

Original Papers

A pilot study of cerebral and haemodynamic physiological changes during sedation with dexmedetomidine or propofol in patients with acute brain injury

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SUMMARY

Sedation for the mechanically-ventilated, brain-injured patient remains challenging. The purpose of this pilot study was to compare the cerebral physiologic effects of sedation with propofol versus dexmedetomidine in mechanically-ventilated, brain-injured patients. Using a randomised, crossover, unblinded clinical trial, we enrolled patients with severe brain injury (Glasgow Coma Score ≤ 8) from traumatic injury, subarachnoid haemorrhage or intracerebral haemorrhage undergoing multimodal monitoring (intracranial pressure, brain temperature, oximetry and microdialysis). Patients received an infusion of either propofol or dexmedetomidine for six hours and then a crossover for the subsequent six hours after sufficient washout/in. Clinical and physiological measurements were recorded hourly. In eight patients, (four traumatic injury, three subarachnoid haemorrhage and one intracerebral haemorrhage), the mean dose of propofol used was 25.5 $\mu\text{g/kg/minute}$ while the mean dose of dexmedetomidine was 0.54 $\mu\text{g/kg/hour}$. All subjects were effectively sedated to a goal of Richmond Agitation Sedation Scale -2 and Bispectral Index of 50–70 throughout the study period. We did not observe any statistically significant differences between the groups in systemic or cerebral physiologic metrics. Though differences were noted in cerebral metabolic substrates (lactate/pyruvate ratio), none were statistically significant. In our pilot cohort, dexmedetomidine and propofol appear equally effective in sedating severely brain-injured patients and neither is associated with adverse physiologic effects as measured by multimodal monitoring. Larger long-term studies are required to determine whether clinically favourable benefits demonstrated in the medical critical care setting also apply to this patient population.

Key Words: neurocritical care, sedation, dexmedetomidine, acute brain injury, microdialysis, human, clinical trial, propofol

Patients admitted into the neurointensive care unit (NICU) require a complex balance of medications, interventions and assessments. Patients admitted with severe cerebral damage exemplify this group. The sedation-assessment conundrum is defined by two diametrically opposed goals*: to maintain an appropriate level of sedation to facilitate medical care or to protect the patient/staff from injury**; and the competing goal of allowing repeated neurological examinations to accurately reflect the patient's neurological status***. In short, it is imperative to know

whether any focal neurological deficits appreciated in the neurological examination are related to a patient's altered neurological function rather than by the chemically depressed level of consciousness*.

Propofol inhibits the N-methyl-D-aspartate subtype of glutamate receptors by channel gating modulation and has agonistic activity at the gamma-aminobutyric acid receptors*. Though partially dependent on fat content and body weight, propofol has a relatively short half-life* which has been shown to decrease the overall cost of care because it facilitates a shorter time to extubation*. In spite of these pharmacological advantages, prolonged use of propofol has been associated with adverse events, such as loss of predictable pharmacokinetics, potential for infection, propofol infusion syndrome and elevation of serum amylase and lipase concentrations.

Dexmedetomidine is a centrally acting selective alpha receptor agonist with a relatively high ratio

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Traumatic Brain Injury Papers

Wenbin Ji



Traumatic Brain Injury Papers:

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