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**HYPERACUTE BLOOD-BASED PROTEOMIC AND MACHINE
LEARNING SIGNATURES FOR STROKE SUBTYPE
DIFFERENTIATION: A TRANSLATIONAL POINT-OF-CARE
DIAGNOSTIC FRAMEWORK**

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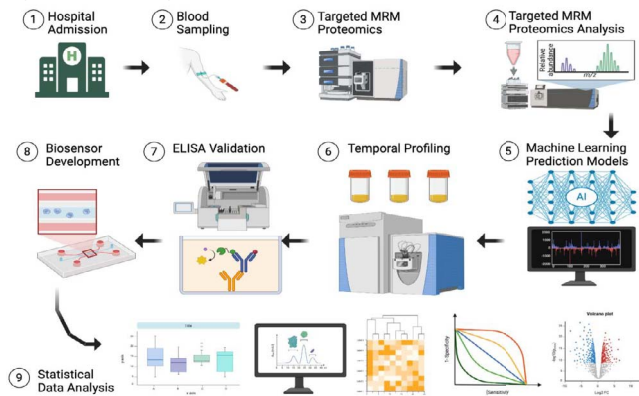
Background and aims: Quickly distinguishing between ischemic stroke, intracerebral hemorrhage, transient ischemic attack, and stroke mimics during the hyperacute phase remains a vital unmet need in stroke treatment. This issue is especially significant in settings where immediate neuroimaging is unavailable or delayed, hindering timely reperfusion and proper triage.

To validate blood-based protein biomarkers for early stroke subtype differentiation using targeted proteomics and machine learning, and to translate the most informative biomarkers into a rapid point-of-care diagnostic assay.

Methods: In this prospective, multi-center diagnostic study, patients presenting within 6 hours of symptom onset with suspected stroke (ischemic stroke, intracerebral hemorrhage), transient ischemic attack, or stroke mimics will be recruited. Serum samples will undergo multiple reaction monitoring-based targeted proteomics to quantify 22 candidate proteins identified in prior high-throughput discovery studies. Supervised machine learning models—including logistic regression, random forest, support vector machines, gradient boosting, and deep neural networks—will be trained to differentiate stroke subtypes and predict 90-day functional outcomes. Temporal profiling of top-performing biomarkers will be conducted in a predefined patient subset. The most informative biomarkers will undergo ELISA validation and be integrated into a gold nanoparticle-based multiplex point-of-care biosensor designed for prehospital and emergency department deployment.

Results: We hypothesize that a multi-protein biomarker panel combined with machine learning will show high diagnostic accuracy for distinguishing ischemic stroke from hemorrhagic stroke, transient ischemic attack, and stroke mimics within 6 hours of onset. Currently, proteomics studies are underway and the prototype for the point-of-care biosensor device is being developed with several iterations underway.

Conflict of interest: Manabesh Nath: nothing to disclose, Deepti Vibha: nothing to disclose

Figure 1 - belongs to Methods**Figure 2 - belongs to Results**