Original Article

Utility of bispectral index in the management of multiple trauma patients

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Abstract

Background: Bispectral index (BIS) monitoring in multiple trauma patients has become a common practice in monitoring the sedation levels. We aimed to assess the utility of BIS in the trauma intensive care unit (ICU).

Methods: A prospective observational study was conducted in the trauma ICU at Hamad General Hospital in Qatar between 2011 and 2012. Patients were divided in two groups: Group I (without BIS monitoring) and Group II (with BIS monitoring). The depth of sedation was clinically evaluated with Ramsey Sedation Scale, changes in vital signs and Glasgow Coma Scale (GCS) level. Use of sedatives, analgesics, and muscle relaxants were also recorded. Data were compared using Chi-square and Student *t*-tests.

Results: A total of 110 mechanically ventilated trauma patients were enrolled with a mean age of 36 ± 14 years. The rate of head injury was greater in Group I when compared with Group II (94% vs. 81%, P = 0.04). In comparison to Group I, patients in Group II had lower GCS and higher mean Injury Severity Score (ISS) ($6.3 \pm 2.5 \text{ vs. } 7.4 \pm 2.7 \text{ and } 25.5 \pm 8.5 \text{ vs. } 21.2 \pm 4.7$, respectively, P = 0.03). The used midazolam dose was less in Group II in comparison to Group I ($5.2 \pm 2.3 \text{ vs. } 6.1 \pm 2.1$, P = 0.03). Also, fentanyl dose was less in Group II ($152 \pm 58 \text{ vs. } 187 \pm 59$, P = 0.004). The rate of agitation, failure of extubation and tracheostomy in Group II were lower than those in Group I, P = 0.001. The length of stay for patients Group I was longer ($14.6 \pm 7.1 \text{ vs. } 10.2 \pm 5.9 \text{ days}$) in comparison to group II, P = 0.001.

Conclusion: Management of multiple trauma patients in the trauma ICU with BIS monitoring was found to be associated with better outcomes. BIS monitoring is a guide for adjusting the dosage of sedative agents. It can also minimize agitation, failure of extubation, and length of stay in ICU.

Key words: Analgesia, bispectral index, head injury, sedation, trauma



INTRODUCTION

Optimal management of multi-traumatized patients in intensive care units (ICUs), especially in those who are connected to mechanical ventilators includes use of sedation and analgesia. Analgesia and sedation are needed in order to tolerate clinical procedures, synchrony with ventilators, optimize oxygenation and to secure patient safety and comfort.^[6] A recent study reported that approximately 69% of patients in ICU were inappropriately sedated (54% were over-sedated, and 15% were under-sedated).^[15] Inadequate level of sedation may lead to patient's anxiety, patient-ventilator dyssynchrony, agitation, and posttraumatic stress disorder (PSTD). Undersedation may also cause increased stress symptoms, sodium and water retention, increased catabolism and mobilization of substrates to provide energy sources. It also associated with tachycardia, hypertension, increased oxygen consumption, and altered levels of respiratory rates. Moreover, it induces changes in gastro intestinal mobility and variations in coagulability such as clotting time, and platelet aggregation, in addition to delayed wound healing.^[24] On the other hand, oversedation may lead to delayed weaning that may result in ventilator associated pneumonia. Furthermore, it may prolong artificial ventilation and length of ICU stay, and consequently increases the cost associated with patient care.[14]

Sedation monitoring can be challenging because of variations in patient's requirement of medications. The synergistic action of drugs in different combinations also makes it difficult to achieve the goal of optimal sedation. Moreover, there is no gold standard for measuring sedation in ICU patients. Most of the time, clinicians monitor vital signs such as heart rate and blood pressure and use subjective sedation scales to monitor sedation level. Commonly used subjective methods include Riker sedation-agitation scale (SAS), motor activity assessment (MAAS), Richmond agitation-sedation scale scale (RASS), Adaptation to the intensive care environment scale (ATICE), and Ramsay sedation scale (RSS).^[14] Vital signs do not reflect the depth of sedation always. RSS and SAS rely on patient movement and response and therefore may not precisely measure the deep states of sedation. Weinert et al. added that subjective assessment of sedation is not optimal because of the influence of social, personal, and professional factors on sedation monitoring.^[26] Furthermore, differences in the subjective assessment of sedation are more likely to occur.^[11] An objective measure such as bispectral Index (BIS) monitoring may be a useful tool in monitoring optimal sedation.^[6-8] BIS index is derived from patient's electroencephalogram (EEG) data and scaled from 100 to 0 denoting fully awake to unconscious. BIS value decreases linearly with increasing doses of anesthesia.

Studies among anesthetized patients pointed out the acceptable target ranges; BIS of 40-60 for deep surgical anesthesia and 70-90 for light anesthesia.^[14] Beneficial outcomes of use of BIS monitors include improved sedation, decreased sedative requirements, and faster recovery times. BIS monitoring system is useful in minimizing the incidence of awareness with recall in adults. Awareness is a leading cause of patient dissatisfaction with anesthesia. BIS technology enables to assess consciousness and sedation separately from cardiovascular reactivity. It has been demonstrated that BIS monitored patients wake up faster, extubated sooner, and more oriented on arrival at the postanesthesia care unit (PACU). PACU discharge was sooner among BIS monitored patients. Moreover, BIS-guided anesthesia lead to reductions in use of hypnotics.^[1,4,11] Recently, its use has been extended to ICU patients.^[14,15,24,26] The present study analyzes the utility of BIS monitors in mechanically ventilated patients in trauma ICU.

MATERIAL AND METHODS

The present study was conducted prospectively from December 2011 to December 2012. All adult patients admitted to the trauma ICU at Hamad General Hospital, Doha, Qatar who required mechanical ventilation, sedation, and analgesia were included in the study. Sedation depth was clinically evaluated with RSS, observation of vital signs and Glasgow Coma Scale (GCS). Information on age, gender, mechanism of injury, type of injury, Injury Severity Score (ISS), comorbidity, dose of analgesic and sedative agents used, and outcome measures such as mortality, agitation, failure of extubation, and length of stay in the ICU were recorded. Data were compared between two groups of patients, that is, with BIS monitoring versus without monitoring. Group frequencies were compared with the Chi-square and Student t-test, while the level of significance was set at P < 0.05 for all tests.

In our study, the brain electrical activity was measured using BIS-monitor system (Covidien, Gunbarrel Ave, Boulder, USA). The sensor (BIS Quatro Sensor) used in our study has four electrodes, which were placed at an angle over the forehead of the patient. The first electrode was positioned at the midline, approximately 5 cm above the nose, and the second electrode was placed just lateral and inferior to the first electrode. The third electrode was placed over the temporal region behind the angle of the eye, and the fourth electrode was positioned directly above and adjacent to the eyebrow [Figure 1]. An internal algorithm based on a database of EEGs calculate a number between 0 (absence of brain electrical activity) and 100 (fully awake). This provides a direct measure of level of consciousness in the patients. This study was approved by Hamad Medical Corporation, Medical Research Center, IRB#11222/12.

RESULTS

A total of 110 patients admitted to the trauma ICU were enrolled; of them, 97% were males with a mean age of 36 ± 14 years. All these patients were intubated and connected to mechanical ventilators Table for different indications. l summarizes demographics, presentation and interventions in all the study population. Sixty-five percent of injuries were traffic-related such as motor vehicle crashes (MVCs) and pedestrian injuries. The main injured body region was the head (88%), followed by chest (44%), and abdomen (19%). The rate of head and abdominal injuries were significantly greater in Group I when compared



Figure 1: Bispectral index monitoring device

Table 1: Overall demographics, presentation, and interventions (n=110)

Age	35.6±14.3
Male	105 (97.2)
Mechanism of injury (%)	
Motor vehicle crashes	54 (50)
Pedestrian injury	17 (15.7)
Fall from height	26 (24.1)
Motor cycle crash	2 (1.9)
Fall of heavy object	4 (3.7)
All terrain vehicle	4 (3.7)
Assault	1 (0.9)
Head injury	97 (88.2%)
Chest injury	48 (43.6%)
Abdominal injury	21 (19.1%)
Injury severity score	23.1±6.9
Glasgow coma score	6.9±2.7
Length of stay	12.6 ± 6.9
Propofol	46 ± 10.4
Remifentanil	$0.16 {\pm} 0.06$
Bispectral index	47 (42.7%)
Bispectral index (on admission)	54±16.2
Bispectral index (follow-up)	45 ± 6.4
Intracranial pressure	15.3±5.2
Failed extubation	33 (30%)
Agitation	34 (30.9%)
Tracheostomy	12 (10.9%)

with Group II (94% vs. 81%, P = 0.04 and 27% vs. 8.5%, P = 0.01, respectively). In comparison to Group I, patients with BIS monitoring (Group II) had significant lower GCS and higher mean ISS $(6.3 \pm 2.5 \text{ vs}, 7.4 \pm 2.7,$ P = 0.039 and 25.5 ± 8.5 vs. 21.2 ± 4.7 , P = 0.003, respectively) [Table 2]. The used midazolam dose was less in Group II in comparison to Group I (5.2 \pm 2.3 vs. 6.1 ± 2.1 , P = 0.029). Also, fentanyl dose was less in Group II (152 \pm 58 vs. 187 \pm 59, P = 0.004). The rate of agitation (44% vs. 13%, P = 0.001) and failure of extubation (40% vs. 17%, P = 0.001), and tracheostomy (19% vs. 0%) were higher in Group I when compared with Group II. The hospital length of stay in Group I was shorter in Group II (10.2 ± 5.9 days), in comparison to group I (14.6 \pm 7.1), P = 0.001. The present study observed no correlation between BIS values and clinical sedation scale scores (RSS) (P = 0.49).

DISCUSSION

The current study assesses the utility of BIS monitoring among intubated trauma patients and demonstrates that BIS monitoring has reduced the dose of used analgesics and sedation with a subsequent reduction of the hospital

Table 2: Comparison of ICU patients with and without BIS

	Group-1 (<i>n</i> =63)	Group-2 (<i>n</i> =47)	<i>P</i> value
Age (mean±SD)	35.1 ± 13.4	36.4 ± 15.7	0.643
Male (%)	100	93.3	0.070
Mechanism of injury (%)			0.879
Motor vehicle crashes	52.4	46.7	
Pedestrian injury	14.3	17.8	
Fall from height	23.8	24.4	
Motor cycle crash	1.6	2.2	
Fall of heavy object	4.8	2.2	
All Terrain vehicles	3.2	4.4	
Assault	0	2.2	
Head injury (%)	93.7	80.9	0.040
Chest injury (%)	46	40.4	0.348
Abdominal injury (%)	27	8.5	0.012
Injury severity score (mean \pm SD)	21.2 ± 4.7	25.5 ± 8.5	0.003
Glasgow coma score (mean±SD)	7.4 ± 2.7	6.3 ± 2.5	0.039
Length of stay (mean \pm SD)	14.6 ± 7.1	10.2 ± 5.9	0.001
Sedation (mean \pm SD)			
Midazolam	6.1 ± 2.1	5.3 ± 1.2	0.029
Propofol	46.7 ± 7.7	44.3 ± 16.2	0.720
Analgesia (mean \pm SD)			
Fentanyl	186.7 ± 58.8	156.5 ± 38.9	0.004
Remifentanil	0.14 ± 0.05	$0.21\!\pm\!0.06$	0.082
Failed extubation (%)	39.7	17	0.008
Agitation (%)	44.4	12.8	0.001
Tracheostomy (%)	19	0	0.001

SD: Standard deviation, ICU: Intensive care unit, BIS: Bispectral index

Surgical Neurology International 2014,5:141

length of stay. It also reduced the rate of agitation, failed extubation, and the need for tracheostomy.

In general, BIS was used to titrate sedative agents in ICU patients where it showed decreased use of sedatives and less oversedation when compared with use of subjective methods.^[14] However, a previous study reported no difference in ventilator days or hospital mortality by the use of BIS method compared with standard care.^[17] deWit et al.^[7] reported that BIS scores were unable to predict extubation success or failure in medical ICU patients. In trauma patients, BIS was found reliable and advantageous over subjective scales such as RASS during a propofol spontaneous awakening trial (SAT).^[18] However, Ogilvie et al.^[21] reported that subjective interpretations of BIS are more likely in patients with traumatic brain injury (TBI) or patients receiving paralytic agents. Agitation, irritability, and aggression are the major behavioral complications related to TBI in the ICU. However, the frequency of TBI was significantly higher among non-BIS group in our study. This indicates that the higher incidence of agitation, extubation failure, and tracheostomy observed in our cases might be associated with TBI irrespective of the BIS monitoring. Therefore, patient characteristics differences alone could be associated with higher rate of complication and increased length of hospital stay. Also, it is evident from our analysis that more severely injured patients underwent BIS monitoring, which helps in lowering the rate of sedation and analgesia and reduces the overall hospital length of stay. Consistently, it has been demonstrated that BIS-monitored patients wake up faster, extubated sooner, and more oriented on arrival at the PACU.^[11]

Brocas *et al.*^[3] studied the effect of an alfentanil bolus in mechanically ventilated critically ill patients and showed that BIS was lower with alfentanil bolus. Riker *et al.*^[23] compared BIS and Suppression ratio (SR) with burst suppression of the EEG during Pentobarbital infusions in adult intensive care patients and found BIS was correlated well with SR ($r^2 = 0.79$).

The usefulness of the BIS in monitoring the degree of sedation in ICU patients can be demonstrated by its degree of correlation to commonly used sedation scales. Nasraway *et al.*^[20] conducted a prospective study to validate the BIS to SAS. Patients with SAS of 3 or less were included, that is, deeply sedated patients. The correlation was low, but statistically significant between the BIS and SAS. The correlation was increased when the BIS associated with high Electromyographic (EMG) activity was removed from analysis. The study demonstrated that BIS may be useful in patients who are deeply sedated (SAS score ≤ 3), that is, in patients with lower EMG activity.^[20] Ma *et al.*^[16] compared the reliability of BIS with SAS in assessing the depth of sedation in mechanically ventilated patients in ICU and

found that SAS was well correlated with BIS; and BIS monitoring was more reliable in measuring the depth of sedation especially when SAS reaches 2-4. Furthermore, a study that compared subjective RSS with BIS concluded that BIS monitoring enables more effective titration of sedatives and offers an objective, safe, and reliable measure of sedation.^[9]

A significant correlation between BIS scores and the level of consciousness in brain-injured ICU patients was reported by Jung *et al.*^[13] The investigators pointed out the fact that BIS scores could not indicate a patient's mental status without vagueness except in coma patients. However, the study added that BIS in conjunction with GCS or other neurological evaluation scales will be useful for assessing the level of consciousness in brain-injured patients.^[13]

Paul *et al.*^[22] also reported the significant correlation between BIS and GCS scores in patients with mild-to-moderate head injuries. GCS and BIS were measured before surgery, after surgery and once a day for the first 10 days in 29 patients with mild (GCS 13-15) and moderate (GCS 9-12) head injuries who underwent craniotomy. Hsia *et al.*^[12] reported on the positive correlation that existed between GCS and BIS among critically ill pediatric patients. Gill *et al.*^[10] showed the weak correlation between GCS and BIS in emergency department patients. Notably, the level of consciousness and BIS are two scores obtained from different brain regions that may not correlate with each other as BIS score is the measure of frontal lobe electrical activity.^[13]

BIS monitors are in fact cerebral function monitors since it records the electrical activity of the cerebral cortex. In severe brain injury patients, the electrical activity of the brain may vary according to cerebral perfusion, cerebral metabolism, hypoxia, sedative pharmacologic agents, and seizure activity.^[8,27] Dunham *et al.*^[8] showed that BIS ≥ 60 suggest that patients with severe brain injury are likely to have an acceptable ICP and cerebral perfusion pressure (CPP). Xifeng *et al.*^[27] and Myles *et al.*^[19] reported that BIS provides useful information that may lead to the identification of patients with good chance of recovery after ischemic hypoxic brain injury.^[19,27]

Limitations: BIS values are affected by the choice of anesthetic agents and the depth of sedation among the patients may differ for the same BIS score but with different combinations of drugs.^[14,15] BIS scores have not been proved as a useful tool in certain populations such as critically ill patients with unstable body temperatures and patients with dementia.^[2] In many studies, different investigators used different versions of BIS monitor that makes it difficult to compare the results obtained because of the changes in the algorithm used. BIS-XP version attempts to filter electro-oculographic and other

Surgical Neurology International 2014,5:141

Table 3: Factors that affect bispectral index value

	Expected BIS value
Sedation	Decrease
Analgesia	Decrease
Neuromuscular blocking agents	Decrease
Deep sleep	Lower (20-70)
REM sleep	Higher (75-92)
Hypothermia	Decrease
Cerebral ischemia	Decrease
High-frequency electrical artifact	Increase

BIS: Bispectral index, REM: Rapid eye movement

EMG artifacts. Deogaonkar *et al.*^[5] reported that XP version of BIS showed much higher correlation with sedation scales when comparing with version 2.11. However, Vivien *et al.*^[25] showed overestimation of the BIS due to high EMG activity with both versions and this resulted in undersedation. The data regarding the use of neuromuscular blocking agents, cerebral ischemia, and information for emergence from coma are lacking in our study. Table 3 shows the factors that should be considered during the evaluation of BIS results.

CONCLUSION

Clinically, use of a BIS monitor may help standardize clinical practice and improve patients care. BIS is a guide for adjusting the dosage of sedative agents. It can also minimize the clinical consequences such as sedation overdose, prolonged intubation, failed extubation, and hospital stay. It can also provide financial benefits by limiting excessive use of costly sedatives and decreasing time to extubation. Although the BIS monitor is relatively new to the intensive care setting, it can be a beneficial tool in monitoring neurological patients.

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Commentary

GOALS AND USES

Originally designed for use in the operating theater, the bispectral index monitor may be adapted in the neurocritical care unit as an adjunct to monitor states of unconsciousness. It is used alongside with clinical parameters obtained from clinical examination and sedation scales.

Critically ill patients in the neurocritical care unit represent a heterogeneous group, with individualized goals for levels of analgesia, awareness, and recall ability. In general, three levels of awareness are present: (i) Conscious awareness with pain perception, (ii) conscious awareness without pain perception, and (iii) perception without conscious awareness. Recall ability can be classified into: (i) Explicit recall with conscious recollection of events and (ii) implicit recall with behavioral changes but no conscious recollection of events. Anesthetic goals include: (i) Analgesia with lack of response to noxious stimuli and (ii) hypnosis with lack of awareness and recall.

In the neurocritical care unit, bispectral index monitor is useful for monitoring and ensuring hypnosis in patients on deep sedation with neuromuscular blockade used to decrease cerebral metabolic rate in those with refractory intracranial hypertension and ventilator management. It is also useful for sedation titration in patients with deep coma and locked-in states. The bispectral index value trend may also be useful for sedation weaning in those emerging from unawareness.

MECHANISMS

In bispectral index monitoring, EEG information is converted to digital signals with rejection of noise artifacts. Multivariable statistical models are used to derive a score between 0 and 100. Bispectral index values between 45 and 60 indicate high likelihood of unconsciousness and unawareness. Sedation titration to values below 40 may be associated with sedation-related hemodynamic instability. In those emerging from unaware states, muscle activity often interferes with bispectral index scores above 70. Bispectral index suppression ratio is also useful. It indicates the percentage of the previous 63 s of EEG present as suppression (with 15-30 s time lag).

Bispectral analysis takes into account the frequency, amplitude, and phase angle of the sinusoidal components of EEG. EEG frequencies can overlap with low frequency EMG signals in the range of 30-50 Hz. This produces EMG-induced EEG interference. Focal neurologic pathologies (including EEG changes induced by cerebral blood flow alterations and focal seizures) may be detected if bilateral bispectral index monitor leads are placed. It is also important to note that hypnotic agents, such as propofol, may induce sleep and changes in cortical EEG readings, whereas opioid analgesia suppresses movement but have small effects on EEG.

PITFALLS

Bispectral index monitoring cannot be used in the following situation:

- Extensive frontotemporal contusions and subdural hematomas
- Extensive forehead soft tissue injuries
- Frontotemporal skull fractures
- After decompressive craniectomies, and
- Other conditions preventing application of adherent sensors.

Differing values may be produced by different bispectral index monitoring software.

Unreliable bispectral index values may also be seen in the following conditions:

- Extremes of age with altered sensitivity of bispectral index values
- Body temperature fluctuations (reflects altered protein binding, liver and kidney drug metabolism, and cerebral metabolic rate)
- Ambient noise and equipment affecting signal transmission
- Stimuli (including painful stimuli and tracheal suctioning)
- Muscle activity (including grimacing, shivering and eye movements)
- Anesthetic agents that do not affect EEG (such as ketamine)
- Metabolic states that alter drug metabolism (including liver disease, states of chronic alcoholism and drug use, withdrawal syndromes)
- Metabolic conditions with neurologic manifestations (including encephalopathy associated with multiorgan failure, sepsis, toxic metabolic states, delirium, anoxia with EEG slowing).

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