The most recent trials regarding the acute management of stroke have highlighted that treatment with endovascular therapy is associated with better functional outcome at 90 days.23 Of the 5 recent stent retriever trials, all but one (SWIFT PRIME) included patients with carotid occlusions and tandem distal occlusions treated with endovascular therapy. They reviewed the type of approach employed for endovascular intervention and clinical outcomes reported with emphasis on the revascularization technique. They also present an illustrative case of AIS and concurrent proximal cervical carotid occlusion and distal middle cerebral artery occlusion from their own experience in order to outline the management dilemma for similar cases.

Revascularization of tandem occlusions in acute ischemic stroke: review of the literature and illustrative case

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OBJECTIVE Tandem occlusions continue to represent a major challenge in patients with acute ischemic stroke (AIS). The anterograde approach with proximal to distal revascularization as well as the retrograde approach with distal to proximal revascularization have been reported without clear consensus or standard guidelines.

METHODS The authors performed a comprehensive search of the PubMed database for studies including patients with carotid occlusions and tandem distal occlusions treated with endovascular therapy. They reviewed the type of approach employed for endovascular intervention and clinical outcomes reported with emphasis on the revascularization technique. They also present an illustrative case of AIS and concurrent proximal cervical carotid occlusion and distal middle cerebral artery occlusion from their own experience in order to outline the management dilemma for similar cases.

RESULTS A total of 22 studies were identified, with a total of 790 patients with tandem occlusions in AIS. Eleven studies used the anterograde approach, 3 studies used the retrograde approach, 4 studies used both, and in 4 studies the approach was not specified. In the studies that reported Thrombolysis in Cerebral Infarction (TICI) grades, an average of 79% of patients with tandem occlusions were reported to have an outcome of TICI 2b or better. One study found good clinical outcome in 52.5% of the thrombectomy-first group versus 33.3% in the stent-first group, as measured by the modified Rankin Scale (mRS). No study evaluated the difference in time to reperfusion for the anterograde and retrograde approach and its association with clinical outcome. The patient in the illustrative case had AIS and tandem occlusion of the internal carotid and middle cerebral arteries and underwent distal revascularization using a Solitaire stent retrieval device followed by angioplasty and stent treatment of the proximal cervical carotid occlusion. The revascularization was graded as TICI 2b; the postintervention National Institutes of Health Stroke Scale (NIHSS) score was 17, and the discharge NIHSS score was 7. The admitting, postoperative, and 30-day mRS scores were 5, 1, and 1, respectively.

CONCLUSIONS In stroke patients with tandem occlusions, distal to proximal revascularization represents a reasonable treatment approach and may offer the advantage of decreased time to reperfusion, which is associated with better functional outcome. Further studies are warranted to determine the best techniques in endovascular therapy to use in this subset of patients in order to improve clinical outcome.

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KEY WORDS tandem occlusion; ischemic stroke; revascularization; endovascular therapy

ABBREVIATIONS ACA = anterior cerebral artery; AP = anteroposterior; AIS = acute ischemic stroke; CCA = common carotid artery; CTA = CT angiography; DSA = digital subtraction angiography; ICA = internal carotid artery; IV = intravenous; MCA = middle cerebral artery; mRS = modified Rankin Scale; NIHSS = National Institutes of Health Stroke Scale; TICI = Thrombolysis in Cerebral Infarction; tPA = tissue plasminogen activator.


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literature on the subject of the optimal strategy in regard to the technical approach to cases of stroke with tandem occlusions.

We present a review of the recent literature regarding tandem occlusions in acute ischemic stroke with a focus on the endovascular technique and related clinical outcome. We also present an illustrative case involving a patient with tandem occlusions of the cervical internal carotid artery (ICA) and middle cerebral artery (MCA) who was treated with distal to proximal revascularization.

Literature Review
Methods
A search of the PubMed database identified all studies between 2011 and 2016 that included patients with tandem occlusions in AIS treated with endovascular therapy. Tandem occlusion was defined as a proximal carotid artery occlusion with concomitant distal intracranial occlusion. Anterograde therapy was defined as proximal to distal revascularization using stent placement and/or angioplasty of the proximal occlusion before distal revascularization. Distal revascularization techniques included stent retrieval, intra-arterial thrombolysis, microwire thrombus manipulation, or thromboaspiration. Retrograde therapy was defined as distal to proximal revascularization with the above methods applied in the reverse order (treatment of the distal occlusion before treatment of the proximal occlusion). The clinical outcome identified for each subset of patients was also recorded as defined by the respective studies.

Results
A review of the literature identified 22 studies that met the inclusion criteria. These studies involved a total of 837 patients, 805 of whom had AIS with tandem occlusion (Table 1). Eleven studies reported use of the anterograde approach, 3 studies used the retrograde approach, 4 studies used both, and in 4 studies the approach was not specified. Seventeen studies were single- and multicenter retrospective reviews and 1 was a prospective study. Twenty studies reported the TICI revascularization grade as a measure of clinical outcome, with an average grade of 2b or 3 obtained in 79% of all cases for which TICI grade was reported (Table 1). The mRS was used as a primary outcome measure, and the majority of studies reported a mRS score of 2 or better in cases treated with endovascular therapy (Table 1).

Soize et al. reported a retrospective analysis of both tandem and solitary occlusions in AIS and demonstrated that tandem occlusions had a higher mortality rate at 3 months and more symptomatic intracranial hemorrhage at 24 hours. Duijsens et al. looked at tandem occlusions and further categorized them as true occlusions and pseudo-occlusions (those with low flow). The authors identified 16 tandem occlusions on CTA, and in 15 of these 16 cases, the tandem occlusions were subsequently confirmed on digital subtraction angiography (DSA). Of the patients with true occlusions (10 patients), 60% had TICI 2/3, 40% had TICI 0, and 40% had an average NIHSS score at discharge of 3. Of the patients with pseudo-occlusions (6 patients), 83% had a TICI 2/3 or 3 after recanalization, and the average NIHSS score at discharge was 2.

Throughout the literature, proximal revascularization consisted of stent placement and/or angioplasty for residual stenosis or passage of distal reperfusion devices. Distal revascularization was achieved with the MERCI retriever (Stryker), stent retrievers, Penumbra aspiration, balloon angioplasty and/or intracranial stent placement, and intra-arterial thrombolysis. The majority of those performing the anterograde approach placed the proximal stent without distal protection due to the presence of the distal occlusion and did not use proximal flow arrest. Some studies reported sporadic cases of spontaneous distal recanalization following the placement of a stent in the proximal carotid occlusion.

Only one study specifically advocated for the retrograde approach and included the specific cohort of carotid artery dissections in tandem occlusions. In this study, Marnat et al. found that only 25% of patients with dissection required cervical stent placement, and no early ipsilateral stroke occurrence was found in those who did not have the stent placed. The authors advocated for a retrograde approach where intracranial revascularization is established before evaluation of circle of Willis insufficiency and making the decision to place a stent. Although they compared onset to puncture time between those with tandem and solitary occlusions, time to reperfusion was not specifically reported.

Two studies compared the anterograde and retrograde approach. Stampfl et al. found that the mean time to reperfusion was 20 minutes longer in the anterograde approach. Lockau et al. had similar results, reporting a shorter median time from groin puncture to final reperfusion with the retrograde approach. Furthermore, for patients with tandem occlusions, Lockau et al. demonstrated good outcome (mRS score of 0–2) in 52.5% of the thrombectomy-first group versus 33.3% in the stent-first group. Otherwise, the studies reporting use of the anterograde approach reported similar revascularization scores (Table 1).

While some studies mentioned the predictors of outcome with solitary occlusions, only one specifically addressed this in the cohort of tandem occlusions. Grigoryan et al. performed a multivariate analysis and found that the strongest predictor of good outcome in patients with tandem occlusions was young age and low presenting NIHSS score. Reperfusion score of TICI 3 and high ASPECTS (Alberta Stroke Program Early CT Score) at presentation were also found to have a strong association with good clinical outcome. Time to reperfusion was notably not specifically evaluated as a predictor of clinical outcome in the review of the literature.

Illustrative Case
Presentation, Evaluation, and Initial Treatment
This 72-year-old man presented to an outside facility with acute onset of right hemiparesis and aphasia within 3 hours of symptom onset. He was treated with intravenous administration of tissue plasminogen activator (tPA, Alteplase) after noncontrast CT of the head demonstrated a dense left middle cerebral artery (MCA) sign and no acute
<table>
<thead>
<tr>
<th>Authors &amp; Year</th>
<th>Level of Evidence</th>
<th>No. of Pts w/ Tandem Occlusions</th>
<th>Treatment†</th>
<th>Description of Outcomes‡</th>
</tr>
</thead>
<tbody>
<tr>
<td>Malik et al., 2011</td>
<td>3, retrospective, single center</td>
<td>77</td>
<td>Anterograde</td>
<td>TICI ≥2 in 75.3%; mRS ≤2 in 41.6% at 3 mos</td>
</tr>
<tr>
<td>Machi et al., 2011</td>
<td>3, retrospective</td>
<td>10</td>
<td>Retrograde</td>
<td>TICI 2b or better in 8 of 10 pts; mRS 2 or better in 4</td>
</tr>
<tr>
<td>Kwak et al., 2013</td>
<td>3, retrospective</td>
<td>35</td>
<td>Anterograde</td>
<td>TICI 3, 54.3%; TICI 2b, 20%; TICI 2a, 25.7%; mRS 0-2 at 3 mos, 62.9%; median NIHSS 4 at 3 mos (range 1-17)</td>
</tr>
<tr>
<td>Dalyai et al., 2013</td>
<td>3, retrospective</td>
<td>17</td>
<td>Anterograde</td>
<td>94% recanalization;§ mean mRS 2.88 at discharge</td>
</tr>
<tr>
<td>Soize et al., 2014</td>
<td>3, prospective</td>
<td>11¶</td>
<td>Retrograde</td>
<td>Pts w/ tandem occlusions had poorer functional outcomes than those w/ solitary anterior circulation occlusions (18.2% vs 67.7%, p = 0.01) &amp; higher mortality rate at 3 mos (45.5% vs 12.9%, p = 0.03)</td>
</tr>
<tr>
<td>Tütüncü et al., 2014</td>
<td>3, retrospective, single center</td>
<td>30**</td>
<td>Not specified</td>
<td>TICI 2b/3 in 64%; major neurological improvement†† in 64%; mRS &lt;2 in 54%</td>
</tr>
<tr>
<td>Dababneh et al., 2014</td>
<td>3, retrospective, single center</td>
<td>7</td>
<td>Anterograde</td>
<td>TICI 2b/3 in 6 of 7 pts; NIHSS ≤1 at 90 days in 4 of 7; mRS 0 at 90 days in 4 of 7</td>
</tr>
<tr>
<td>Stampfli et al., 2014</td>
<td>3, retrospective, single center</td>
<td>24</td>
<td>Anterograde in 21, retrograde in 3</td>
<td>TICI 2b in 62.5%; median mRS 3.0 at 3 mos</td>
</tr>
<tr>
<td>Puri et al., 2015</td>
<td>3, retrospective</td>
<td>28</td>
<td>Anterograde</td>
<td>TICI 2a or better in 96.4%; mRS ≤ 2 in 52.5% at 3 mos</td>
</tr>
<tr>
<td>Duijsens et al., 2015</td>
<td>3, retrospective, single center</td>
<td>15‡‡</td>
<td>Anterograde</td>
<td>True occlusions: TICI 2/3 in 60%; pseudo-occlusions: TICI 2/3 in 83%; mean NIHSS 3 for true occlusions &amp; 2 for pseudo-occlusions</td>
</tr>
<tr>
<td>Spiotta et al., 2015</td>
<td>3, retrospective</td>
<td>16</td>
<td>Anterograde</td>
<td>TICI 2/3 in 100%; mRS ≤2 at 3 mos in 56.3%</td>
</tr>
<tr>
<td>Cohen et al., 2015</td>
<td>3, retrospective</td>
<td>24</td>
<td>Anterograde</td>
<td>TICI 2b in 79%; TICI 2a in 13%; TICI 1 in 8%; mRS ≤2 76% at 3 mos</td>
</tr>
<tr>
<td>Maurer et al., 2015</td>
<td>3, retrospective</td>
<td>43</td>
<td>Not specified</td>
<td>TICI 2b/3 in 76.7%; mRS ≤2 at discharge in 32.6%</td>
</tr>
<tr>
<td>Lockau et al., 2015</td>
<td>3, retrospective</td>
<td>37</td>
<td>Retrograde</td>
<td>TICI 2b/3 in 73% w/ anterograde Tx; mRS ≤2 at 3 mos in 8.3% of anterograde cases vs 16%, retrograde</td>
</tr>
<tr>
<td>Mishra et al., 2015</td>
<td>3, retrospective</td>
<td>7</td>
<td>Anterograde</td>
<td>TICI 2b in 100%; mean NIHSS 4.9 at discharge; mRS 1 at 3 mos in 100%</td>
</tr>
<tr>
<td>Heck &amp; Brown, 2015</td>
<td>3, retrospective, single center</td>
<td>23</td>
<td>Anterograde</td>
<td>TICI 2b/3 in 91%; mRS ≤2 at 3 mos in 52%</td>
</tr>
<tr>
<td>Lescher et al., 2015</td>
<td>3, retrospective</td>
<td>39</td>
<td>Not specified</td>
<td>TICI 2b in 64%; mRS ≤2 at 3 mos in 36%</td>
</tr>
<tr>
<td>Behme et al., 2015</td>
<td>3, retrospective, multicenter</td>
<td>170</td>
<td>Anterograde in 151 (89%)</td>
<td>TICI 2b in 77%; TICI 2a in 12%; mRS ≤ 2 at 3 mos in 36%</td>
</tr>
<tr>
<td>Weiner et al., 2017</td>
<td>4, case series, single center</td>
<td>2</td>
<td>Anterograde</td>
<td>Case 1: TICI 2b, mRS 1 at 3 mos; Case 2: TICI 2b, mRS at 6 mos</td>
</tr>
<tr>
<td>Marnat et al., 2016</td>
<td>3, retrospective, single center</td>
<td>20</td>
<td>Retrograde</td>
<td>TICI 2b/3 in 70%; mRS ≤2 at 3 mos in 70%</td>
</tr>
<tr>
<td>Grigoryan et al., 2016</td>
<td>3, retrospective, multicenter</td>
<td>100</td>
<td>Anterograde &amp; retrograde but not specified per cases</td>
<td>TICI 2b/3 in 88%; mRS ≤2 at 3 mos in 42%</td>
</tr>
<tr>
<td>Fahed et al., 2016</td>
<td>3, retrospective, single center</td>
<td>70</td>
<td>Not specified</td>
<td>mRS 2 or better in 50.8%; TICI 2b/3 in 67%</td>
</tr>
</tbody>
</table>

Ant = anterior; circ = circulation; EVT = endovascular therapy; Pts = patients; Tx = treatment.

* Of 837 patients reported on in the listed papers, a total of 805 patients had tandem occlusions.
† Approach: 11 anterograde, 3 retrograde, 4 both, 4 did not specify.
‡ Mean of 79% TICI 2b or better in studies that reported TICI grade.
§ No TICI scores reported.
¶ In addition to the 11 patients with tandem occlusions, the authors also reported on 31 patients with solitary anterior circulation occlusions and compared outcomes.
** Of 30 patients with tandem occlusions, 14 were treated with EVT and 16 were treated with IV thrombolysis.
†† Decrease in NIHSS by 8 points or NIHSS of 1/0 at discharge.
‡‡ Confirmed cases. One additional case was initially thought to be a tandem occlusion on presenting CTA and later seen as a solitary lesion on DSA. Of the original 16 cases, 10 involved true occlusions and 6 involved pseudo-occlusions.
hemorrhage. Following transfer of the patient to our facility, he had a National Institutes of Health Stroke Scale (NIHSS) score of 20 and modified Rankin Scale (mRS) score of 5.

CT angiography (CTA) demonstrated left cervical ICA occlusion with reconstitution at the supraclinoid segment and occlusion within the left M1 segment of the MCA. A CT perfusion study demonstrated a large area of perfusion mismatch within the left MCA territory as defined by decreased blood flow and increased blood volume (Fig. 1). After discussion with the family, the decision was made to proceed with endovascular revascularization.

Endovascular Technique

The patient was placed under monitored anesthesia care (MAC) sedation. The right femoral artery was accessed, a 5-Fr catheter was advanced into the left common carotid artery (CCA), and angiographic runs confirmed proximal cervical carotid artery occlusion. The sheath and diagnostic catheter were exchanged for a 9-Fr sheath and a 9-Fr concentric balloon guide catheter, which was advanced into the left common carotid artery (CCA). A triaxial system was used with a 5MAX ACE reperfusion catheter (Penumbra), Marksman microcatheter (ev3/Covidien Vascular Therapies), and Glidewire Gold 0.016-inch–diameter microwire (Terumo) and navigated slowly and progressively into the occluded lumen of the ICA. The concentric balloon catheter was not inflated at this point. Serial arteriograms were obtained to ensure that the endovascular devices remained within the intravascular lumen. Contrast injection at the level of the petrous ICA revealed filling of the anterior cerebral artery (ACA) territory with distal thrombus and a left M1 occlusion (Fig. 2). The microcatheter and microwire were navigated distal to the M1 occlusion, and the Penumbra 5MAX ACE catheter was advanced over the microcatheter and brought to the clot interface. A Solitaire FR device (4 × 20 mm; Covidien, ev3 Neurovascular) was deployed and a Penumbra aspiration system was applied to the 5MAX ACE catheter. When it was seen that there was no flow under aspiration, the catheter was removed under suction. Distal access was obtained again, and subsequent angiography revealed reperfusion of the MCA territory with a revascularization score of TICI 2b with delayed filling in distal M1 branches (Fig. 3).
To localize the distal extension of the ICA occlusion, the microcatheter was slowly and progressively withdrawn into the distal ICA with serial microinjections of contrast (Fig. 4A–C). Once the extent of the ICA occlusion was identified, we attempted thromboaspiration with the 5 MAX ACE, but the attempt was not successful. The Traxcess microwire (MicroVention) extension was added to maintain access, and the microcatheter was exchanged with an Aviator plus balloon dilatation catheter (4 × 20 mm, Cordis) for angioplasty of the area of focal stenosis under proximal flow arrest to prevent distal embolization. Next, a carotid Wallstent endoprosthesis (8 × 21 mm, Boston Scientific Corp.) was deployed across the area with abnormal lumen. Postdilatation angioplasty was performed for approximately 50% residual stenosis.

The patient’s blood pressure was in the 200 mm Hg range and left untreated until this point to help with collateral circulation before recanalization. Members of the anesthesia department administered intravenous (IV) antihypertensive treatment at this time to lower the systolic blood pressure. To reduce the risk of hemorrhagic conversion after recanalization, several aspirations were performed to remove debris, and once the systolic blood pressure was less than 160 mm Hg, the concentric proximal balloon was deflated. A subsequent cervical angiogram showed revascularization of the cervical ICA (Fig. 4D and E) and a cerebral angiogram showed thrombosis in the distal A3 segment (Fig. 5A and B). The triaxial system was navigated into the A2 segment just proximal to the site of occlusion. The Marksman microcatheter was exchanged for a 3MAX reperfusion catheter (Penumbra). The Neuron 3MAX separator (Penumbra) was used for thrombus manipulation and a total of 4 mg of tissue plasminogen activator (tPA) was administered intrarterially. Final control cerebral angiogram showed recanalization of the A2 segment from a TICI grade of 0 to a TICI grade of 3 and maintained patency of the MCA territory with a TICI grade of 2b (Fig. 5C–E).

An 8-Fr Angio-Seal (Terumo Interventional Systems) was applied to the groin for closure. The patient was given aspirin 325 mg during the procedure and intravenous heparin was administered to maintain an activated clotting time (ACT) greater than 250 seconds.

Postoperative Course

CT showed hyperdensity within the left basal ganglia, internal capsule, thalamus, and areas in the left MCA re-
gion in the posterior frontal and parietal cortex (Fig. 6A). The patient was maintained on aspirin at 325 mg and IV heparin over 24 hours with an activated partial thromboplastin time (aPTT) goal of 40–50 seconds. Subsequent CT scans demonstrated clearance of hyperdensity that appeared to be contrast staining, and treatment with clopidogrel was initiated. The IV heparin was discontinued once the clopidogrel treatment was started, and the patient was maintained on clopidogrel and aspirin (Fig. 6B and C).

The immediate follow-up NIHSS score was 17. Carotid artery Doppler ultrasonography confirmed stent patency, and the patient was discharged to rehabilitation on Day 4 with an NIHSS score of 7 and an mRS score of 1. At 30-day follow-up, the patient still had an mRS score of 1 with only minimal residual aphasia.

Discussion

As techniques for endovascular revascularization evolve, there is uncertainty as to how to approach tandem occlusions in AIS. The literature suggests that tandem occlusions are associated with worse outcome than solitary intracranial occlusions. Historically, patients with tandem occlusions have had a mortality rate ranging between 24% and 27% and a good clinical outcome measured by an mRS of 0–2 in only 10%–29% of cases. Intravenous thrombolysis in these patients has not demonstrated a significant difference in recanalization rates or clinical outcome in comparison with placebo. More recently, despite the recognized poor clinical outcome, it has been suggested through systematic reviews that these patients may fare better with endovascular management. Although we have used both approaches at our center, we are not certain about the best practice in similar cases, and the reported literature on this topic has no established recommendations.

Furthermore, the underlying pathology is variable, and proximal occlusions can be related to a wide spectrum of disorders, such as dissections, atherosclerotic occlusions, embolic events, or even occlusions in patients who have previously undergone carotid artery stent placement. In a review of recent stroke studies in the American Heart Association/American Stroke Association guidelines update, Powers et al. report that 30 of the 75 patients with carotid artery stenosis or occlusion in MR CLEAN were treated with stent placement during the acute intervention. Nine of the 19 patients with carotid occlusion in REVASCATS had stents placed at the time of thrombectomy. However,
it was not made clear what factors were considered regarding the decision to place a stent and what approach was used (anterograde versus retrograde revascularization) to recanalize the proximal and distal occlusions.

We noted that the majority of published approaches involved anterograde revascularization of the proximal carotid occlusion with angioplasty and/or stent placement followed by distal revascularization of the intracranial le-

FIG. 5. Revascularization of ACA. A: Left ICA injection (lateral view) angiogram obtained after placement of stent demonstrating distal A3 segment thrombosis (white arrow). B: Microcatheter angiogram demonstrating thrombus at A3 segment bifurcation and occlusion of distal ACA branches (white arrow). C: Left ICA injection (lateral view) angiogram obtained after microwire thrombus manipulation and intra-arterial tPA treatment showing TICI 3 revascularization of the ACA territory. D and E: Left ICA injection final control angiograms (lateral [D] and AP [E] views) showing TICI 3 revascularization of the ACA territory and TICI 2b revascularization of the MCA territory.

FIG. 6. Postintervention axial noncontrast CT image. A: Image from scan performed immediately after intervention showing hyperdensity in the putamen, globus pallidus (white arrow), internal capsule, and thalamus. Areas of hyperdensity were also seen in regions of the posterior frontal cortex and parietal cortex supplied by the left MCA (not shown here). B and C: Images from scans performed 6 hours (B) and 24 hours (C) after intervention showing resolution of likely contrast staining in the area of the left basal ganglia.
sion (Table 1). The cited advantages included the ability to jail the likely atheromatous friable plaque against the vessel wall proximally with angioplasty and/or stent placement, preventing showering of emboli distally. Additionally, the recanalization of the proximal occlusion results in anterograde blood flow and is thought to assist in thrombolysis for the distal lesion.

Previously published anterograde approaches involved an assessment of the intracranial circulation for improved flow after revascularization of the proximal occlusion. The time required for stent placement and/or angioplasty in combination with the time to assess for distal occlusions could certainly delay intracranial reperfusion that could be obtained from distal collateral circulation across the circle of Willis and may contribute to neurological morbidity and worse functional outcome. Moreover, proximal revascularization can result in embolism of thrombus distally to a branch that was not occluded, highlighting the importance of using proximal protection in these cases. Decreasing “door to needle time” and time to reperfusion represents a driving force in establishing stroke management guidelines. A recent meta-analysis looked at the relationship between time to treatment and endovascular intervention, and time to treatment was defined as the time from symptom onset to time of reperfusion (TICI 2b or 3). The rate of functional independence was 64% with reperfusion at 3 hours and 46% with reperfusion at 8 hours.

As in our illustrative case, and in cases with proximal and distal tandem occlusions with patency of the carotid terminus and proximal intracranial segments on CTA, we prefer to perform a distal to proximal approach for revascularization. In this specific subset of patients, we believe that revascularization could be obtained faster from distal collaterals, and this might be associated with reduced time to recanalization and possibly a better outcome. As previously mentioned, Lockau et al. reported a higher rate of good outcome for patients with tandem occlusions when distal thrombectomy was performed first rather than stent placement first (52.5% vs 33.3%). However, their study was retrospective, with inherent selection bias.

Although future studies can be designed to compare the time to reperfusion for these 2 approaches, technical nuances should be established, and care should be individualized based on the specifics of the underlying pathology. For example, occlusions that involve the carotid terminus and extend proximally with no underlying pathology of the cervical ICA might be better addressed with proximal flow arrest regardless of technique for clot retrieval to prevent distal embolization. Even large suction catheters are not guaranteed to remove the clot without leaving pieces that can be displaced after restoration of flow. Clots can be moved distally to more important branches, and stroke cases can be converted from ischemic penumbra of non-eloquent brain to infarct of highly functional and eloquent brain with irreversible damage.

Additionally, technical considerations exist along with significant differences in these approaches. When deploying a stent in the proximal ICA first, there is an advantage to establishing a true lumen for navigation of reperfusion devices distally. However, there is the possibility of challenges for stent deployment, and some stents with open-cell design can compromise the ability to advance the reperfusion devices distally. Stent retrievers can also get caught on the stent if not retrieved distally. All of this can potentially prolong the time to distal revascularization and might increase the morbidity of these challenging procedures. In our center, we try to avoid the use of stents in the carotid artery unless there is a dissection flap that could not be maintained open or a clot that could not be aspirated. We attempt balloon angioplasty first to allow for advancement of a large suction catheter with the microcatheter and microwire. In our practice we use proximal protection with flow arrest at the ICA if possible to reduce the risk of distal embolization. The balloon is inflated in the CCA in occlusions at the bifurcation, and this strategy may still allow for retrograde flow through the external carotid artery.

Hemorrhagic conversion is also a recognized potential complication of acute stroke treatment, and the risk is magnified in the setting of stent and/or angioplasty requiring dual antiplatelet therapy regardless of anterograde or retrograde approach. For the studies evaluated in this review, the rate of symptomatic intracranial hemorrhage among patients treated for tandem occlusions ranged from 0%–18% but was not specifically evaluated as a predictor of clinical outcome. This risk can be minimized by ensuring a normotensive blood pressure before restoration of flow and following the procedure. As mentioned above, we try to avoid stent placement whenever possible, and the decision of whether or not to use a stent is delayed until after attempts of revascularization of distal occlusions. When stent deployment is necessary, we maintain patients on aspirin and intravenous heparin with low PTT goal (40–60) and perform serial CT imaging to rule out acute hemorrhagic conversion before administering a loading dose of clopidogrel.

Currently, the combination of angioplasty and stent placement for proximal stenosis or occlusion is not considered useful in the most recent American Heart Association guidelines on management of patients with AIS (Class III: Level of Evidence C). The use of carotid artery stents placement in acute stroke is considered an “off-label” application and is not covered by the Centers for Medicare and Medicaid Services (CMS) guidelines in regard to national indications for coverage in percutaneous transluminal angioplasty (PTA). Specifically, patients who have had a disabling stroke (mRS score ≥ 3) shall be excluded from coverage. We have not used carotid artery stent placement routinely in the acute setting, and we usually avoid using stents in this setting given the need for dual anticoagulation and risk of hemorrhagic conversion. However, we have found angioplasty helpful in patients with severe underlying ICA stenosis that likely progressed and caused the occlusion. In patients with a dissection flap in whom a patent ICA cannot be maintained or in those with a large thrombus that we cannot aspirate, we use a stent as a method of maintaining patency of the pathological ICA segment, as the stent pushes the dissecting flap or the clot outside and helps reestablish flow within the lumen.

This study has several limitations. The majority of studies included in the literature review represent Level
3 evidence with heterogeneous and nonstandardized description of the specific endovascular technique of revascularization. Together with the illustrative case, this review calls for the need to better individualize techniques to specific cases of tandem occlusions and perhaps establish recommendations based on clinical outcome.

Conclusions

Distal to proximal revascularization of tandem occlusions is worth a special consideration, especially in patients with patent collaterals through the circle of Willis. Compared with the anterograde approach, this technique may result in better clinical outcomes, especially with evidence that time to reperfusion is a significant predictor of functional outcome. Technical variations should be individualized to the specific distribution and configuration of the occlusions, as tandem occlusions have not been analyzed systematically in the most recent stroke trials. Future studies involving the endovascular management of these patients may be able to further elucidate the predictors of good clinical outcome in this group.

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Disclosures

Dr. Tawk reports direct stock ownership in Medtronic.

Author Contributions

Acquisition of data: Tawk, Brown, Miller. Analysis and interpretation of data: Mbabuike, Gassie. Drafting the article: Mbabuike. Critically revising the article: Tawk. Reviewed submitted version of manuscript: Tawk. Approved the final version of the manuscript on behalf of all authors: Tawk. Administrative/technical/material support: Gassie, Brown, Miller. Study supervision: Tawk, Brown, Miller.

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