

RADIOLOGICAL METHODS AND METHODOLOGIES FOR ASSESSING COLLATERAL BLOOD CIRCULATION OF THE BRAIN

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Collateral circulation is assigned a **decisive role** in the final outcome of ischemic stroke. Accordingly, the refinement of various methods and methodologies for assessing this system is becoming increasingly relevant. Currently, there are numerous **radiological methods** that allow us to assess not only the anatomy of collateral circulation but also to gain an understanding of its quantitative characteristics. Furthermore, the modern radiological literature publishes the latest studies comparing various methodologies for the **qualitative and quantitative assessment** of collateral circulation to determine which of them is more accurate and yielding better results.

აბსტრაქტი

ტვინის კოლატერალური სისხლის მიმოქცევის შეფასების რადიოლოგიური
მეთოდები და მეთოდოლოგიები

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კოლატერალურ სისხლის მიმოქცევას ენიჭება გადამწყვეტი როლი იშემიური ინსულტის საბოლოო გამოსავალში. შესაბამისად დღითიდღე აქტუალური ხდება სისტემის შეფასების სხვადასხვა მეთოდების და მეთოდოლოგიების დახვეწა. დღესდღეობით არსებობს მრავალი რადიოლოგიური მეთოდი, რომლებიც საშუალება სგვაძლევენ შევაფასოთ არა მხოლოდ კოლატერალური სისხლის მიმოქცევის

ანატომია, არამედ წარმოდგენა შევიქმნათ მის რადენობრივ მახასიათებლებზეც. ასევე, თანამედროვე რადიოლოგიური ლიტერატურაში ქვეყნდება უახლესი კვლევები, რომლის საშუალებითაც ერთმანეთს დარდება კოლატერალური სისხლის მიმოქცევის თვისოვრივი და რადენობრივი შეფასების სხვადასხვა მეთოდოლოგიები, რათა დადგინდეს, თუ რომელი მათგანია უფრო ზუსტი და შედეგის მომცემი.

Key words: Cerebral infarction, Collateral circulation, Radiological methods of collateral assessment, Normal anatomy of collaterals.

Well-perfused blood flow via collaterals may **increase the benefit of endovascular treatment** in acute ischemic stroke and reduce the risk of hemorrhagic transformation. It also significantly **reduces the risk of recurrent stroke** in patients with symptomatic intracranial atherosclerotic stenosis and **decreases the volume of the infarct core** during ischemic stroke.

Accurate assessment of the **anatomy and function of cerebral collateral circulation** is an important prerequisite for the individualized management of stroke patients. Currently, the assessment and intervention of collateral circulation in ischemic stroke is a subject of **active research**. Various imaging criteria have been developed to assess collateral status and correlate with prognosis in stroke patients. There are also interventions aimed at **enhancing collateral circulation** in stroke patients.

Abbreviations

Abbreviation	Full Name
DSA	Digital Subtraction Angiography
TCD	Transcranial Doppler
TOF-MRA	Time-of-Flight Magnetic Resonance Angiography

Abbreviation	Full Name
CTA	Computed Tomography Angiography
SPECT	Single-Photon Emission Computed Tomography
PET	Positron Emission Tomography
CTP	Computed Tomography Perfusion
QMRA	Quantitative Magnetic Resonance Angiography
ASL	Arterial Spin Labelling
MCA	Middle Cerebral Artery
ACA	Anterior Cerebral Artery
PCA	Posterior Cerebral Artery
ASITN/SIR	American Society of Interventional and Therapeutic Neuroradiology / Society of Interventional Radiology

Overview of Assessment Methods

Currently, many radiological methods exist for assessing collateral circulation (Table 1).

The **gold standard** for assessment is **Digital Subtraction Angiography (DSA)**. However, due to the invasiveness and high cost of the procedure, it is not frequently used (Fig. 1) [3].

Table 1: Various Radiological Methods for Assessing Collateral Circulation

1. Transcranial Doppler (TCD)
2. Transcranial Color Duplex Scanning
3. Routine, Single-Phase Computed Tomography (CT)
4. Computed Tomography Angiography (CTA)
5. Computed Tomography Perfusion (CTP)
6. Non-contrast MRI Angiography (TOF-MRA)
7. Contrast-enhanced MRI Angiography
8. Digital Subtraction Angiography (DSA)
9. Xenon Computed Tomography (Xenon-CT)
10. Single-Photon Emission Computed Tomography (SPECT)
11. Positron Emission Tomography (PET)
12. MRI Perfusion (MRP)

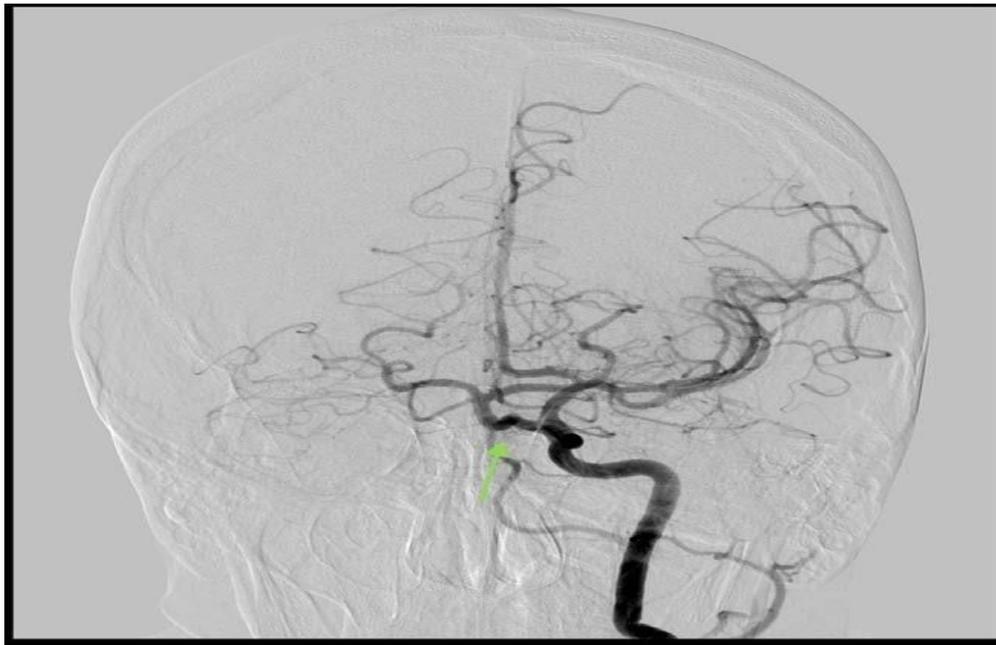


FIG. 1. DIGITAL SUBSTRACTION ANGIOGRAPHY(DSA)

TCD, as a non-invasive examination method, reflects the cerebral blood flow velocity, collateral status, and cerebrovascular reactivity in real-time. It is characterized by low cost, but the accuracy of TCD in diagnosing cerebrovascular anomalies is highly dependent on the operator's (radiologist's) experience. Collateral flow via the anterior communicating artery, posterior communicating artery, ophthalmic artery, and leptomenigeal arteries can be detected directly or indirectly by TCD. According to some studies, the sensitivity and specificity of TCD in detecting collateral flow via the anterior communicating artery and basilar artery were 95% and 87%, and 100% and 95%, respectively, compared to the gold standard, which was considered DSA.

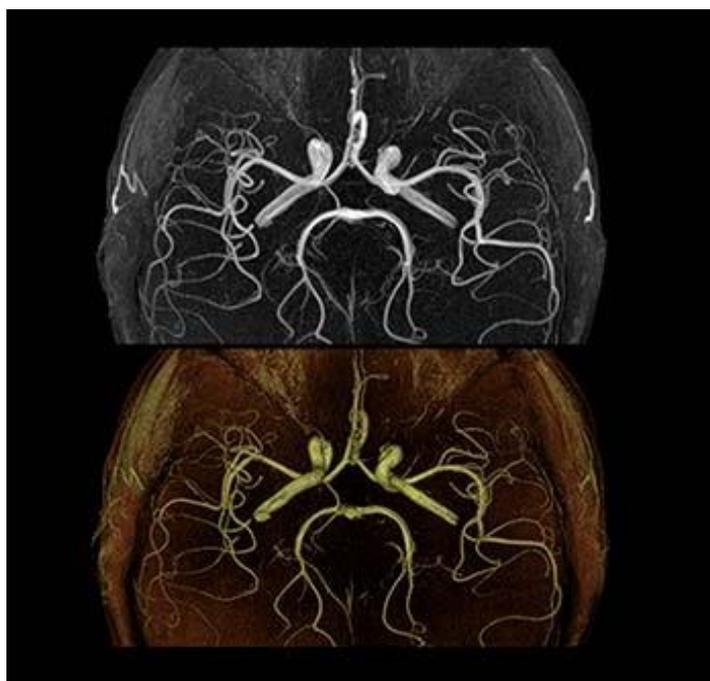


FIG. 2. NON CONTRAST MRI ANGIOGRAPHY, TOF (TIME-OF-FLIGHT)

TCD, as a non-invasive examination method, reflects the cerebral blood flow velocity, collateral status, and cerebrovascular reactivity in real-time. It is characterized by low cost, but the accuracy of TCD in diagnosing cerebrovascular anomalies is highly dependent on the operator's (radiologist's) experience. Collateral flow via the anterior communicating artery, posterior communicating artery, ophthalmic artery, and leptomeningeal arteries can be detected directly or indirectly by TCD. According to some studies, the sensitivity and specificity of TCD in detecting collateral flow via the anterior communicating artery and basilar artery were 95% and 87%, and 100% and 95%, respectively, compared to the gold standard, which was considered DSA.

TOF-MRA (Non-contrast MRI Angiography) is another non-invasive method commonly used to assess the structure of cerebral collateral circulation. The reliability of TOF-MRA for assessing leptomeningeal collaterals is limited due to its relatively low spatial resolution. TOF-MRA is typically used to assess **first-order collaterals** in the Circle of Willis. In various studies, compared to DSA as the gold standard, the sensitivity and specificity of TOF-MRA for detecting collateral flow via the anterior circulation of the Circle of Willis were 83% and 77%, and for the posterior part of the Circle of Willis, 33% and 88%, respectively (Fig. 2) [4].

CTA is also a non-invasive method that has high accuracy in assessing the patency of arterial segments in the Circle of Willis, showing >90% concordance with DSA, but its sensitivity (53%) is limited for imaging hypoplastic arterial segments. Blood flow via collaterals may be delayed compared to normal antegrade flow. Thus, traditional single-phase CTA may underestimate the compensatory collateral flow. Currently, **Multi-Phase CTA** (or Dynamic CTA, or Four-Dimensional CTA) is increasingly used in clinical studies to assess cerebral collateral status [5].

Various visualization methods exist to assess the function of cerebral collateral circulation, such as cerebrovascular reserve using **TCD**, **Xenon-CT**, **SPECT**, **PET**, **CTP**, **QMRA**, traditional dynamic susceptibility contrast-enhanced **MR Perfusion (MRP)**, and **Arterial Spin Labeling (ASL)**. These visualization methods typically assess the direction/velocity/volume of cerebral blood flow or the perfusion status. Consequently, they accurately reflect the compensatory function of collateral blood flow. Some modern visualization methods can simultaneously detect both the structure and function of collateral circulation; for example, QMRA can reveal the direction of blood flow via collateral channels and quantitatively determine the total/regional cerebral blood flow.

Currently, there are many different methodological scales for assessing collateral circulation, which are adapted according to various radiological examination methods (Table 2).

CT	<ol style="list-style-type: none"> 1. Miteff collateral grading method 2. Maas collateral grading method 3. Tan collateral grading system 4. The Alberta Stroke Program Early CT Score 20-point methodology 5. Regional leptomeningeal collateral (rLMC) score 6. ACA-MCA and PCA-MCA regional collateral score
MRI	<ol style="list-style-type: none"> 1. PCCS 2. Good CS and Poor CS 3. ASITN/SIR 4. 3 point scale by Tan 5. Present vs absent ACOMA 6. Presence or absence of FVH-DWI mismatch 7. FHV-ASPECTS 8. 5-point scale 9. TMACS 10. Flow map collateral grade based on ASITN/SIR 11. Maas scoring
DSA	<ol style="list-style-type: none"> 1. The ASITN/SIR collateral scale 2. Christoforidis collateral grading system

CTA-Based Grading Systems

With the **Miteff collateral grading method** (Fig. 3), collateral status is assessed on maximum intensity projection reconstructions in axial, coronal, and sagittal planes of single-phase CTA in patients with **MCA occlusion** and is graded as: **Good** – if the main MCA branches are reconstituted distal to the occlusion, **Moderate** – if some MCA branches are visualized in the Sylvian fissure, **Poor** – if only distal superficial MCA branches are reconstituted.

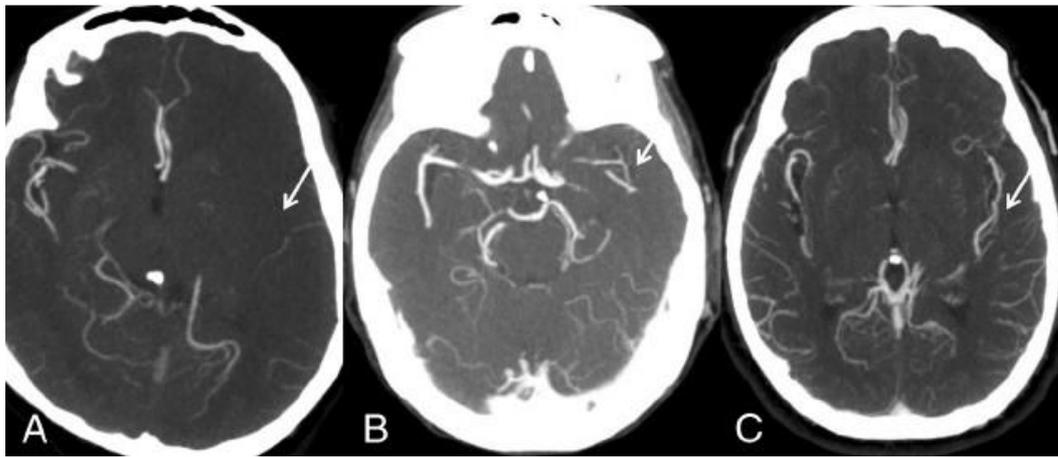
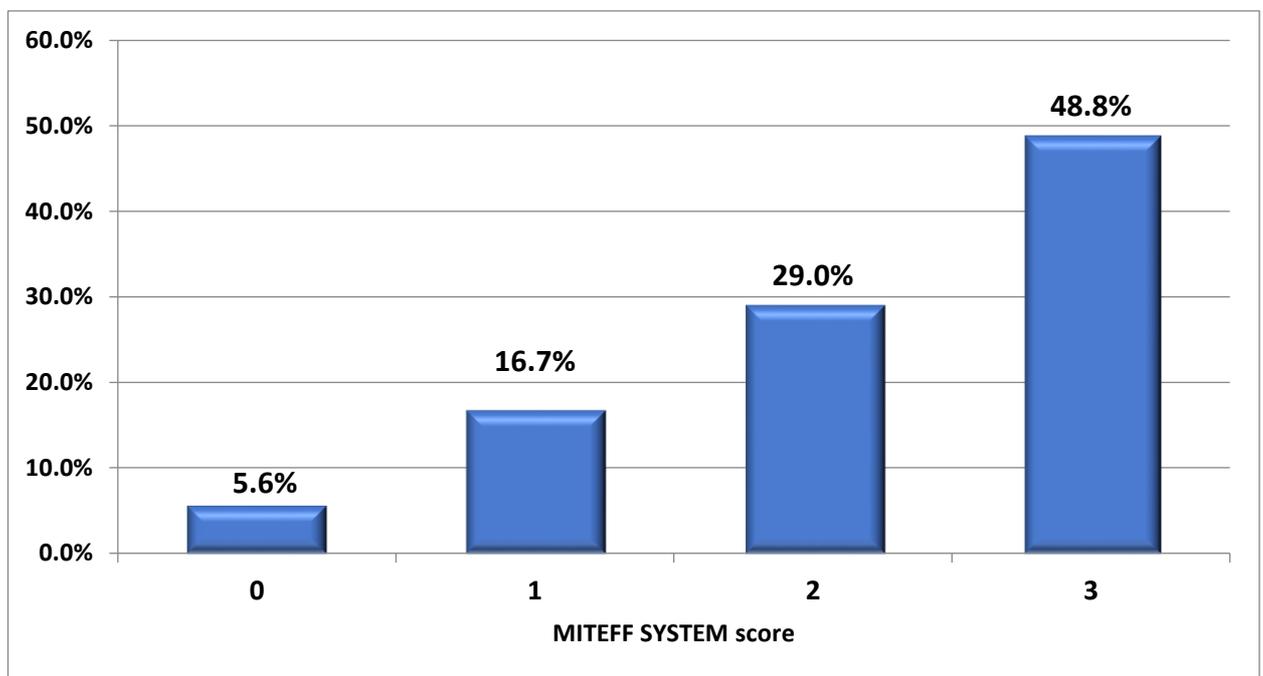
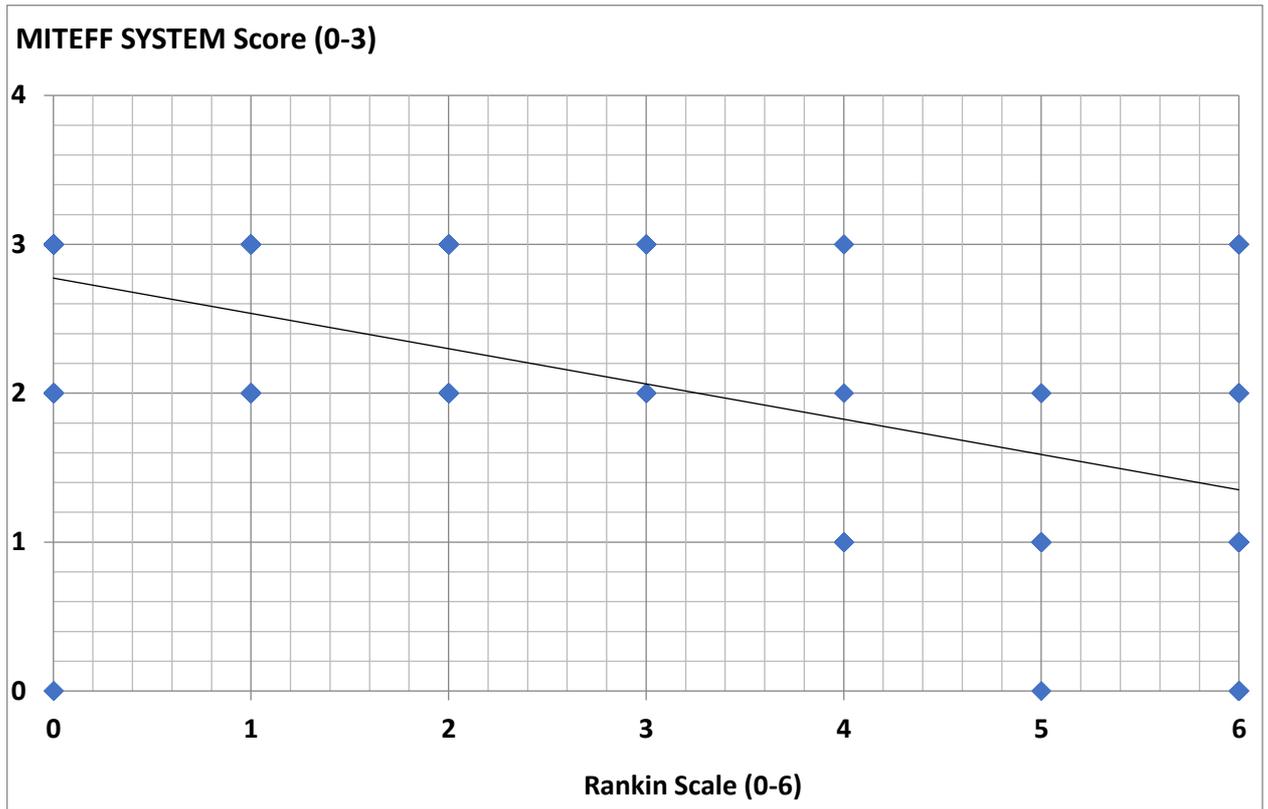


FIG. 3. Miteff System A – contrast in distal branches. B – vessels are visible in the sylvian fissure C – vessels are recanalized distal to the occlusion.



MITEFF SYSTEM	N, %	Chi2; p
0	9; 5.6%	66.64; p<0.001
1	27; 16.7%	
2	47; 29.0%	
3	79; 48.8%	



$r = - 0.585, p < 0.001$

SYSTEM	Poor outcome	Moderate outcome	Good outcome
MITEFF SYSTEM score – 0-1	34 (94.4%)	0 (0.0%)	2 (5.6%)
MITEFF SYSTEM score – 2-3	17 (13.5%)	33 (26.2%)	76 (60.3%)
OR (95%CI), p	109.0 (24.0-495.9), p<0.001	N/A	25.8 (5.9-112.4), p<0.001

With the **Maas collateral grading method** (Fig. 4), the leptomeningeal vessels in the Sylvian fissure are assessed on CTA images and compared to the contralateral side: **1** – None, **2** – Less than the contralateral normal side, **3** – Equal to the contralateral normal side, **4** – More than the contralateral normal side, **5** – Hyper-abundant. The presence and condition of the anterior and

posterior communicating arteries are assessed as follows: **1** – Absent, **2** – Probably present, **3** – Very faint, **4** – Definitely present, **5** – Strong.

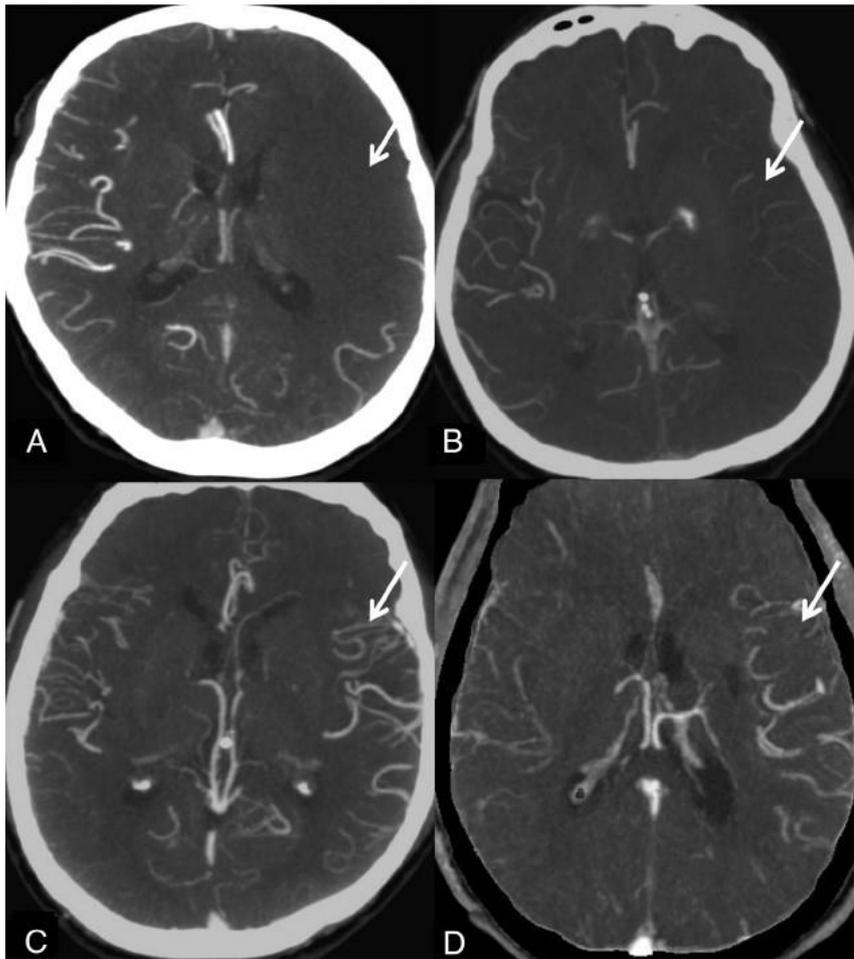
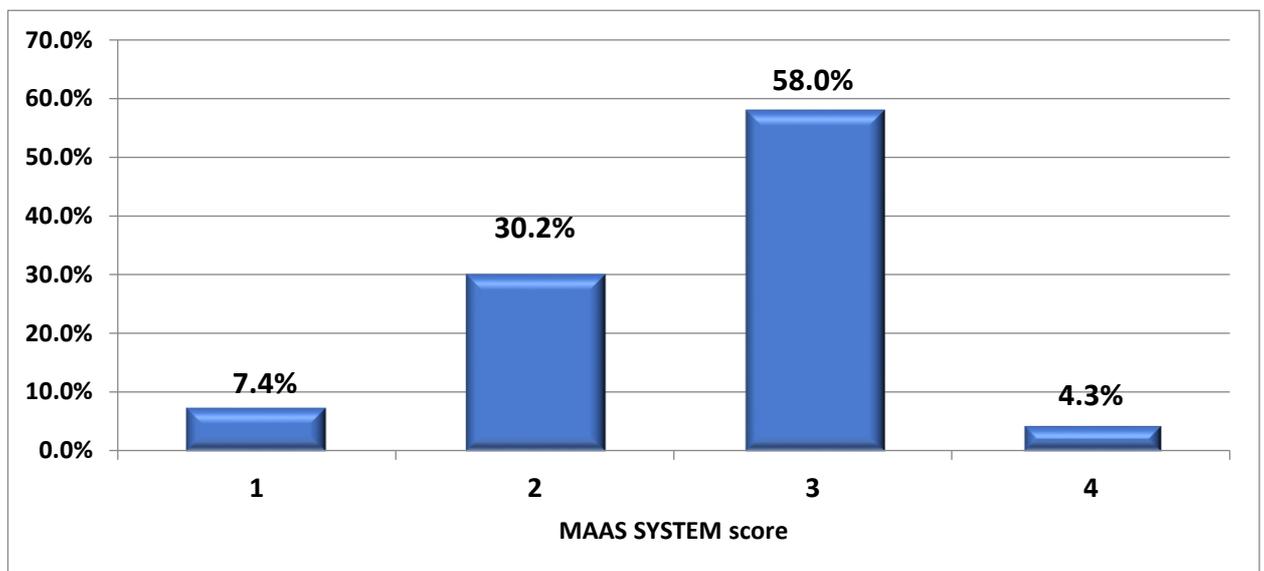
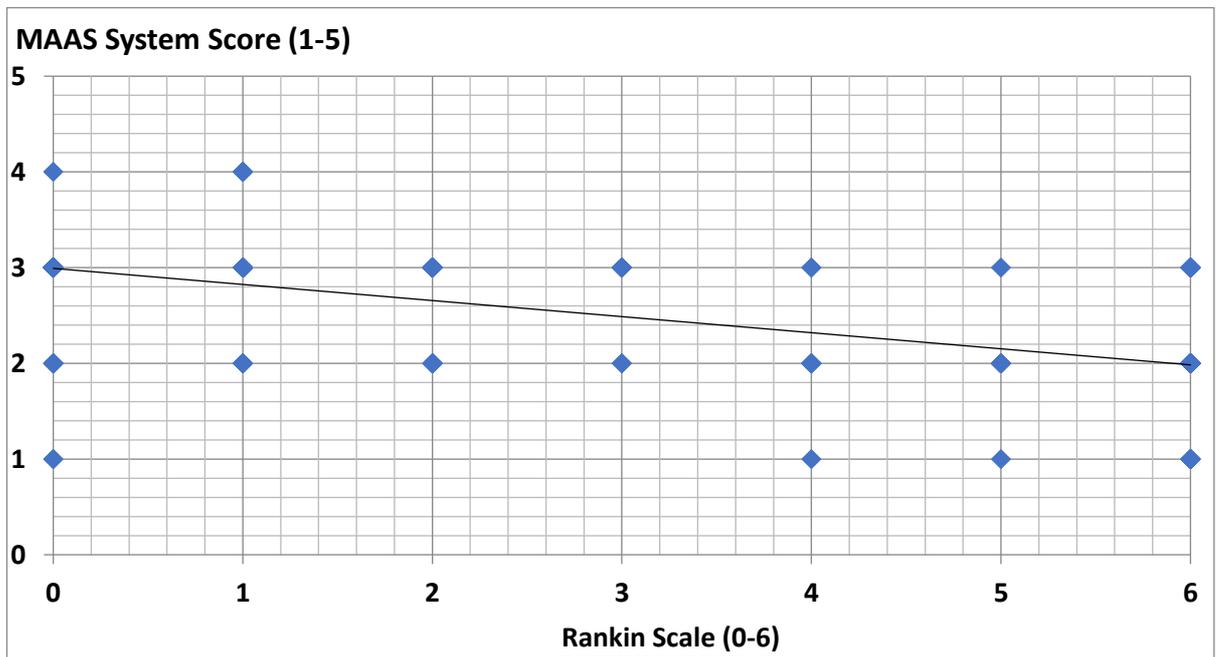


FIG. 4. Maas system. A – no vessels, B – less than contralateral side, C – more than contralateral side, D – Hyper-abundant



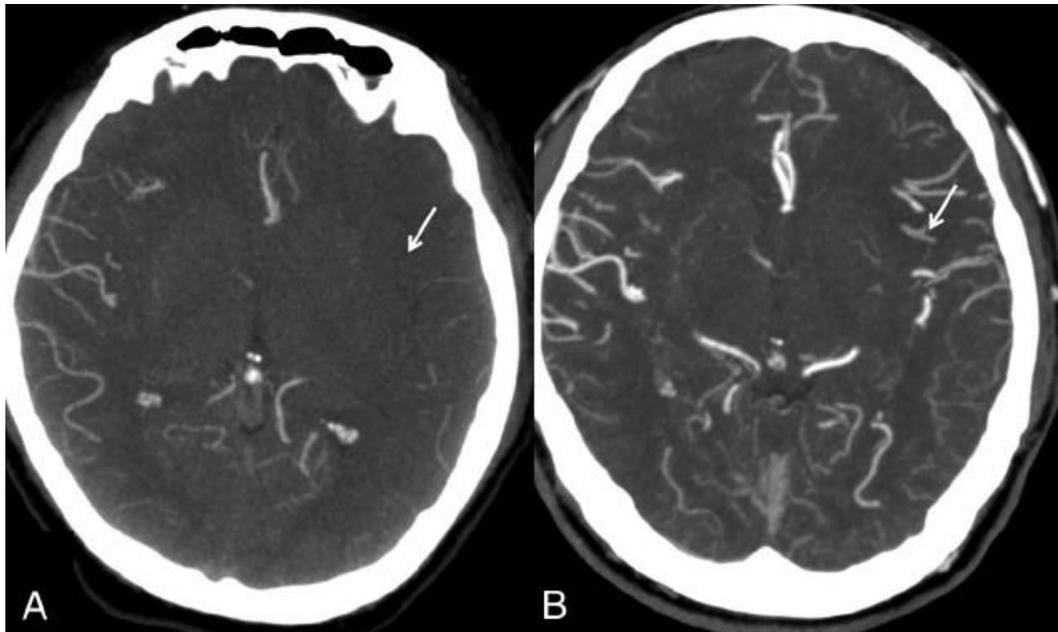
MAAS SYSTEM	N, %	Chi2, p
1	12; 7.4%	120.22; p<0.001
2	49; 30.2%	
3	94; 58.0%	
4	7; 4.3%	



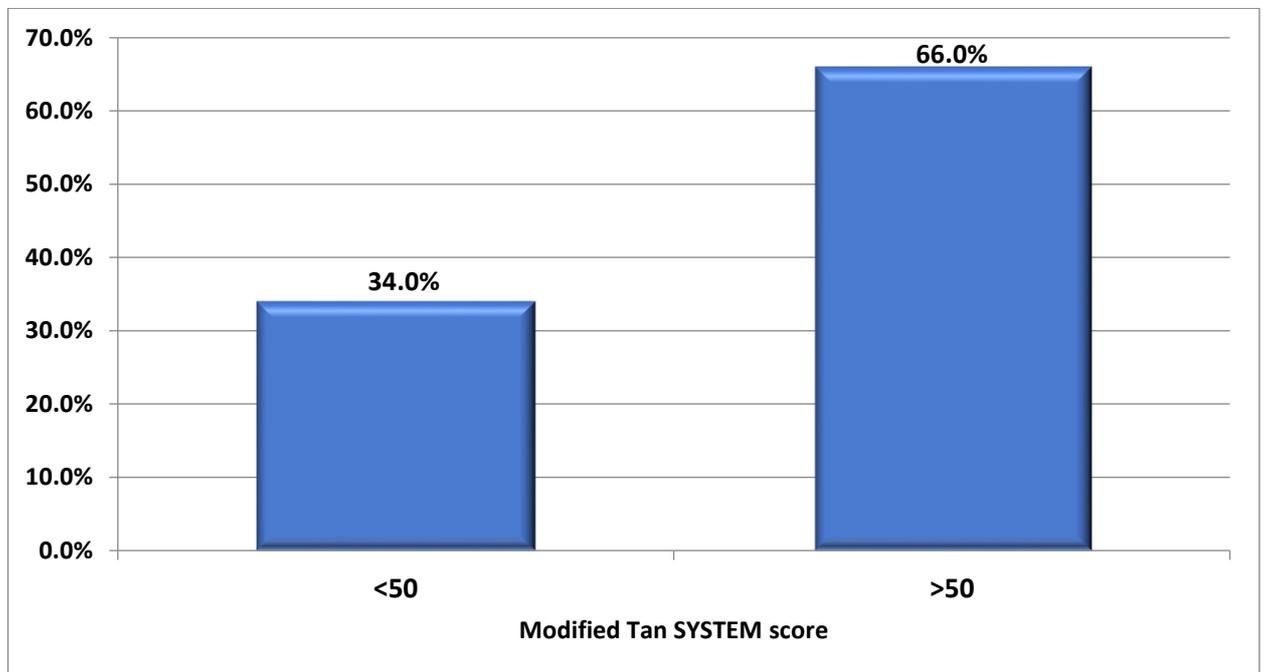
r = - 0.549, p<0.001

SYSTEM	Poor outcome	Moderate outcome	Good outcome
MAAS SYSTEM score – 1-2	41 (67.2%) 12 (19.7%)	8 (13.1%)	12 (19.7%)
MAAS SYSTEM score – 3-5	10 (9.9%)	25 (24.8%)	66 (65.3%)
OR (95%CI), p	7.7 (3.6-16.3), p<0.001	2.2 (0.9-5.2), p=0.079	18.6 (8.0-43.4), p<0.001

The **Modified Tan Collateral Grading System** (Fig. 5) assesses leptomeningeal collateral status on source CTA images in maximum intensity projection reconstructions, good if there is more than 50% of MCA territory and poor if there is less than 50% MCA territory.



sur. 5. **Modified Tan Collateral Grading System** A – MCA less than 50% MCA territory, B – more than 50% of MCA territory

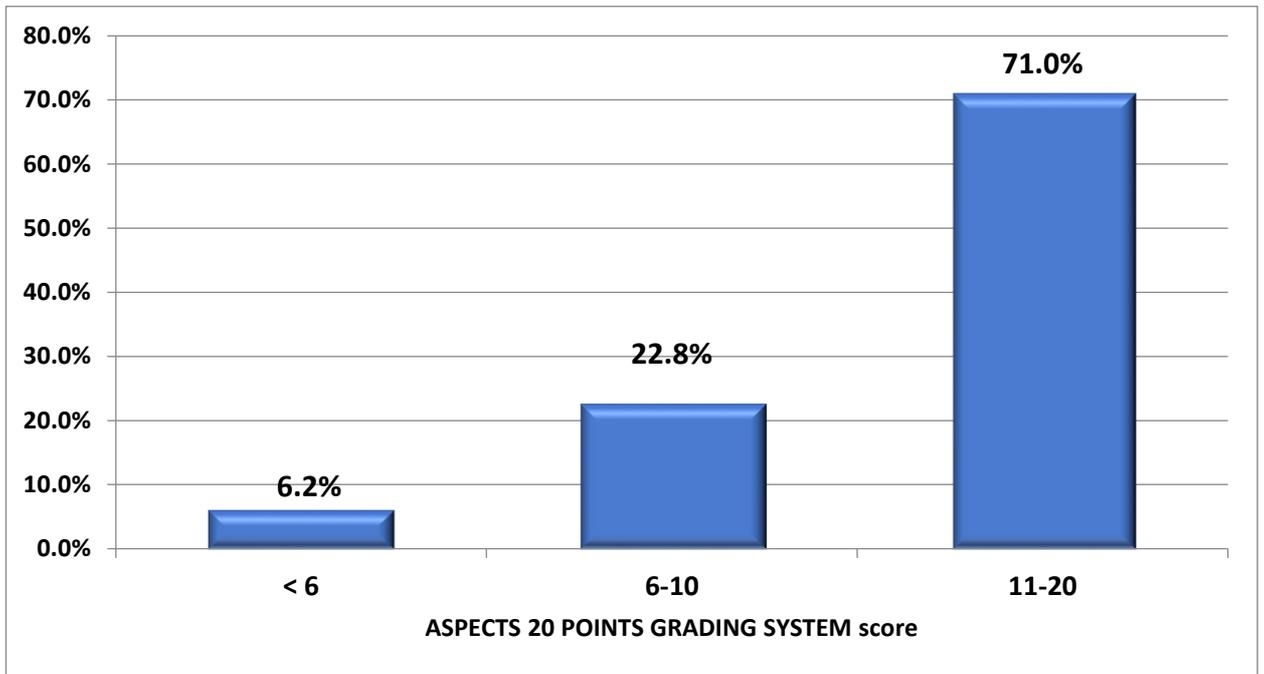


Modified Tan Scale	N, %	Chi2-, p
< 50	55; 34.0%	16.69; p<0.001

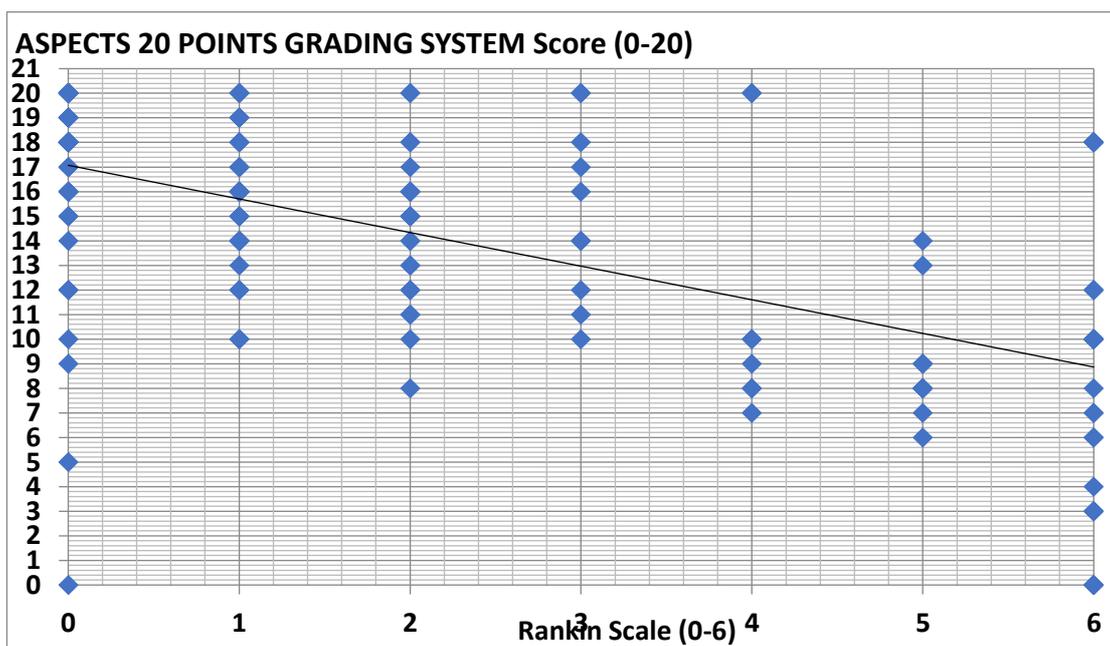
> 50	107; 66.0%		
SYSTEM	Poor outcome	Moderate outcome	Good outcome
Modified Tan SYSTEM score - <50	37 (67.3%)	10 (18.2%)	37 (67.3%)8 (14.5%)
Modified Tan SYSTEM score - >50	14 (13.1%)	23 (21.5%)	14 (13.1%)70 (65.4%)
OR (95%CI), p	11.1 (4.8-26.0), p<0.001	1.2 (0.5-2.8), p=0.620	13.7 (6.2-30.3), p<0.001

The **Regional Leptomeningeal Collateral (rLMC) Score** (20-point scale) compares the degree of contrast opacification in arteries distal to an **M1 MCA occlusion** (internal carotid artery occlusion) in the symptomatic hemisphere to the contralateral hemisphere, using reformatted CTA reconstructions. A higher score indicates better collateral status. The degree of contrast opacification is graded as **0** (artery not visible), **1** (less pronounced), or **2** (equal to or more pronounced than the opposite hemisphere) for six **ASPECTS** cortical regions. the parasagittal **ACA** territory, and the basal ganglia. The areas in the Sylvian fissure are assigned a higher score of **0, 2, or 4**.

The **ACA-MCA and PCA-MCA Regional Collateral Score** assesses the extent and visibility of pial arteries in the **ACA-MCA and PCA-MCA regions** in stroke patients with **M1 MCA occlusion** (intracranial internal carotid artery occlusion), using multi-phase reconstructions of Dynamic CTA. Collaterals in the ACA-MCA and PCA-MCA regions are each scored from 0-5 compared to the contralateral hemisphere, with the total score ranging from 0-10: **0** – Absent, **1** – Minimal, **2** – Vessel visibility and extent significantly reduced, **3** – Vessel visibility and extent moderately reduced, **4** – Vessel visibility and extent mildly reduced, **5** – Normal or increased visibility and extent.



ASPECTS 20 POINTS GRADING SYSTEM	N, %	Chi2, p
< 6	10; 6.2%	110.11; p<0.001
6-10	37; 22.8%	
11-20	115; 71.0%	



$r = - 0.606, p < 0.001$

DSA-Based Grading Systems

Among the Digital Angiography methods, the **ASITN/SIR Collateral Scale** is the most frequently used. This method is the most widely recognized assessment system. The collateral scale developed by the **American Society of Interventional and Therapeutic Neuroradiology / Society of Interventional Radiology (ASITN/SIR)**, based on DSA, is graded from **0 to 4 points**: **0** – No collaterals visible in the ischemic area, **1** – Slow, peripheral filling of the ischemic area, with some defects, **2** – Rapid, peripheral filling of the ischemic area, with some defects, **3** – Slow, but complete filling of the ischemic area with blood flow in the late venous phase, **4** – Complete and rapid collateral flow in the ischemic area, driven by retrograde perfusion.

The **Christoforidis Collateral Grading System**, also based on DSA, assesses collateral status with **5 points**: **1** – The entire distal part of the occluded vessel is reconstituted by collaterals, **2** – The proximal part of the adjacent segment of the occluded vessel is reconstituted by collaterals, **3** – The distal part of the adjacent segment of the occluded vessel is reconstituted by collaterals, **4** – Blood flow separated by two segments from the closed vessel is reconstituted by collaterals, **5** – Minimal or practically non-reconstituted blood circulation.

Conclusion

There is a strong correlation between collateral circulation assessed by all four systems and final neurological outcome, assessed by the modified Rankin scale. for MITEFF SYSTEM $r = - 0.585, p < 0.001$, for MAAS SYSTEM $r = - 0.549, p < 0.001$, for Modified Tan SYSTEM score $r = 0.58828, P < 0,001$, for ASPECTS 20 POINTS GRADING SYSTEM $r = - 0.606, p < 0.001$,

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