* (2019) 26:401–8. doi: 10.1007/s10140-019-01686-z
74. Lyden P. Using the national institutes of health stroke scale: a cautionary tale. *Stroke.* (2017) 48:513–9. doi: 10.1161/STROKEAHA.116.015434
75. Heldner MR, Hsieh K, Broeg-Morway A, Mordasini P, Bühlmann M, Jung S, et al. Clinical prediction of large vessel occlusion in anterior circulation stroke: mission impossible? *J Neurol.* (2016) 263:1633–40. doi: 10.1007/s00415-016-8180-6
76. Asdaghi N, Yavagal DR, Wang K, Mueller-Kronast N, Bhatt N, Gardener HE, et al. Patterns and outcomes of endovascular therapy in mild stroke. *Stroke.* (2019) 50:2101–7. doi: 10.1161/STROKEAHA.118.023893
77. Duloquin G, Crespy V, Jakubina P, Giroud M, Vergely C, Béjot Y. Large vessel occlusion in patients with minor ischemic stroke in a population-based study. The Dijon stroke registry. *Front Neurol.* (2022) 12:796046. doi: 10.3389/fneur.2021.796046
78. Kim J-T, Park MS, Chang J, Lee JS, Choi KH, Cho KH. Proximal arterial occlusion in acute ischemic stroke with low NIHSS scores should not be considered as mild stroke. *PLoS One.* (2013) 8:e70996. doi: 10.1371/journal.pone.0070996
79. Seners P, Ben Hassen W, Lapergue B, Arquiza C, Heldner MR, Henon H, et al. Prediction of early neurological deterioration in individuals with minor stroke and large vessel occlusion intended for intravenous thrombolysis alone. *JAMA Neurol.* (2021) 78:321. doi: 10.1001/jamaneurol.2020.4557
80. Dargazanli C, Arquiza C, Gory B, Consoli A, Labreuche J, Redjem H, et al. Mechanical thrombectomy for minor and mild stroke patients harboring large vessel occlusion in the anterior circulation: a multicenter cohort study. *Stroke.* (2017) 48:3274–81. doi: 10.1161/STROKEAHA.117.018113
81. Goyal N, Tsvigoulis G, Malhotra K, Ishfaq MF, Pandhi A, Frohler MT, et al. Medical management vs mechanical thrombectomy for mild strokes: an international multicenter study and systematic review and meta-analysis. *JAMA Neurol.* (2020) 77:16–24. doi: 10.1001/jamaneurol.2019.3112
82. Volny O, Zerna C, Tomek A, Bar M, Roczek M, Padr R, et al. Thrombectomy vs medical management in low NIHSS acute anterior circulation stroke. *Neurology.* (2020) 95:E3364–72. doi: 10.1212/WNL.0000000000010955
83. McCarthy DJ, Tonetti DA, Stone J, Starke RM, Narayanan S, Lang MJ, et al. More expansive horizons: a review of endovascular therapy for patients with low NIHSS scores. *J Neurointerv Surg.* (2021) 13:146–51. doi: 10.1136/neurintsurg-2020-016583
84. Bilgin C, Bolseguí ML, Ghosy S, Hassankhani A, Kobeissi H, Jabal MS, et al. Common design and data elements reported in active mechanical thrombectomy trials focusing on distal medium vessel occlusions and minor strokes: a systematic review. *J Neurointerv Surg.* (2024) 17(5):530–8. doi: 10.1136/JNIS-2023-021073
85. Rotschild O, Honig A, Halleli H, Horev A, Seyman E, Ben-Assayag E, et al. Endovascular thrombectomy is beneficial for functional nonagenarians—a multicenter cohort analysis. *J Stroke Cerebrovasc Dis.* (2022) 31(10):106699. doi: 10.1016/j.jstrokecerebrovasdis.2022.106699
86. Derex L, Cho TH. Mechanical thrombectomy in acute ischemic stroke. *Rev Neurol (Paris).* (2017) 173:106–13. doi: 10.1016/j.neurol.2016.06.008
87. Meyer L, Alexandrou M, Leischner H, Flottmann F, Deb-Chatterji M, Abdullayev N, et al. Mechanical thrombectomy in nonagenarians with acute ischemic stroke. *J Neurointerv Surg.* (2019) 11:1091–4. doi: 10.1136/neurintsurg-2019-014785
88. Pinto MM, Nunes AP, Alves M, Papoila AL, Pamplona J, Carvalho R, et al. Mechanical thrombectomy in stroke in nonagenarians: useful or futile? *J Stroke Cerebrovasc Dis.* (2020) 29(9):105015. doi: 10.1016/j.jstrokecerebrovasdis.2020.105015
89. Sugar BP, Drasler NE, Lee J, Beutler BD, Moody AE, Cadavona JJP, et al. Utility of thrombectomy in nonagenarians: a scoping review. *Eur Neurol.* (2024) 87:122–9. doi: 10.1159/000539789
90. Lasek-Bal A, Binek Ł, Żak A, Student S, Krzan A, Puz P, et al. Clinical and non-clinical determinants of the effect of mechanical thrombectomy and post-stroke functional status of patients in short and long-term follow-up. *J Clin Med.* (2021) 10(21):5084. doi: 10.3390/jcm10215084
91. Bai X, Zhang X, Zhang Y, Yang W, Wang T, Feng Y, et al. Mechanical thrombectomy in nonagenarians: a systematic review and meta-analysis. *Transl Stroke Res.* (2021) 12:394–405. doi: 10.1007/s12975-021-00894-5
92. Hendrix P, Killer-Oberpfalzer M, Broussalis E, Melamed I, Sharma V, Mutzenbach S, et al. Outcome following mechanical thrombectomy for anterior circulation large vessel occlusion stroke in the elderly. *Clin Neuroradiol.* (2022) 32:369–74. doi: 10.1007/s00062-021-01063-9
93. Turc G, Tsvigoulis G, Audebert HJ, Boogaarts H, Bhogal P, De Marchis GM, 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. *Eur Stroke J.* (2022) 7(1):1–XXVI. doi: 10.1177/23969873221076968
94. Venditti L, Chassin O, Ancelet C, Legris N, Sarov M, Lapergue B, et al. Pre-procedural predictive factors of symptomatic intracranial hemorrhage after thrombectomy in stroke. *J Neurol.* (2021) 268:1867–75. doi: 10.1007/s00415-020-10364-x
95. Li X, Li C, Shi M, Qu Y, Huo L, Hao Z, et al. Which glucose parameter best predicts poor outcome after mechanical thrombectomy for acute large vessel occlusion stroke? *Intern Med J.* (2022) 52:1374–80. doi: 10.1111/imj.15259
96. Xu J, Jiang X, Liu Q, Liu J, Fang J, He L. Lower serum uric acid to serum creatinine ratio as a predictor of poor functional outcome after mechanical thrombectomy in acute ischaemic stroke. *Eur J Neurol.* (2024) 31(7):e16296. doi: 10.1111/ene.16296
97. Tsvigoulis G, Katsanos AH. TAG for symptomatic intracerebral haemorrhage following mechanical thrombectomy. *J Neurol Neurosurg Psychiatry.* (2019) 90:1307. doi: 10.1136/jnnp-2019-321807
98. Zi W, Wang H, Yang D, Hao Y, Zhang M, Geng Y, et al. Clinical effectiveness and safety outcomes of endovascular treatment for acute anterior circulation ischemic stroke in China. *Cerebrovasc Dis.* (2017) 44:248–58. doi: 10.1159/000478667
99. Kim S-Y, Yi HJ, Shin D-S, Kim B-T. Prognostic significance of platelet-to-lymphocyte and platelet-to-neutrophil ratios in patients with mechanical thrombectomy for acute ischemic stroke. *J Cerebrovasc Endovasc Neurosurg.* (2022) 24:221–31. doi: 10.7461/jcen.2022.E2021.10.003
100. Ji Y, Xu X, Wu K, Sun Y, Wang H, Guo Y, et al. Prognosis of ischemic stroke patients undergoing endovascular thrombectomy is influenced by systemic inflammatory Index through malignant brain edema. *Clin Interv Aging.* (2022) 17:1001–12. doi: 10.2147/CIA.S365553
101. Zheng F, Gao W, Xiao Y, Guo X, Xiong Y, Chen C, et al. Systemic inflammatory response index as a predictor of stroke-associated pneumonia in patients with acute ischemic stroke treated by thrombectomy: a retrospective study. *BMC Neurol.* (2024) 24:287. doi: 10.1186/s12883-024-03783-0
102. Matuszevicius M, Cooray C, Holmin S, Bottai M, Ahmed N. Association between systolic blood pressure course and outcomes after stroke thrombectomy. *BMJ Neurol Open.* (2021) 3(2):e000183. doi: 10.1136/bmjno-2021-000183
103. Inauen C, Boss JM, Katan M, Luft AR, Kulcsar Z, Willms JF, et al. Blood pressure variability indices for outcome prediction after thrombectomy in stroke by using high-resolution data. *Neurocrit Care.* (2022) 37:220–9. doi: 10.1007/s12028-022-01519-x
104. Maier B, Gory B, Chabanne R, Tavernier B, Balanca B, Audibert G, et al. Effect of an individualized versus standard blood pressure management during mechanical thrombectomy for anterior ischemic stroke: the DETERMINE randomized controlled trial. *Trials.* (2022) 23:598. doi: 10.1186/s13063-022-06538-9
105. Petersen NH, Silverman A, Strander SM, Kodali S, Wang A, Sansing LH, et al. Fixed compared with autoregulation-oriented blood pressure thresholds after mechanical thrombectomy for ischemic stroke. *Stroke.* (2020) 51:914–21. doi: 10.1161/STROKEAHA.119.026596
106. Thomas S, de la Pena P, Butler L, Akbilgic O, Heiferman DM, Garg R, et al. Machine learning models improve prediction of large vessel occlusion and mechanical thrombectomy candidacy in acute ischemic stroke. *J Clin Neurosci.* (2021) 91:383–90. doi: 10.1016/j.jocn.2021.07.021
107. Hu Y, Yang T, Zhang J, Wang X, Cui X, Chen N, et al. Dynamic prediction of mechanical thrombectomy outcome for acute ischemic stroke patients using machine learning. *Brain Sci.* (2022) 12(7):938. doi: 10.3390/brainsci12070938
108. Zhou S, Wei J, Tang L, Gu C, Liang J. A prognostic model for interventional thrombectomy in patients with acute ischemic stroke based on a BP neural network, random forest model and decision tree model. *Am J Transl Res.* (2023) 15:3290–9.
109. Teo YH, Lim ICZY, Tseng FS, Teo YN, Kow CS, Ng ZHC, et al. Predicting clinical outcomes in acute ischemic stroke patients undergoing endovascular thrombectomy with machine learning: a systematic review and meta-analysis. *Clin Neuroradiol.* (2021) 31:1121–30. doi: 10.1007/s00062-020-00990-3
110. Al-Mufti F, Schirmer CM, Starke RM, Chaudhary N, De Leacy R, Tjoumakaris SI, et al. Thrombectomy in special populations: report of the society of neurointerventional surgery standards and guidelines committee. *J Neurointerv Surg.* (2022) 14:1033–41. doi: 10.1136/neurintsurg-2021-017888

111. Rikhtegar R, Mosimann PJ, Weber R, Wallocha M, Yamac E, Mirza-Aghazadeh-Attari M, et al. Effectiveness of very low profile thrombectomy device in primary distal medium vessel occlusion, as rescue therapy after incomplete proximal recanalization or following iatrogenic thromboembolic events. *J Neurointerv Surg.* (2021) 13:1067–72. doi: 10.1136/neurintsurg-2020-017035
112. Fischer S, Will L, Phung T, Weber W, Maus V, Nordmeyer H. The tigertriever 13 for mechanical thrombectomy in distal and medium intracranial vessel occlusions. *Neuroradiology.* (2022) 64:775–83. doi: 10.1007/s00234-021-02792-x
113. Weafer FM, Duffy S, Machado I, Gunning G, Mordasini P, Roche E, et al. Characterization of strut indentation during mechanical thrombectomy in acute ischemic stroke clot analogs. *J Neurointerv Surg.* (2019) 11:891–7. doi: 10.1136/neurintsurg-2018-014601
114. Jovin TG, Albers GW, Liebeskind DS, STAIR IX Consortium. Stroke treatment academic industry roundtable: the next generation of endovascular trials. *Stroke.* (2016) 47:2656–65. doi: 10.1161/STROKEAHA.116.013578
115. Turk AS, Frei D, Fiorella D, Mocco J, Baxter B, Siddiqui A, et al. ADAPT FAST study: a direct aspiration first pass technique for acute stroke thrombectomy. *J Neurointerv Surg.* (2018) 10:14–7. doi: 10.1136/neurintsurg-2014-011125.rep
116. Peschillo S, Tomasello A, Diana F, Hernandez D, Toccaceli G, Rosal-Fontana M, et al. Comparison of subacute vascular damage caused by ADAPT versus stent retriever devices after thrombectomy in acute ischemic stroke: histological and ultrastructural study in an animal model. *Interv Neurol.* (2018) 7:501–12. doi: 10.1159/000491028
117. Dornbos D, Arthur AS. Current state of the art in endovascular stroke treatment. *Neurol Clin.* (2022) 40:309–19. doi: 10.1016/j.ncl.2021.11.008
118. Mohammaden MH, Haussen DC, Perry da Camara C, Pisani L, Olive Gadea M, Al-Bayati AR, et al. Hyperdense vessel sign as a potential guide for the choice of stent retriever versus contact aspiration as first-line thrombectomy strategy. *J Neurointerv Surg.* (2021) 13:599–604. doi: 10.1136/neurintsurg-2020-016005
119. Bernava G, Rosi A, Boto J, Brina O, Kulcsar Z, Czarnetzki C, et al. Direct thromboaspiration efficacy for mechanical thrombectomy is related to the angle of interaction between the aspiration catheter and the clot. *J Neurointerv Surg.* (2020) 12:396–400. doi: 10.1136/neurintsurg-2019-015113
120. Deshaies EM. Tri-axial system using the solitaire-FR and penumbra aspiration microcatheter for acute mechanical thrombectomy. *J Clin Neurosci.* (2013) 20:1303–5. doi: 10.1016/j.jocn.2012.10.037
121. Maus V, Behme D, Kabbasch C, Borggrefe J, Tsogkas I, Nikoubashman O, et al. Maximizing first-pass complete reperfusion with SAVE. *Clin Neuroradiol.* (2018) 28:327–38. doi: 10.1007/s00062-017-0566-z
122. Lapergue B, Blanc R, Gory B, Labreuche J, Duhamel A, Marnat G, et al. Effect of endovascular contact aspiration vs stent retriever on revascularization in patients with acute ischemic stroke and large vessel occlusion: the ASTER randomized clinical trial. *JAMA.* (2017) 318:443–52. doi: 10.1001/jama.2017.9644
123. Turk AS, Siddiqui A, Fifi JT, De Leacy RA, Fiorella DJ, Gu E, et al. Aspiration thrombectomy versus stent retriever thrombectomy as first-line approach for large vessel occlusion (COMPASS): a multicentre, randomised, open label, blinded outcome, non-inferiority trial. *Lancet.* (2019) 393:998–1008. doi: 10.1016/S0140-6736(19)30297-1
124. Lapergue B, Blanc R, Costalat V, Desal H, Saleme S, Spelle L, et al. Effect of thrombectomy with combined contact aspiration and stent retriever vs stent retriever alone on revascularization in patients with acute ischemic stroke and large vessel occlusion: the ASTER2 randomized clinical trial. *JAMA.* (2021) 326:1158–69. doi: 10.1001/jama.2021.13827
125. Bourcier R, Marnat G, Dargazanli C, Zhu F, Consoli A, Shotar E, et al. Safety and efficacy of stent retrievers plus contact aspiration in patients with acute ischaemic anterior circulation stroke and positive susceptibility vessel sign in France (VECTOR): a randomised, single-blind trial. *Lancet Neurol.* (2024) 23:700–11. doi: 10.1016/S1474-4422(24)00165-0
126. Nogueira RG, Frei D, Kirmani JF, Zaidat O, Lopes D, Turk AS, et al. Safety and efficacy of a 3-dimensional stent retriever with aspiration based thrombectomy vs aspiration-based thrombectomy alone in acute ischemic stroke intervention: a randomized clinical trial. *JAMA Neurol.* (2018) 75:304–11. doi: 10.1001/jamaneurol.2017.3967
127. Kaneko N, Sakuta K, Imahori T, Gedion H, Ghovvati M, Tateshima S. Devices and techniques. *J Neuroendovasc Ther.* (2023) 17:257–62. doi: 10.5797/jnet.ra.2023-0054
128. Brinjikji W, Starke RM, Murad MH, Fiorella D, Pereira VM, Goyal M, et al. Impact of balloon guide catheter on technical and clinical outcomes: a systematic review and meta-analysis. *J Neurointerv Surg.* (2018) 10:335–9. doi: 10.1136/neurintsurg-2017-013179
129. Liu J, Zhou Y, Zhang L, Li Z, Chen W, Zhu Y, et al. Balloon guide catheters for endovascular thrombectomy in patients with acute ischaemic stroke due to large-vessel occlusion in China (PROTECT-MT): a multicentre, open-label, blinded-endpoint, randomised controlled trial. *Lancet.* (2024) 404:2165–74. doi: 10.1016/S0140-6736(24)02315-8
130. Jolly SS, Yusuf S, Cairns J, Niemelä K, Xavier D, Widimsky P, et al. Radial versus femoral access for coronary angiography and intervention in patients with acute coronary syndromes (RIVAL): a randomised, parallel group, multicentre trial. *Lancet.* (2011) 377:1409–20. doi: 10.1016/S0140-6736(11)60404-2
131. Khanna O, Mouchtouris N, Sweid A, Chalouhi N, Ghosh R, Al Saiegh F, et al. Transradial approach for acute stroke intervention: technical procedure and clinical outcomes. *Stroke Vasc Neurol.* (2020) 5:103–6. doi: 10.1136/svn-2019-000263
132. Esente P, Giambartolomei A, Simons AJ, Levy C, Caputo RP. Overcoming vascular anatomic challenges to cardiac catheterization by the radial artery approach: specific techniques to improve success. *Catheter Cardiovasc Interv.* (2002) 56:207–11. doi: 10.1002/ccd.10233
133. Snelling BM, Sur S, Shah SS, Caplan J, Khandelwal P, Yavagal DR, et al. Transradial approach for complex anterior and posterior circulation interventions: technical nuances and feasibility of using current devices. *Oper Neurosurg (Hagerstown).* (2019) 17:293–302. doi: 10.1093/ons/opy352
134. Kuroiwa M, Hanaoka Y, Koyama JJ, Yamazaki D, Kubota Y, Kitamura S, et al. Transradial mechanical thrombectomy using a radial-specific neurointerventional guiding sheath for anterior circulation large-vessel occlusions. Preliminary experience and literature review. *World Neurosurg.* (2023) 171:e581–9. doi: 10.1016/j.wneu.2022.12.060
135. Almansi A, Alqato S, Yassin MNA, Taher LH, Batarseh SF, Nashwan AJ. Transradial versus transfemoral artery access in mechanical thrombectomy for acute ischemic stroke: an updated systematic review and meta-analysis. *Clin Neurol Neurosurg.* (2024) 246:108585. doi: 10.1016/j.clineuro.2024.108585
136. Chen SH, Snelling BM, Sur S, Shah SS, McCarthy DJ, Luther E, et al. Transradial versus transfemoral access for anterior circulation mechanical thrombectomy: comparison of technical and clinical outcomes. *J Neurointerv Surg.* (2019) 11:874–8. doi: 10.1136/neurintsurg-2018-014485
137. Snelling BM, Sur S, Shah SS, Chen S, Menaker SA, McCarthy DJ, et al. Unfavorable vascular anatomy is associated with increased revascularization time and worse outcome in anterior circulation thrombectomy. *World Neurosurg.* (2018) 120:e976–83. doi: 10.1016/j.wneu.2018.08.207
138. Mendiz OA, Sampaulesi AH, Londero HF, Fava CM, Lev GA, Valdivieso LR. Initial experience with transradial access for carotid artery stenting. *Vasc Endovascular Surg.* (2011) 45:499–503. doi: 10.1177/153857441140554
139. Cord BJ, Kodali S, Strander S, Silverman A, Wang A, Chouairi F, et al. Direct carotid puncture for mechanical thrombectomy in acute ischemic stroke patients with prohibitive vascular access. *J Neurosurg.* (2021) 135:53–63. doi: 10.3171/2020.5.JNS192737
140. Mokin M, Snyder KV, Levy EI, Hopkins LN, Siddiqui AH. Direct carotid artery puncture access for endovascular treatment of acute ischemic stroke: technical aspects, advantages, and limitations. *J Neurointerv Surg.* (2015) 7:108–13. doi: 10.1136/neurintsurg-2013-011007
141. Sokolowski JD, Mastorakos P, Fisher J, Kellogg RT. Direct carotid puncture for neuroendovascular procedures. *Acta Neurochir (Wien).* (2022) 164:1293–6. doi: 10.1007/s00701-022-05146-5