Systolic pressure target after endovascular thrombectomy in acute large-vessel occlusion ischemic stroke patients: Comment on ENCHANTED2/MT

Yunpeng Yuan¹ | Baozhu Wei^{2,3} | Wenyun Zhu⁴ | Yang Liu⁵ | Yingfeng Wan⁶

¹Department of Neurosurgery, Baoshan Campus, Shuguang Hospital, Shanghai University

of Traditional Chinese Medicine, Shanghai, China

²Department of Cardiology, Zhongnan Hospital, Wuhan University, Wuhan, China

³Institute of Myocardial Injury and Repair, Wuhan University, Wuhan, China

⁴Department of Gastroenterology, Renjin People's Hospital, Ruijin, China

⁵Department of Endocrinology, China Resources and WISCO General Hospital, Wuhan,

China

⁶Department of Neurosurgery, University of Michigan, Ann Arbor, Michigan, USA

Correspondence

Yingfeng Wan, Department of Neurosurgery, Medical School, University of Michigan,
109 Zina Pitcher Place, R5016 Biomedical Science Research Building, Ann Arbor, MI
48109, USA.

E-mail: yingfenw@med.umich.edu

1

This is the author manuscript accepted for publication and has undergone full peer review but has not been through the copyediting, typesetting, pagination and proofreading process, which may lead to differences between this version and the Version of Record. Please cite this article as doi: 10.1002/brx2.4.

Patients with large-vessel occlusion ischemic stroke continue to have high mortality and disability rates after mechanical thrombectomy with or without intravenous alteplase treatment. Elevated blood pressure during the perioperative period is associated with higher mortality and disability prevalence rates^[1]. Thus, lowering post-procedure systolic pressure is a potential approach to improving patients' outcomes. The guideline^[2] recommends a systolic pressure of < 180 mmHg before and after mechanical thrombectomy without randomized clinical trial evidence.

Recently, Yang and coauthors^[3] reported the results of the multicenter, open-label, blinded-endpoint, randomized controlled trial ENCHANTED2/MT, which aimed to determine if a more aggressive blood pressure goal is beneficial in patients with acute ischemic stroke. Patients were required to have a diagnosis of large-vessel occlusion acute ischemic stroke and a successful endovascular thrombectomy procedure followed by hypertension (defined as ≥ 2 successive measurements of systolic blood pressure ≥ 140 mmHg for > 10 min). The randomization arms were < 120 mmHg vs 140–180 mmHg for 72 hours, and the primary outcome was assessed by a shift analysis of the modified Rankin Scale at 90 days. The 821 patients who were prospectively enrolled between 2020 and 2022 were randomized, and the source populations were derived from 44 hospitals in China. The trial was suspended in June 2022 due to safety concerns after an independent data and safety monitoring board reviewed the data. Unexpectedly, the primary results were that the more intensive treatment group was more likely to have worse outcomes and higher early neurological deterioration and disability rates compared to the less intensive treatment

group.

This study proved again the complex relationship between blood pressure and functional outcomes after acute large-vessel occlusion ischemic stroke^[1]. This is an important trial that was built on accumulating observational data in the field and provided randomized trial evidence that more intensive blood pressure lowering (< 120 mmHg) is not only neutral but harmful. In this regard, this study is of broad general interest to emergency departments and stroke centers, as blood pressure is a parameter that must be managed in all stroke patients post-procedure.

However, several points need to be noted and comprehensively discussed before interpreting the results and applying them clinically. First, Figure 2 from the ENCHANTED2/MT trial paper^[3] suggested that a systolic pressure of < 120 mmHg was only narrowly achieved in the more intensive treatment group during the first 3 days. Although current guidelines^[2] recommend a blood pressure of < 180 mmHg, no optimal blood pressure target for patients with ischemic stroke who undergo mechanical thrombectomy has been identified to date^[1]. Taken together, doctors may need to employ individualized targets for systolic pressure control after mechanical thrombectomy based on the patient's prior medical history and current blood pressure condition as well as the success of the occluded vessel recanalization. Second, blood pressure is elevated in approximately 75% of patients with acute stroke and usually decreases spontaneously over the subsequent few days^[4]. In the ENCHANTED2/MT trial^[3], 267 (66%) patients in the more intensive treatment group and 261 (64%) patients in the less intensive treatment group

had premorbid hypertension, limiting the likelihood that the poststroke outcomes were due to premorbid hypertension rates. Spontaneous poststroke hypertension may be a compensatory response to the ischemic stroke; thus, relatively low blood pressure may be partly responsible for the worse functional outcomes in the more intensive treatment group. Third, it is noteworthy that there was no difference in symptomatic intracerebral hemorrhage rates between the two groups, consistent with the previously published BP-TARGET trial^[5], in which an aggressive systolic pressure goal of 100–129 mmHg after successful reperfusion in patients with ischemic stroke did not reduce intracerebral hemorrhage rates at 24–36 hours when compared to a standard care systolic blood pressure goal of 130–185 mmHg, suggesting either a lack of protection at this level of more intensive treatment of systolic pressure or a lack of harm at this level of less intensive treatment of systolic pressure regarding symptomatic intraparenchymal hemorrhage transformation.

To answer the question of what the optimal systolic pressure target should be after mechanical thrombectomy in patients with acute large-vessel occlusion stroke, the results of ongoing relevant clinical trials and future more rigorously designed clinical trials are required.

Funding: None

Ethical Approval: Not applicable

Conflict of Interest: Yunpeng Yuan, Baozhu Wei, Wenyun Zhu, Yang Liu, and Yingfeng

Wan declare no conflicts of interest.

Author Contribution:

Author contribution: Yunpeng Yuan (Writing-original draft); Baozhu Wei (Writing-original draft);

Wenyun Zhu (Writing-original draft); Yang Liu (Writing-original draft); Yingfeng Wan (Writing-review &

editing).

References:

- 1. Anadani M, de Havenon A, Mistry E, Anderson CS. Blood Pressure Management After Endovascular Therapy: An Ongoing Debate. *Stroke*. Jun 2021;52(6):e263-e265. doi:10.1161/STROKEAHA.121.034995
- 2. Powers WJ, Rabinstein AA, Ackerson T, 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: A Guideline for Healthcare Professionals From the American Heart Association/American Stroke Association. *Stroke*. Dec 2019;50(12):e344-e418. doi:10.1161/STR.0000000000000011
- 3. Yang P, Song L, Zhang Y, et al. Intensive blood pressure control after endovascular thrombectomy for acute ischaemic stroke (ENCHANTED2/MT): a multicentre, open-label, blinded-endpoint, randomised controlled trial. *Lancet*. Nov 5 2022;400(10363):1585-1596.

doi:10.1016/S0140-6736(22)01882-7

- 4. Osaki Y, Matsubayashi K, Yamasaki M, et al. Post-stroke hypertension correlates with neurologic recovery in patients with acute ischemic stroke. *Hypertens Res.* Sep 1998;21(3):169-73. doi:10.1291/hypres.21.169
- 5. Mazighi M, Richard S, Lapergue B, et al. Safety and efficacy of intensive blood pressure lowering after successful endovascular therapy in acute ischaemic stroke (BP-TARGET): a multicentre, open-label, randomised controlled trial. *Lancet Neurol*. Apr 2021;20(4):265-274. doi:10.1016/S1474-4422(20)30483-X