Systemic Intravenous Thrombolysis for Massive Pulmonary Embolism after Intracranial Surgery for Traumatic Head Injury: When Absolute Becomes Relative!

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Abstract
Intravenous thrombolysis is the preferred treatment for massive pulmonary embolism with hemodynamic instability, when there are no contraindications to the therapy. However, when absolute contraindications to thrombolysis are present, clinicians are at a crux, especially when mechanical thrombectomy is also not possible. We present a case of a 40-year-old man with polytrauma with head injury who developed massive pulmonary embolism requiring intravenous thrombolysis on day 15 after decompressive craniotomy—possibly a first in the literature. The patient, however, successfully recovered from this near-fatal episode. With this report, we wish to highlight that at a near-dying situation, no contraindication is absolute, and all possible efforts must be made by the clinician to save the patient.

Keywords
► massive pulmonary embolism
► alteplase
► traumatic head injury
► thrombolysis

Introduction
Trauma patients have a high risk of venous thromboembolism, with the incidence of deep vein thrombosis (DVT) reaching as high as 58% in case of traumatic head injury.¹ P<ul class="download-pdf"

Case Description
A 40-year-old man, road traffic accident victim, with no known/existing comorbidities, was brought to the hospital with an arrival GCS score of E2M4Vt. Noncontrast computed tomography (CT) of the head was suggestive of multiple frontal bone fractures, anterior cranial fossa fractures, multiple hemorrhagic contusions, subarachnoid hemorrhage (SAH), and pneumocephalus in bilateral frontal and basifrontal lobes (Fig. 1). CT of the whole spine was normal. The patient had multiple rib fractures but no long bone fractures. Complete blood count, liver and renal function tests, and two-dimensional (2D) echo were normal with ejection fraction of 65%. Sequential compression stockings were applied as per the ICU protocol to prevent DVT.

On day 1, the patient underwent bifrontal craniotomy, right frontal contusectomy, anterior cranial fossa repair

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(by neurosurgical team), and open reduction and internal fixation of the frontal, zygomatic and maxillary bones (by plastic surgery team). On day 2, he underwent redo bilateral frontal craniotomy for hematoma evacuation. From days 3 to 12, he had intermittent fever spikes (for which he underwent cerebrospinal fluid [CSF] analysis by lumbar puncture on days 5 and 11, and meningitis was ruled out) and one episode of seizures on day 5. GCS score during this time fluctuated between E2VtM3 and E2VtM2. Enoxaparin at 40 mg once daily for thromboembolism prophylaxis was started on day 12. By day 15, the patient was successfully weaned off ventilator to T-piece for 48 hours. Fever spikes (up to 38.9°C) intermittently continued.

On day 16 morning, the patient had fever (39°C), was tachypneic (respiratory rate ≈ 25 breaths/min) and was started on continuous positive airway pressure (CPAP) support. Systolic blood pressure (SBP) was low (= 90–100 mm Hg) that responded to fluid boluses. At this time, 2D echo was normal with no obvious lower limb swellings. Serum procalcitonin was measured (suspecting a new-onset infection), which came 1.0 ng/mL (similar to the value 4 days back). In the evening, however, the patient suddenly became severely tachypneic (respiratory rate = 35–40 breaths/min), with desaturation (SpO₂ up to 92% on FiO₂ 50%) and needed progressively higher ventilatory and inotropic support. At this time, 2D echo done was suggestive of severe pulmonary hypertension, dilated right atrium and ventricle, and right ventricular ejection fraction (RVEF) 35%. CT pulmonary angiography was suggestive of bilateral pulmonary artery thrombosis in the main, ascending, descending, and subsegmental branches (►Fig. 2). CT venography showed partial filling defect of the left common femoral, popliteal, and tibial veins and right superficial femoral and popliteal vein.

As the patient’s hemodynamics became progressively unstable (SBP = 60 mm Hg on triple inotropes), he was considered unsuitable for mechanical thrombectomy or catheter-guided thrombolysis. With no better alternative other than intravenous thrombolysis, after consulting and explaining the eventuality to family including high risk of intracranial bleed, the patient was started on alteplase 100 mg over 6 hours followed by heparin infusion for 24 hours. His blood pressure (BP) gradually improved over 6 hours, and he was off inotropes in next 24 hours. GCS was E1VtM1 for next 3 days but then gradually improved. The patient underwent vena cava filter (IVC) filter placement on day 17 and was later started on rivaroxaban. He was shifted to the ward from ICU on day 35 with GCS of E3VtM5 where his GCS improved to E3VtM6 by day 60.

**Discussion**

The authors believe that the use of alteplase for thrombolysis in case of traumatic brain injury with recent decompressive craniectomy for massive PE has not been described previously in the literature. A significant head trauma in previous 3 months, history of intracranial hemorrhage, and recent intracranial surgery are (absolute) exclusion criteria for alteplase administration after acute stroke because of the risk of worsening intracranial hemorrhage. Therefore, massive PE in neurocritical care unit becomes a challenging situation.

Heparinization followed by warfarinization for PE has been done successfully previously in case of a traumatic brain injury with nephrotic syndrome by Akimoto et al. They noticed slight increase in the hemorrhagic contusions with this therapy. Bottinor et al used tPA in case of a massive PE to save life of a 60-year-old who had suffered from hemorrhagic
stroke 8 weeks prior. Their daring trial achieved much-needed appreciation, so we should not follow the guidelines blindly but must also make clinical judgments as per the need of situation. Similarly, thrombolytic bolus was successful in saving a patient’s life following intracranial surgery for glioblastoma multiforme who suffered from cardiac arrest while being prepared for the mechanical thrombolysis via ultrasonography for pulmonary embolus dissolution.

With this case report, we wish to highlight that although intravenous thrombolysis is contraindicated in cases of recent head injury and intracranial bleed, even this absolute contraindication should be considered relative in case of an eventually dying patient. We were left with no option to save the patient’s life but to try intravenous thrombolysis. In this case, our patient fortunately recovered from PE with no complication of thrombolysis. Having said that, we acknowledge the wait in starting enoxaparin early and diagnosing PE earlier where we could have stepped up our antithrombotic measures. Therefore, one must be more vigilant in suspecting PE in neurosurgical ICU where antithrombotic measures are usually delayed or underdosed. Fever, a consistent sign in our patient was probably due to DVT, and its association carries worse outcome.

Conclusion
We thus recommend that every effort must be made for prevention and early diagnosis of DVT and PE in neurocritical care unit, as role of thrombolysis in case of massive PE is limited. If, however, a dire situation arises to save a patient, all the absolute contraindications for thrombolysis become relative (as recent intracranial surgery and cerebral contusions in this case). The clinician must not be hesitant in counseling the family and taking the best possible course at that moment.

References
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