Abstract

Title: Identification of an Isolated Epidural Hematoma Using Blood-Based Biomarkers in Traumatic Brain Injury

Authors: Katja Malmi¹, Dana Pisica^{2,3}, Iftakher Hossain^{1,9}, Mehrbod Mohammadian^{1,4}, Teemu M. Luoto⁵, Riikka S.K. Takala⁶, Olli Tenovuo⁷, Thomas van Essen^{8,9,10}, John K. Yue¹¹, Andrew I.R. Maas¹², David K. Menon¹³, András Büki¹⁴, Jussi P. Posti¹, and CENTER-TBI Investigators and Participants.

Affiliations: ¹Neurocenter, Department of Neurosurgery and Turku Brain Injury Center, Turku University Hospital and University of Turku, Turku, Finland, ²Department of Neurosurgery, Erasmus MC - University Medical Center, Rotterdam, the Netherlands, ³Center for Medical Decision Making, Department of Public Health, Erasmus MC -University Medical Center, Rotterdam, the Netherlands, ⁴Department of Radiology, Massachusetts General Hospital, Boston MA, USA, ⁵Department of Neurosurgery, Tampere University Hospital and Tampere University, Tampere, Finland ⁶Perioperative Services, Intensive Care Medicine and Pain Management, Turku University Hospital and University of Turku, Turku, Finland, ⁷Division of Clinical Neurosciences, University of Turku, Turku, Finland, ⁸University Neurosurgical Center Holland, Leiden University Medical Center, Haaglanden Medical Center, HAGA, Leiden and The Hague, the Netherlands ⁹Division of Neurosurgery, Department of Clinical Neurosciences, University of Cambridge and Addenbrooke's Hospital, Cambridge, United Kingdom, ¹⁰Department of Surgery, Division of Neurosurgery, QEII Health Sciences Centre and Dalhousie University, Halifax, Nova Scotia, Canada, ¹¹Brain and Spinal Injury Center, Department of Neurological Surgery, Zuckerberg San Francisco General Hospital, University of California San Francisco, San Francisco, USA, ¹²Department of Neurosurgery, Antwerp University Hospital and University of Antwerp, Edegem, Belgium, ¹³Division of Anaesthesia, University of Cambridge, Addenbrooke's Hospital, Cambridge, UK, ¹⁴School of Medical Sciences, Örebro University, Örebro, Sweden, CENTER-TBI Investigators and Participants

Background: Blood-based biomarkers have shown promise in identifying head injury patients with acute traumatic intracranial findings, however their capacity in detecting patients with isolated extra-axial hematomas remains unknown. The objective of this study was to assess the potential S100 calcium-binding protein B (S100B), glial fibrillary acid protein (GFAP), total tau (t-tau), neurofilament light (NF-L), ubiquitin C-terminal hydrolase L1 (UCH-L1), and neuron specific enolase (NSE) to identify mild TBI (mTBI) patients with epidural hematoma (EDH).

Materials and methods: This prospective study included 1043 patients (median age 46 years, interquartile range 29-63) with mild traumatic brain injury [mTBI (GCS \geq 13)] with 1) all six serum blood biomarker levels measured and 2) acute head computed tomography (CT) scan within 24 hours of injury. The main outcome measure was the area under the curve of the receiver operating characteristic (AUC) for each biomarker in distinguishing between patients with CT-negative findings and those with EDH without coexisting surgical findings, i.e. subdural or intraparenchymal haemorrhage. An AUC \geq 0.7 was considered clinically adequate. A distinction was made between i) isolated EDH and ii) EDH occuring concomitantly with non-surgical findings (e.g. subarachnoid haemorrhage, intraventricular haemorrhage, and/or signs of axonal injury).

Results: Of the patients, 37 (3,5%) patients had an isolated EDH, 30 (2,5%) patients had EDH with non-surgical lesions and 976 patients had CT-negative findings (94%). In discriminating between any EDH versus CT-negative, the AUCs were 0.83 for GFAP, 0.65 for NSE, 0.66 for t-tau, 0.60 for NF-L, 0.57 for S100B, and 0.67 for UCH-L1. For

discriminating between isolated EDH and CT-negative, the AUCs were 0.77 for GFAP, 0.63 for NSE, 0.58 for t-tau, 0.52 for NF-L, 0.51 for S100B, and 0.60 for UCH-L1. **Conclusion:** Blood levels of GFAP adequately discriminated mTBI patients with EDH.