Mechanical thrombectomy (MT) for acute ischemic stroke (AIS) in COVID-19 pandemic: a systematic review

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Abstract

**Introduction:** Coronavirus disease 2019 (COVID-19) is a disease caused by severe acute respiratory syndrome coronavirus 2 (SARS-CoV-2). Initially, COVID-19 is a disease that attacks the respiratory tract, but now the clinical manifestations of COVID-19 are various, including acute ischemic stroke (AIS). Emergency surgeries such as mechanical thrombectomy (MT) for AIS must be performed without any delay even during the COVID-19 pandemic, to reduce morbidity and mortality. Besides the focus on patient’s health, the safety of healthcare workers must also be considered. The aim of the study was to evaluate and summarize the scientific literature systematically to explore MT for AIS in the COVID-19 pandemic.

**Data synthesis:** The independent reviewers searched the literature through 12 electronic databases, searching for articles fulfilling inclusion and exclusion criteria. The data from all included studies were presented in a summary table featuring key points of each study. The authors independently assessed the risk of bias of 15 included articles.

**Conclusion:** Although MT procedure has been prolonged during the pandemic, clinical outcomes and procedure-related serious adverse events have remained unchanged during the COVID-19 pandemic. The screening process and the implementation of the PCS algorithm must be performed to reduce the spread of COVID-19 infection without threatening patient safety and clinical outcomes. The standard precaution of infection and the health assurance of healthcare workers and their families (including mental health) are also important factors that must be given special attention and consideration in the COVID-19 pandemic.

**Keywords:** Acute ischemic stroke, COVID-19, Ischemic stroke, Mechanical thrombectomy

Introduction

Coronavirus disease 2019 (COVID-19) is a disease caused by severe acute respiratory syndrome coronavirus 2 (SARS-CoV-2) [1–4]. Phylogenetic analysis showed that SARS-CoV-2 had 79.5% and 51% similarity with severe acute respiratory coronavirus syndrome (SARS-CoV) and Middle East respiratory coronavirus syndrome (MERS-CoV) [2, 5]. The first case of COVID-19 occurred in December 2019 in Wuhan, Hubei Province, China with sources of transmission linked to the fish market and several wild animals (birds, snakes, and bats) in Wuhan [1, 6, 7]. SARS-CoV-2 can spread through direct contact (droplet and transmission between humans) and indirect contact (airborne transmission and contaminated objects) [2]. SARS-CoV-2 has spread widely in more than 250 countries, until 12 March 2020 COVID-19 was established as a global pandemic by the World Health Organization (WHO) with a world fatality rate of 5.28% [8–11]. Until June 20, 2020, there were 8,745,570 cases worldwide with a total of 461,760 deaths [9].

Initially, COVID-19 is a disease that attacks the respiratory tract with angiotensin-converting enzyme 2 (ACE2) as the main receptor, but now the clinical manifestations of COVID-19 are various, including neurological disorders. A neurological disorder that needs...
special attention because of their morbidity and mortality that can be suppressed if treated in the golden period is acute stroke [12–18]. The cytopathic effect of the virus and dysregulation of the immune system can cause severe inflammation, including inflammatory cytokine storms that cause COVID-19-associated coagulopathy (CAC) or thrombosis, and can cause acute ischemic stroke (AIS) [13]. SARS-CoV-2 has also been shown to be associated with thromboembolic disease and hypercoagulability state through mechanisms of hypoxia, inflammation, and disseminated intravascular coagulation (DIC) [19]. Inflammation, CAC, and endothelial dysfunction can also increase the permeability of the blood-brain barrier, so that the virus can enter the central nervous system (CNS) through transcellular, paracellular, and retrograde transport of axons through the sensory and olfactory nerves (lamina cribiform and olfactory bulb) [13, 16, 20]. The incidence of COVID-19 patients with stroke in China and Europe was about 2.5% to 6% [13, 21]. The most common type of stroke is ischemic stroke (84.6%), followed by central venous thrombosis (7.7%), and hemorrhagic stroke (7.7%) [16, 22].

In the beginning, mechanical thrombectomy (MT) for AIS had not shown a better outcome in comparison with conventional medical treatments. But now with the better understanding and procedure of MT, MT is reported to have a clear advantage for AIS over conventional medical treatments [23–25]. Currently, MT has been widely accepted as the first-line therapy for patients with emergent large vessel occlusion (LVO) stroke [23, 26, 27]. During the COVID-19 pandemic, many operations have been postponed or even canceled due to the need for SARS-CoV-2 infection screening and relocation of limited medical resources. Even though, emergency surgeries such as MT for AIS must be performed without any delay even during the pandemic, to reduce morbidity and mortality [28, 29]. Besides the focus on patient’s health, the safety of healthcare workers must also be considered in the COVID-19 pandemic. Protected code stroke is a term used during the COVID-19 pandemic to prioritize acute assessment and management of AIS patients and to provide safety and protection of healthcare workers and also patients [30]. Expert consensuses from Chinese and European Federation recommended that all patients including those receiving MT should undergo screening process (preoperative chest CT scan and multidisciplinary consultations to exclude COVID-19) and the PCS algorithm [29–32].

MT for AIS in COVID-19 pandemic have been reported and explained in various studies, but there was no systematic review about it. The aim of the study was to evaluate and summarize the scientific literature systematically to explore MT for AIS in the COVID-19 pandemic.

Main text (review of the literature, results, discussion)

Review of literature

Inclusion and exclusion criteria

Inclusion criteria in this systematic review were publication type was full-text articles discussing MT for AIS in COVID-19 pandemic and primary studies of every design (descriptive study, such as case report and case series; observational study such as cross-sectional, case-control, and cohort; and experimental study such as clinical trial); the language of publication was English; time of publication was in December 2019–December 2020; and objective, methodology, and outcome measure must explain MT for AIS in the COVID-19 pandemic. Exclusion criteria in this systematic review was confounding variables were related to outcome in MT for AIS in the COVID-19 pandemic.

Literature search

This systematic review was conducted according to the Cochrane handbook for systematic reviews and the guideline of preferred reporting items for systematic review and meta-analysis (PRISMA) [33, 34]. A systematic search literature was used in these electronic databases: Cambridge Core, Clinical Key, Ebsco, Emerald Insight, JSTOR, Medline, Nature, Proquest, PubMed, Science Direct, Scopus, and Springer Link. The search was conducted using the following keywords for title and abstract: (thrombectomy OR mechanical thrombectomy) AND (stroke OR ischemic stroke) AND (COVID-19 OR coronavirus OR SARS-CoV-2). The reference lists from retrieved literature were also examined to avoid missing any published data.

Data collection and analysis

Studies were selected for evaluation after two independent reviewers (AK and DT) had collected titles and abstracts identified in the electronic database. The results of the two independent reviewers were compared by a third independent reviewer (YA), and any differences of opinion were resolved by discussion. Full papers from potential studies were independently assessed by the investigators (R and RA). All studies selected for this systematic review were screened by two reviewers independently to validate the results (AK and JB). The data from all retrieved studies were presented in a summary table featuring key points of each study. The key points of each study were: first author, country, and year; study design; sample; outcome measure; and result.

Quality assessment

The lead author independently assessed quality assessment and risk of bias of each included study and discussed them with other authors. Quality assessment and
risk of bias within studies were assessed using criteria developed by Hawker and colleagues [35, 36]. Ratings were concluded (very poor, poor, fair, good, and not applicable) across nine different categories: abstract and title; introduction and aim; method and data; sampling; data analysis; ethic and bias; result; generalisability; and implication and usefulness. The risk of bias potentially affecting the cumulative evidence across studies was determined by examining study methods, ethics committee approvals, study funding, and conflicts of interest [35, 36]. Newcastle–Ottawa scale for cohort study was also used to assess the methodological quality of prospective study; interpretation of total score was ≥7 points were included in good studies, 5–6 points were included in fair studies, <5 points were included in poor studies. Newcastle-Ottawa scale adapted for cross-sectional study was used to assess the methodological quality of the cross-sectional study. Interpretation of total score was 9 to 10 points were considered in very good studies, 7 to 8 points were considered in good studies, 5 to 6 points were considered in satisfactory studies, and 0 to 4 points were considered in unsatisfactory studies [37–41]. The Joanna Briggs Institute (JBI) critical appraisal checklist was used to assess the methodological quality of the case report and case series [42–44].

Results

Selection of articles for review

Figure 1 summarized the identified, screened, and included articles for review. Initially, 288 peer-reviewed articles were identified from electronic databases and an additional 5 articles were identified through other sources (search engine). After removing duplicates, 168 articles remained for the title and abstract screening. Articles that did not meet the inclusion and exclusion criteria were not further screened. Eighteen articles were screened for eligibility of which 15 articles met all the inclusion criteria.

Assessment of study validity (quality assessment and risk of bias)

All eligible studies were associated with MT for AIS in the COVID-19 pandemic. Table 1 provided quality assessment and risk of bias by Hawker and colleagues, and all of the studies are fair and good. Table 2 provided quality scores for cohort study and the study got 7 points that were considered in good study. Table 3 provided quality scores for cross-sectional study and all of the studies got 5–8 points that were considered satisfactory and good studies. Tables 4 and 5 showed JBI critical appraisal checklist for case report and case series; all of the studies had overall appraisal in “included studies” for systematic review.
Study characteristic

The study characteristics for the included studies could be seen in Table 5. There were one cohort study, eight cross-sectional studies, three case series studies, and one case report. The studies reported about patient characteristics (baseline characteristic, clinical presentation, laboratory examination, and radiology examination), MT procedure, clinical outcome, the care delays, and PCS of AIS in the COVID-19 pandemic (Table 6).

Discussion

MT for AIS in COVID-19 pandemic

There are some guidelines related to MT for AIS in the COVID-19 pandemic but there is no systematic review about it. The differences about MT for AIS in COVID-19 pandemic and non-pandemic are about patient characteristics (baseline characteristic, clinical presentation, laboratory examination, and radiology examination), MT procedure, clinical outcome, the care delays, and PCS. The differences in patient characteristics were not discussed in this systematic review because they were more related to stroke in COVID-19 generally. All of the AIS types were LVO because MT was frequently used in LVO case [23, 26, 27].

Kerleroux and colleagues (2020) reported about 21% decrease in MT cases in the first month of the COVID-19 pandemic in French [28]. Kwan and colleagues (2020) reported a decrease of 21% in MT procedure from the external referral hospital in UK [49]. McConachie and colleagues (2020) reported about 27.7% decrease of MT procedure in April 2020 compared with the first 3 months of 2020 in the UK [51]. Qureshi and colleagues (2020) reported significant reduction of MT procedure in the USA [57]. The most possible reason was the strict guidelines of stroke care centers to perform MT in eligible patients. Patients with indications outside of strict guidelines may have not been accepted for MT [28, 49]. Other plausible reasons were an increase in healthy lifestyles during the COVID-19 pandemic (reduced the incidence of stroke), the strict PCS and screening procedure, or it could be due to the patient’s reluctance to come to the hospital/emergency room (especially in ‘mild’

Table 1 Quality assessment and risk of bias by Hawker and colleagues

| No. | First author, country | Abstract and title | Introduction and aim | Method and data | Sampling and analysis | Ethic and bias | Finding Generalisability | Implication and usefulness |
|-----|-----------------------|--------------------|----------------------|----------------|-----------------------|---------------|-----------------------------|-----------------------------|
| 1.  | Al Kasab S; North and South America, Europe [45] | Good | Good | Good | Good | Good | Good | Good | Good |
| 2.  | Cox M, USA [46] | Good | Good | Fair | Fair | Fair | Good | Good | Good |
| 3.  | Escalard E, France [47] | Good | Good | Good | Good | Fair | Good | Good | Good |
| 4.  | Havenon A, USA [48] | Good | Good | Good | Good | Fair | Good | Good | Good |
| 5.  | Kerleroux B, France [28] | Good | Good | Good | Good | Good | Good | Good | Good |
| 6.  | Kwan J, UK [49] | Good | Good | Fair | Fair | Fair | Good | Good | Good |
| 7.  | Mansour OY, Egypt [50] | Good | Fair | Fair | Fair | Good | Fair | Good | Good |
| 8.  | McConachie D, UK [51] | Good | Good | Good | Good | Good | Good | Good | Good |
| 9.  | Pop R, France [52] | Good | Good | Good | Good | Good | Good | Good | Good |
| 10. | Tiedt, Germany [53] | Good | Good | Good | Good | Good | Good | Good | Good |
| 11. | Qureshi A L, USA [54] | Fair | Fair | Fair | Fair | Fair | Good | Fair | Fair |
| 12. | Wang A, USA [19] | Good | Good | Fair | Fair | Good | Fair | Good | Good |
| 13. | Yaeger K A, USA [55] | Good | Good | Good | Fair | Good | Good | Good | Good |
| 14. | Yang B, China [29] | Good | Good | Good | Good | Good | Good | Good | Good |
| 15. | Yeboah K, USA [56] | Fair | Fair | Fair | Fair | Good | Fair | Good | Good |

Table 2 Newcastle-Ottawa scale (cohort study)

| No. | First author, year | Selection | Comparability | Outcome | Total |
|-----|---------------------|-----------|---------------|---------|-------|
| 1.  | Tiedt, Germany [53] | * * * * | * * * | * | 7 |

Maximum points for selection number 4, comparability, and outcome number 1 were 2
Selection: (1) representativeness of the sample, (2) sample size, (3) non-respondents, (4) risk factor measurement tool
Outcome: (1) assessment of the outcome, (2) statistical test
symptoms of stroke) for fear of being exposed to the COVID-19. It is important to explain and educate the public that this is not a problem because the hospital has taken standard precautions to ensure that patients and medical personnel are protected and the hospital remains the best and safe place to provide appropriate treatment for emergency cases such as AIS care. Hospital administration must also ensure patient safety through appropriate standard precautions and the use of personal protective equipment (PPE) [10, 58]. PCS is the modified of regular code stroke during the COVID-19 pandemic to provide an additional layer of protection (including PPE) for patients and medical personnel who were engaged in triage, rapid assessment, COVID-19 screening, and treatment of patients [30]. Cox and colleagues (2020), Mansour and colleagues (2020), Yaeger and colleagues (2020), and Yeboah and colleagues (2020) and reported about the importance of healthcare worker to follow PCS algorithm and the use of appropriate PPE to reduce the risk of COVID-19 infection [46, 50, 55, 56].

Pop and colleagues (2020) reported that there were 39.6% reduction of stroke alerts and 27.6% decrease in MT procedure [52]. Yang and colleagues (2020) reported about the prolonged door to puncture time (by 48.5 min, \( p: 0.002 \)) and door to reperfusion time (by 41 min, \( p: 0.047 \)) [29]. Kerleroux and colleagues (2020) reported the delays between imaging to groin time (by 29 min, \( p < 0.001 \)) [28]. Tiedt and colleagues (2020) reported about the longer of door to puncture time in patients (47 min vs. 38 min, \( p: 0.005 \)) [53]. The plausible reasons were the saturation of health medical resources as a result of the COVID-19 pandemic; screening process and the extra standard and isolation precautions applied in PCS; social distancing of each individual; and also the fear of medical worker’s, patient’s, or family’s fear of COVID-19 contamination risk at the hospital [28, 29, 52, 53]. In some research, these care delays did not impact the short-term outcomes. Although there was no significant difference to short-term outcome (follow up with the national institutes of health stroke scale/NIHSS in day 1–3 after MT), there were not long-term follow-up (such as 90 days of the modified rankin scale/mRS) to evaluate the long-term efficacy [28, 29, 49, 50].

MT technique used in COVID-19 pandemic is various related to the operator (direct aspiration first-pass

### Table 3 Newastle-Ottawa scale adapted for cross-sectional study

| No. | First author, country | Selection | Comparability | Outcome | Total |
|-----|-----------------------|-----------|---------------|---------|-------|
| 1.  | Al Kasab S; North and South America, Europe [45] | * | * | * | * | 8 |
| 2.  | Cox M, USA [46] | * | * | * | * | 6 |
| 3.  | Havenon Ad, USA [48] | * | * | * | * | 7 |
| 4.  | Kerleroux B, France [28] | * | * | * | * | 7 |
| 5.  | Kwan J, UK [49] | * | * | * | * | 7 |
| 6.  | McConachie D, UK [51] | * | * | * | * | 7 |
| 7.  | Pop R, France [52] | * | * | * | * | 7 |
| 8.  | Qureshi A I, USA [54] | * | * | * | * | 5 |
| 9.  | Yang B, China [29] | * | * | * | * | 8 |

**Maximum point for comparability was 2**

**Selection:** (1) representativeness, (2) selection of non-exposed, (3) ascertainment of exposure, (4) demonstration that outcome was not present at the beginning

**Outcome:** (1) assessment of the outcome, (2) follow up long enough, (3) adequacy of follow up

### Table 4 JBI Critical Appraisal Checklist for case report

| No. | Major components | Masnour OY, Egypt [50] | Yeboah K, USA [56] |
|-----|------------------|-----------------------|--------------------|
| 1.  | Were patient’s demographic characteristics clearly described? | Yes | Yes |
| 2.  | Was the patient’s history clearly described and presented as a timeline? | Yes | Yes |
| 3.  | Was the current clinical condition of the patient on presentation clearly described? | Yes | Yes |
| 4.  | Were diagnostic tests or assessment methods and the results clearly described? | Yes | Yes |
| 5.  | Was the intervention(s) or treatment procedure(s) clearly described? | Yes | Yes |
| 6.  | Was the post-intervention clinical condition clearly described? | Yes | Yes |
| 7.  | Were adverse events (harm) or unanticipated events identified and described? | Yes | Yes |
| 8.  | Does the case report provide takeaway lessons? | Yes | Yes |

**Overall appraisal**

Include Include
technique/ADAPT, stent retrieval, or stent-aspiration combination/solumbra technique [29]. In Wang A and colleagues (2020), the operator used solumbra technique [29]. Solumbra technique is safe technique and increases the rate of successful reperfusion [24, 29]. The difficulties of the MT process in COVID-19 pandemic were re-occlusion of the vessel soon after reperfusion (related to systemic hypercoagulability, microvascular thrombo-inflammation, and endotheliitis), low rate of fragmentation, and distal embolization (due to clot composition of reddish clot-suggestive high red blood cell content and neutrophil extracellular traps) [19, 47]. Yang and colleagues (2020) reported about the reduction of puncture to reperfusion time (from 40.5 to 32 min) without the effect of patient safety and successful reperfusion rates during the COVID-19 pandemic. A series of ways were performed to reduce the time of the MT procedure, including performing the entire procedure by an experienced neurointerventional specialist and reducing angiography processes based on good CT angiography images. Other possible reasons were the use of local anesthesia and first-line ADAPT strategy [29]. Sedation and anesthesia choice for MT in COVID-19 pandemic present its own challenges and limited published data are available at this time to guide appropriate decisions and conclusions. Al Kasab and colleagues (2020) reported that general anesthesia was associated with poor functional outcome and mortality in COVID-19 patients [45]. General anesthesia with intubation carries the appeal of a closed respiratory circuit and the load of viral particles in the environment is decreased. However, bag-mask ventilation and intubation itself are highly aerosol-generating procedures. The negative pressure of environment and room, the minimum number of staff with full PPE, and the use of video laryngoscopy can be used to reduce the risk of COVID-19 infection. The use of conscious sedation when possible (such as sufficient oxygenation) avoids this intense aerosolization. The appropriate MT technique and anesthesia choice is still being debated and further research is needed [19, 29, 49, 50, 59, 60].

Clinical outcomes in patients with stroke and COVID-19 who undergo MT are various in some studies. Escalard and colleagues (2020) and Wang and colleagues (2020) reported about the poor clinical outcome; otherwise, Mansour and colleagues (2020), Yaeger and colleagues (2020), Yang and colleagues (2020), and Yeboah and colleagues (2020) reported about the good clinical outcome; while Kwan and colleagues (2020) and Tiedt and colleagues (2020) reported about the similar clinical outcome in pre-COVID-19 vs. COVID-19 periods [19, 29, 47, 50, 55, 56]. The difference in patient characteristics affected on this difference and need further research with similar characteristics on the global scale. Havenon and colleagues (2020) reported an increased risk of death in patients who undergo MT with comorbid COVID-19 [48]. The systemic complication (including acute respiratory failure, acute renal failure, and coagulopathy), and the delay of diagnosis and treatment in COVID-19 patient were the likely explanation of it [13, 19, 48].

### Strength and limitation of the study

This systematic review involved studies that reported 15 studies related to MT for AIS in the COVID-19 pandemic. Most of the studies were analytical observational studies (eight) studies and the majority of the studies discussed the comparison of MT in AIS patients before and during the COVID-19 pandemic.

The limitation of the study was the variance of the demography, limited follow-up time, confounding
| No. | First author, country | Study design | Sample (n) | Outcome measure | Result |
|-----|----------------------|--------------|------------|-----------------|--------|
| 1.  | Al Kasab S; North and South America, Europe [45] | Cross-sectional | 458 | The effect of GA on mortality and discharge outcome | GA had longer door to reperfusion time (138 vs. 100 min, \( p < 0.001 \)), higher mortality (RR: 1.871, \( p = 0.029 \)), and lower functional outcome discharge (RR: 0.053, \( p = 0.015 \)). |
| 2.  | Cox M, USA [46] | Cross-sectional | 45 | PCS in MT of AIS patients | The importance of PCS implementation and the use of PPE during MT. |
| 3.  | Escalard E, France [47] | Case series | 10 | Patient outcome | Successful MT was performed in 9 patients, none had good early neurological outcomes, and 6 patients died in the hospital. |
| 4.  | Havenon A, USA [48] | Cross-sectional | 3145 | Comparison of the outcome of MT in COVID-19 and non COVID-19 | There was a 21% reduction in MT case (OR: 0.79, 95% CI: 0.76–0.82, \( p < 0.001 \)), significant delays between imaging to puncture time (mean 144.9 ± 86.8 vs. 126.2 ± 70.9 min, \( p < 0.001 \) in 2019) and imaging to in-transferred patients (mean 182.6 ± 82.0 vs. 153.25 ± 67 min, \( p < 0.001 \)), compared with the same period in 2019. There was a significant negative correlation between the number of COVID-19 hospitalizations and the number of MT cases compared with the same period in 2019 (\( r = 0.51, p = 0.04 \)). |
| 5.  | Kerleroux B, France [28] | Cross-sectional | 1513 | Comparison of MT in AIS patients before and during COVID-19 pandemic | There was a 21% reduction in MT case (OR: 0.79, 95% CI: 0.76–0.82, \( p < 0.001 \)), significant delays between imaging to puncture time (mean 144.9 ± 86.8 vs. 126.2 ± 70.9 min, \( p < 0.001 \) in 2019) and imaging to in-transferred patients (mean 182.6 ± 82.0 vs. 153.25 ± 67 min, \( p < 0.001 \)), compared with the same period in 2019. There was a significant negative correlation between the number of COVID-19 hospitalizations and the number of MT cases compared with the same period in 2019 (\( r = 0.51, p = 0.04 \)). |
| 6.  | Kwan J, UK [49] | Cross-sectional | 61 | Comparison of MT in AIS patients before and during COVID-19 pandemic | During the COVID-19 pandemic, (a) MT rate was maintained at 20% of AIS and there was a non-significant 21% decrease in MT, referred from the external hospital (\( p = 0.067 \)); (b) successful reperfusion rate was maintained at 81% and early neurological outcomes were not significantly different; (c) the use of general anesthesia reduced significantly from 85 to 32% (\( p < 0.05 \)); and (d) time intervals from onset to arrival, puncture, and reperfusion were not significantly different, whereas internal delays for external referrals significantly increased for door to puncture time (48 [IQR 39–57] vs. 33 [IQR 27–44] minutes, \( p = 0.013 \)) and door to reperfusion time (82.5 [IQR 61–110] vs. 60 [IQR 55–70] minutes, \( p = 0.018 \)). |
| 7.  | Mansour OY, Egypt [50] | Case report | 1 | Patient outcome, PCS in MT of AIS patients | The NIHSS score decreased to 2.2 after reperfusion (from 14 before MT). The importance of PCS implementation and the use of PPE during MT. |
| 8.  | McConachie D, UK [51] | Cross-sectional | 27 | Comparison of MT in AIS patients before and during COVID-19 pandemic | Three centers did not perform MT, there was a 27.7% decrease of MT procedures in April 2020, and 22 centers reported delays of stroke care. 17 centers reported the reduction of training opportunities for specialist registrars and 14 centers reported the delay of development plans of MT service. |
| 9.  | Pop R, France [52] | Cross-sectional | 122 | Comparison of MT in AIS patients before and during COVID-19 pandemic | There were 39.6% reduction of stroke alerts and 27.6% decrease in MT procedures in March 2020, compared to the same period in 2019. There were no significant differences in time delays or clinical outcomes for patients treated by MT. |
| 10. | Tiedt S, Germany [53] | Cohort | 795 | Comparison of MT in AIS patients before and during COVID-19 pandemic | There was prolonged door to groin time in 2020, compared with the same period in 2019 (47 min vs. 38 min, \( p = 0.005 \)). Functional outcome of patients treated with MT in 2020 was not significantly different compared to patients treated in 2019 (\( p > 0.05 \)). |
| 11. | Qureshi A I, USA [54] | Cross-sectional | 24 | Comparison of MT in AIS patients before and during COVID-19 pandemic | There was a significant reduction of MT procedures in March 2020 (\( p < 0.05 \)). |
| 12. | Wang A, USA [19] | Case series | 5 | Patient demographic, laboratory value, MT technique, and clinical and outcome | 4 patients with COVID-19 had AIS with occlusion in anterior circulation and 1 patient with occlusions in anterior and posterior circulation, the average of age was 52.8 years, and all patients had coagulation abnormalities. Stent-aspiration combination technique was performed in all patients with poor clinical outcomes. |
| 13. | Yaeger K A, USA [55] | Case | 10 | Patient outcome | Successful MT was performed in 9 patients with the NIHSS score decreased to 2.2 after reperfusion (from 14 before MT). The importance of PCS implementation and the use of PPE during MT. |
variable in each study (there were confounding variables that cannot be controlled in human subjects), and also the limitation of study type (only an observational study, and the cohort study was only one).

**Future implication**
The current systematic review can be a scientific reading, material, and consideration to physician, researcher, and all of the readers related to MT for AIS in the COVID-19 pandemic. Further research is needed for the selection of appropriate MT technique and anesthesia choice, and also the evaluation of long-term follow-up related to MT for AIS in the COVID-19 pandemic.

**Conclusion**
Although MT procedure has been prolonged during the pandemic, clinical outcomes and procedure-related serious adverse events have remained unchanged during the COVID-19 pandemic. The screening process and the implementation of the PCS algorithm must be performed to reduce the spread of COVID-19 infection without threatening patient safety and clinical outcomes. The standard precaution of infection and the health assurance of healthcare workers and their families (including mental health) are also important factors that must be given special attention and consideration in the COVID-19 pandemic.

**Abbreviations**
95% CI: 95% confidence interval; ACE2: Angiotensin-converting enzyme 2; AIS: Acute ischemic stroke; CAC: COVID-19 associated coagulopathy; CNS: Central nervous system; COVID-19: Coronavirus disease 2019; GA: General anesthesia; IQR: Interquartile range; LVO: Large vessel occlusion; min: Minutes; MERS-CoV: Middle East respiratory coronavirus syndrome; MT: Mechanical thrombectomy; NIHSS: The National Institutes of Health Stroke Scale; OR: Odds ratio; p: Probability; PCS: Protected code stroke; PPE: Personal protective equipment; PRISMA: Preferred reporting items for systematic review and meta-analysis; r: Correlation coefficient (correlation test); RR: Relative risk; SARS-CoV: Severe acute respiratory syndrome coronavirus; SARS-CoV-2: Severe acute respiratory syndrome coronavirus 2

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**References**
1. Li H, Liu SM, Yu XH, Tang SL, Tang CK. Coronavirus disease 2019 (COVID-19): current status and future perspectives. Int J Antimicrob Agents. 2020;55(5):1–8.
2. Lofft M, Hamblin MR, Rezaei N. COVID-19: Transmission, prevention, and potential therapeutic opportunities. Clin Chlm Acta. 2020;508:254–66. https://doi.org/10.1016/j.cca.2020.05.044.
3. Sheereen MA, Khan S, Kazmi A, Bashir N, Siddique R. COVID-19 infection: origin, transmission, and characteristics of human coronaviruses. J Adv Res. 2020;24:91–8. https://doi.org/10.1016/j.jare.2020.03.005.
4. Yuki K, Fujiogi M, Koutsogiannak S. COVID-19 pathophysiology: a review. Clin Immunol. 2020;215:1–7.
5. Zhu N, Zhang D, Wang W, Li X, Yang B, Song J, et al. A novel coronavirus from patients with pneumonia in China, 2019. N Engl J Med. 2020;382(8):727–33. https://doi.org/10.1056/NEJMoa2001017.
6. Rothan HA, Byrareddy SN. The epidemiology and pathogenesis of coronavirus disease (COVID-19) outbreak. J Autoimmun. 2020;109:1–4.
7. Fu B, Chen Y. The 2019 novel coronavirus disease with secondary ischemic stroke: two cases report, 2019, p. 1–12.
8. Avula A, Nalleballe K, Narula N, Sapozhnikov S, Dandu V, Toom S, et al. COVID-19 presenting as stroke. Brain Behav Immun. 2020;87:115–9. https://doi.org/10.1016/j.bbi.2020.04.077.
9. WHO. Coronavirus disease 2019 (COVID-19) Situation report. Geneva: WHO; 2020. https://www.who.int/emergencies/diseases/novel-coronavirus-2019/situation-reports.

**Table 6** Study characteristic (Continued)

| No. | First author, country | Study design | Sample (n) | Outcome measure | Result |
|-----|-----------------------|--------------|------------|-----------------|--------|
| 14. | Yang B, China [29]    | Cross-sectional | 55         | Comparison of MT in AIS patients before and during COVID-19 pandemic | There was significant increase in door to puncture time (174 vs. 125.5 min, p = 0.002) and door to reperfusion time (213 vs. 172 min, p = 0.047) in the COVID-19 pandemic, compared with the same period in 2019. The rate of successful MT was not significantly different between the two groups (85.7% (n = 18) vs. 88.2% (n = 30), OR 0.971, 95% CI: 0.785–1.203; p = 1.000). |
| 15. | Yeboah K, USA [56]    | Case report   | 1          | Patient outcome, PCS in MT of AIS patients | The NIHSS score decreased to 5 after reperfusion (from 14 before reperfusion) and reduced to 0 in day 2 post-reperfusion. The importance of PCS implementation and the use of PPE during MT. |
10. Zhao J, Rudd A, Liu R. Challenges and potential solutions of stroke care during the coronavirus disease 2019 (COVID-19) outbreak. Stroke. 2020;51(5):1356–7. https://doi.org/10.1161/STROKEAHA.120.029701.

11. Li X, Geng M, Peng Y, Meng L, Lu S. Molecular immune pathogenesis and diagnosis of COVID-19. J Pharm Anal. 2020;10(2):102–8. https://doi.org/10.1016/j.jpha.2020.03.001.

12. Pinzon RT, Wijaya VO, Buana RB, Al Jody A, Nunsio PN. Neurologic manifestations of COVID-19 and other coronavirus infections: a systematic review. Clin Neurol Neurosurg. 2020;194:1.

13. Hassett C, Gedansky A, Mays M, Uchino K. Acute ischemic stroke and respiratory virus infections. Front Cell Neurosci. 2018;12:1.

14. Mao L, Jin H, Wang M, Hu Y, Chen S, He Q, et al. Neurologic manifestations of COVID-19 and other coronavirus infections: a systematic review. J Neurointerv Surg. 2020;12(6):683–90. https://doi.org/10.1136/jninterveneuro-2020-011127.

15. Montalvan V, Lee J, Bueso T, De Toledo J, Rivas K. Neurological manifestations of COVID-19 and other coronavirus infections: a systematic review. Clin Neurol Neurosurg. 2020;194:1.

16. Pinzon RT, Wilja VO, Buana BB, Al Jody A, Nunsio PN. Neurologic characteristics in coronavirus disease 2019 (COVID-19): a systematic review and meta-analysis. Front Neurol. 2020;11:1–11.

17. Wang HY, Li XL, Yan ZR, Sun XP, Han J, Zhang BW. Potential neurological symptoms of COVID-19. Ther Adv Neurol Disord. 2020;29(8):1–12.

18. Whittaker A, Anson M, Harky A. Neurological manifestations of COVID-19: a systematic review and current update. Acta Neurol Scand. 2020;142(1):14–22. https://doi.org/10.1111/jn.13266.

19. Wang A, Mandigo GK, Yim PD, Meyers PM, Lavine SD. Stroke and mechanical thrombectomy in patients with COVID-19: Technical observations and patient characteristics. J Neurointerv Surg. 2020;12(7):648–53. https://doi.org/10.1136/neurintneo-2020-016220.

20. Bohwinkel K, Gálvez NMS, Ríos M, Kalergis AM. Neurologic alterations due to respiratory virus infections. Front Cell Neurosci. 2018;12:1–15.

21. Harrett C, Gedansky A, Mays M, Uchino K. Acute ischemic stroke and COVID-19. Cleve Clin J Med. 2020;87(1):https://doi.org/10.3944/ccjm.2019-0878.

22. Li Y, Li M, Wang M, Zhou Y, Chang J, Xian Y, Wang D, Mao L, Jin H, Hu B. Acute cerebrovascular disease following COVID-19: a single center, retrospective, observational study. Stroke Vasc Neurol. 2020;5(3):279–84. https://doi.org/10.1177/2055385920935428.

23. Zhao W, Ma P, Zhang P, Yue X. Mechanical thrombectomy for acute ischemic stroke in octogenarians: a systematic review and meta-analysis. Front Neurol. 2020;11:1–5.

24. Munich SA, Vakharia H, Levy EI. Overview of mechanical thrombectomy techniques. Neurosurgery. 2019;85(1S):S60–7. https://doi.org/10.1093/neuros/nyy071.

25. Powers WJ, Rabinstein AA, Ackerson T, Adeoye OM, Bambakidis NC, Becker K, et al. Guidelines for the early management of patients with acute ischemic stroke: 2019 update to the 2018 guidelines for the early management of acute ischemic stroke. Stroke. 2019;50:344–418.

26. Goyal N, Tsivgoulis G, Malhotra K, Ishaq 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. J Neurointerv Surg. 2019;11(7):16–24.

27. Lambinos A, Schaink AK, Dhalla I, Krings T, Casaubon LK, Sikich N, et al. Mechanical thrombectomy in acute ischemic stroke: a systematic review. Can J Neurol Sci. 2016;43(4):455–60. https://doi.org/10.1016/j.cjns.2016.03.001.

28. Kerleroux B, Fabacher T, Bitoucet N, Moise M, Testud B, Vingadassalom S, et al. Mechanical thrombectomy for acute ischemic stroke amid the COVID-19 outbreak. Stroke. 2020;51(7):2012–7. https://doi.org/10.1161/STROKEAHA.120.030373.

29. Yang B, Wang T, Chen J, Chen Y, Wang Y, Gao P, et al. Impact of the COVID-19 pandemic on the process and outcome of thrombectomy for acute ischemic stroke. J Neurointerv Surg. 2020;12(7):664–8. https://doi.org/10.1136/neurintneo-2020-016177.

30. Khorsavani H, Rajendran P, Notario L, Chapman MG, Menon BK. Protected code stroke: hyperacute stroke management during the coronavirus disease 2019 (COVID-19) Pandemic. Stroke. 2020;51(6):1891–3. https://doi.org/10.1161/STROKEAHA.120.029838.

31. Aggour M, White P, Kulcitar Z, Fiehler J, Brouwer P. European Society of minimally invasive neurological therapy (ESMINT) recommendations for optimal interventional neurovascular management in the COVID-19 era. J Neurointerv Surg. 2020;12(6):542–4. https://doi.org/10.1136/neurintneo-2020-016137.

32. He Y, Hong T, Wang M, Jiao L, Ge Y, Haacke EM, et al. Prevention and control of COVID-19 in neurointerventional surgery: Expert consensus from the Chinese Federation of Interventional and Therapeutic Neuroradiology (CFITN) and the International Society for Neurovascular Disease (ISNVD). J Neurointerv Surg. 2020;12(7):686–63. https://doi.org/10.1136/neurintneo-2020-016073.

33. Higgins J, Green S. Cochrane handbook for systematic reviews of intervention 5.2. United Kingdom: Wiley; 2017. p. 1–50.

34. Mohler D, Liberati A, Tetzlaff J, Altman DG, Group TP. Preferred reporting items for systematic reviews and meta-analyses: The PRISMA statement. PLoS Med. 2009;6(7):1–6.

35. Clay-Williams R, Ludlow K, Testa L, Vingadassalom S, Jandaok S, Sanchez A, et al. The newcastle-ottawa scale (NOS) for assessing the quality of nonrandomised studies in meta-analyses. 2019.

36. Herzog R, Álvarez-pasquin MI, Díaz C, Luis J, Barrio D, Estrada JM, et al. Are healthcare workers’ intentions to vaccinate related to their knowledge, beliefs and attitudes? A systematic review. BMC Public Health. 2020;20:https://doi.org/10.1186/s12889-020-08984-2.

37. Aggour M, White P, Kulcitar Z, Fiehler J, Brouwer P. European Society of minimally invasive neurological therapy (ESMINT) recommendations for optimal interventional neurovascular management in the COVID-19 era. J Neurointerv Surg. 2020;12(6):542–4. https://doi.org/10.1136/neurintneo-2020-016137.

38. He Y, Hong T, Wang M, Jiao L, Ge Y, Haacke EM, et al. Prevention and control of COVID-19 in neurointerventional surgery: Expert consensus from the Chinese Federation of Interventional and Therapeutic Neuroradiology (CFITN) and the International Society for Neurovascular Disease (ISNVD). J Neurointerv Surg. 2020;12(7):686–63. https://doi.org/10.1136/neurintneo-2020-016073.

39. Higgins J, Green S. Cochrane handbook for systematic reviews of intervention 5.2. United Kingdom: Wiley; 2017. p. 1–50.

40. Mohler D, Liberati A, Tetzlaff J, Altman DG, Group TP. Preferred reporting items for systematic reviews and meta-analyses: The PRISMA statement. PLoS Med. 2009;6(7):1–6.

41. Clay-Williams R, Ludlow K, Testa L, Vingadassalom S, Jandaok S, Sanchez A, et al. The newcastle-ottawa scale (NOS) for assessing the quality of nonrandomised studies in meta-analyses. 2019.

42. Aggour M, White P, Kulcitar Z, Fiehler J, Brouwer P. European Society of minimally invasive neurological therapy (ESMINT) recommendations for optimal interventional neurovascular management in the COVID-19 era. J Neurointerv Surg. 2020;12(6):542–4. https://doi.org/10.1136/neurintneo-2020-016137.
51. McConachie D, McConachie N, White P, Crossley R, Izzath W. Mechanical thrombectomy for acute ischaemic stroke during the COVID-19 pandemic: changes to UK practice and lessons learned. Clin Radiol. 2020;75(10):795.e7–795.e13. https://doi.org/10.1016/j.crad.2020.07.001.

52. Pop R, Quenardelle V, Hasiu A, Mihoc D, Sellaf F, Dugay MH, et al. Impact of the COVID-19 outbreak on acute stroke pathways - insights from the Alsace region in France. Eur J Neurol. 2020;27(9):1783–7. https://doi.org/10.1111/ene.14316.

53. Tiedt S, Bode FJ, Uphau T, Alegiani A, Gröschel K, Petzold GC. Impact of the COVID-19-pandemic on thrombectomy services in Germany. Neurol Res Pr. 2020;2(44):1–6.

54. Qureshi A, Siddiq F, French B, Gomez C, Jani V, Hassan A, et al. Effect of COVID-19 Pandemic on mechanical thrombectomy for acute ischemic stroke treatment in United States. J Stroke Cerebrovasc Dis. 2020;29(10):1–3.

55. Yaeger KA, Fifi JT, Lara-Reyna J, Rossitto C, Ladner T, Yim B, et al. Initial Stroke Thrombectomy Experience in New York City during the COVID-19 Pandemic. AJNR Am J Neuroradiol. 2020;41(8):1357–60. https://doi.org/10.31 74/ajnr.A6652.

56. Yeboah K, Edgell R, Conway J, Alshekhlee A. Interventional Stroke Management in a Patient With COVID-19. Neurol Clin Pract. 2021;11(2): e199–e201. https://doi.org/10.1212/CNP.0000000000000884.

57. Qureshi AI, Abd-allah F, Al-senani F, Aytac E, Borhani-haghighi A, Ciccone A, et al. Management of acute ischemic stroke in patients with COVID-19 infection: insights from an international panel. Am J Emerg Med. 2020;38:1548.e5–7.

58. Fiehler J, Brouwer P, Díaz C, Hirsch JA, Kulcsar Z, Liebskind D, et al. COVID-19 and neurointerventional service worldwide: a survey of the European Society of Minimally Invasive Neurological Therapy (ESWINT), the Society of NeuroInterventional Surgery (SNIS), the Sociedad Iberoamericana de Neuroradiologia Diagnostica y. J Neurointerv Surg. 2020;0:1–5.

59. Leslie-Mazwi TM, Fargen KM, Levitt M, Derdeyn CP, Feske SK, Patel AB, et al. Preserving Access: A Review of Stroke Thrombectomy during the COVID-19 Pandemic. AJNR Am J Neuroradiol. 2020;41(7):1136–41. https://doi.org/10.31 74/ajnr.A6006.

60. Salahuddin H, Castonguay AC, Zaidi SF, Burgess R, Jadhav AP, Jumaa MA. Interventional stroke care in the era of COVID-19. Front Neurol. 2020;11(468): 1–8.

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