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Reviewer's comments:

The authors reviewed the biochemical markers (protein) of brain injury such as traumatic brain injury, stroke, subarachnoid haemorrhage, and intra cranial space occupying lesions. A few protein markers including S-100 protein, neuron specific enolase (NSE), creatinine phosphokinase isoenzyme BB (CPK-BB) and myelin basic protein (MBP) have been proven to have prognostic and clinical value in the brain injury. This manuscript is interesting, and is well written. The biochemical makers could also include the RNA molecular, and a lot of papers were published on RNA markers of brain injury. I do understand the authors focus on the protein level, it would be necessary to mention the RNA markers of brain injury in background.

Authors reply:

The authors would like to thank the reviewer for the kind review and considering our manuscript favorably. We feel honored that the reviewer has observed that "This manuscript is interesting, and is well written".

As suggested, we have added the RNA markers of brain injury in:

1. Abstract: “while expression of small non-coding RNAs have presented themselves as potential markers of brain injury for future”

2. Core tip: “In addition expression of small non coding RNAs have presented themselves as potential markers of brain injury for future research.

3. New developments: “Proteomic analysis of potential new CSF biomarkers for TBI has not yet identified any such markers that can be used in clinically useful tests^[156]. A number of proteomics studies on potential biomarkers of TBI in peripheral blood have been published. These studies have replicated the findings from targeted analyses of specific candidate biomarkers, but as yet none of the novel biomarker profiles identified in these studies as being associated with TBI has been validated in independent studies using unrelated, non-proteomic or genomic techniques^[157]. Exciting preliminary data on the expression profiles of small noncoding RNAs in peripheral blood mononuclear cells from military personnel exposed to mild TBI have been reported; three small RNAs seem to be primarily associated with mild TBI, but the results require replication^[158].”