Quantification of Alcohol Dehydrogenase Concentration for Traumatic Brain Injury in Alcohol-intoxicated Patients as a Predictor of Severity and Prognosis

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Abstract

BACKGROUND: Traumatic brain injury (TBI) stills the leading cause of public health problem and disabilities in many countries, especially in lower-middle-income countries. Motor vehicle crashes, especially motorized two-wheeler riders, are the major cause of TBI, followed by fall and assault. During 2013–2014, a total of 2108 TBI cases (65.4% of them being caused by motor vehicle crashes) were documented in the Emergency Unit of Dr. Hasan Sadikin Hospital (RSHS) Bandung. Among these, 30.7% of cases were categorized as moderate and severe TBI, with a mortality rate of 29.2%. Half of the cases arrived at RSHS within 6 h and 14.0% of these cases were alcohol-intoxicated. Individuals with low activity of alcohol-metabolizing enzymes have low tolerance to alcohol and are therefore rapidly intoxicated, making them more susceptible to more serious injuries from motor vehicle crashes.

METHODS: A prospective study was designed by consecutive sampling from January to August 2017 to review 30 intoxicated patients with TBI on hospital admission by measuring their serum alcohol dehydrogenase (ADH) concentration using an ELISA and then correlated it with the severity, type of TBI lesions, and the outcome.

RESULTS: Demonstrated an ADH level of l d0.24 ng/ml in patients with TBI with a history of alcohol intoxication, which statistically correlated with lower initial Glasgow coma scale (GCS) scores on admission by approximately 6.53 times (p < 0.001), thus increasing the risk of developing multiple lesions by 3.98 times (p = 0.029) and worse outcomes by 5.27 times (p = 0.003), but it did not correlate with length of stay (p = 0.392).

CONCLUSIONS: ADH concentrations in intoxicated TBI patients can be taken into consideration to decide about the therapeutic action and further informed consent to the family.
emergency unit during 2014, of which 65.4% were caused by motorcycle crashes. Of these cases, 30.7% were categorized as moderate or severe TBI, with a mortality rate of 29.2%. More than half of the cases arrived at the emergency unit within 6 h, and 14.0% of them were under alcohol intoxication [8].

Alcohol dehydrogenase (ADH) and aldehyde dehydrogenase (ALDH) are the principal enzymes involved in the catabolism of ethanol. Alcohol is initially metabolized by ADH to acetaldehyde, which is consequently oxidized by ALDH to acetic acid. Individuals with low activity of alcohol-metabolizing enzymes have low tolerance to alcohol and are therefore rapidly intoxicated [9]. Our study aim was to correlate the concentration of ADH in TBI patients with the severity, type of TBI lesion, Glasgow coma scale (GCS) score in the final outcome, and length of stay.

Methods

This was a single-institution retrospective study conducted at RSHS in Bandung, which serves the city, and patients are also referred from West Java Province. RSHS in Bandung is one of the 15 hospitals that have an emergency department and an accompanying intensive care unit in West Java Province. The hospital has 900 beds, and the neurosurgery unit at RSHS admits an average of 117 trauma patients every month. Data were collected from patients who presented to RSHS from January 1 to August 31, 2017. The inclusion criteria were a confirmed diagnosis based on ICD-10 of TBI with a history of alcohol abuse with intoxication (Concussion S06.0 with F10.129) and agreeing to undergo a blood test. Patients who refused to participate in the study were excluded from the study. A database was constructed by reviewing the medical records, which included de-identified data for each eligible patient unit at RSHSIC information, initial GCS score, helmet used, type of lesions, and prognosis based on GCS score when the patient was discharged home (improvement, no improvement, or death), including the length of stay. De-identified data were entered into Microsoft Excel 2010 and analyzed using Stata 14. Bivariate logistic regression was performed to evaluate the relationship between each of these variables and ADH concentration. Multivariate logistic regression models were constructed to estimate the association of all variables and ADH concentration.

Results

From January to August 2017, a total of 30 head injury cases with alcohol intoxication were registered at the hospital; all used motorized two-wheelers and were males (100%; n = 30). The majority of them were aged <17 years (90%; n = 30) and had mild head injury (GCS score, 14–15; 50%), followed by moderate (GCS score, 9–13; 43%) and severe head injury (GCS score, 3–8; 7%). Patients had craniomaxillofacial lesion (63.3%; n = 19), intracranial lesion (23.3%; n = 7), spinal cord lesion (3.4%; n = 1), and multiple lesions (10%; n = 3). Improvement in final outcome was observed in 86.7% of patients (GCS score, 14/15 (GCS score, 9–13; 43%) and severe head) and 13.3% of patients died (n = 4). Among patients with TBI associated with motorcycle crashes, less than half were wearing a helmet at the time of the crash (46.7%; n = 14). Patients with length of stay patients died (n = 4). A% (n = 11) and mandth of stay patients % (n = 4), and 30% (n = 15) had forced discharge (Table 1).

Table 1: Characteristics of patients with or without intoxicated TBI

| Characteristic | Intoxicated (n=30) | Not intoxicated (n=10) |
|---------------|-------------------|-----------------------|
| Age           |                   |                       |
| <17 years old | 19                | 10                    |
| ≥17 years old | 11                | 0                     |
| Gender        |                   |                       |
| Man           | 30                | 9                     |
| Women         | 0                 | 1                     |
| Initial GCS   |                   |                       |
| Mild (14–15)  | 15                | 8                     |
| Moderate (9–13)| 13               | 2                     |
| Severe (>8)   | 2                 | 0                     |
| Type of lesion|                   |                       |
| Cranio-maxillo-facial* | 19 | 7                   |
| Intracranial* | 7                 | 0                     |
| Spinal cord injury | 1    | 0                    |
| Multiple†     | 3                 | 3                     |
| Outcome       |                   |                       |
| Improvement (GCS 14–15) | 26 | 10                  |
| Death         | 4                 | 0                     |
| No improvement| 0                 | 0                     |
| Protection    |                   |                       |
| Using helmet  | 14                | 7                     |
| No helmet     | 16                | 3                     |
| Length of stay|                   |                       |
| <1 week       | 11                | 9                     |
| >1 weeks      | 4                 | 1                     |
| Forced discharge | 15          | 0                    |

*Bone fracture; laceration; skull base fracture; †Epidural-, subdural-, sub arachnoid-, extradural- hemorrhage, extradural- contusion; ‡a+/b+other extracranial lesion. TBI: Traumatic brain injury, GCS: Glasgow coma scale.

Of all the ten head injury cases without alcohol intoxication, included as the control group, the majority of them also used motorized two-wheelers and were males (90%; n = 9). All were aged <17 years (100%) and had mild head injury (GCS score 143.3%; n = 7), spinal cord, contusion (GCS score 3cases without ao severe head injury (GCS score 3the control group, the majority of them also used motorized two-wheelers and were males (90%; n = 9). All were aged <17 years (100%) and had mild head injury (GCS score 143.3%). All patients (100%) showed improvement in the final outcome (GCS score 14–15; n = 10). Among patients with TBI associated with motorcycle crashes, 70% (n = 7) of them were wearing a helmet at the time of the crash. Patients with length of stay s with TBI associated with motorcycle crashes, less than half were wearing a helmet at the time of the crash (46.7%; n = 14). Patients with length of stay patients died (n = 4). A% (n = 11) and mandth of stay patients % (n = 4), and 30% (n = 15) had forced discharge (Table 1).

Regarding ADH concentration in the trauma patients, all the 30 patients with alcohol intoxication had an average of 17.17 ing ADHng/ml (range 4.93
Drinking and driving have been pointed out as one of the determining factors for the occurrence of crashes. Faced with this problem, the Republic of Indonesia the determining factors concerning Road Traffic and Transportation in Article 106 of the Law in Paragraph 1 states that everyone who drives a motor vehicle on the road must drive their vehicle in alert and full concentration. The explanation of the article and this paragraph states that the meaning of led the risk of involvement in a fatal crash more than (%) had multiple lesions 80%), followed by monob distracted due to illness, fatigue, sleepiness, using a telephone or watching television or videos installed in a vehicle, or drinking liquid containing alcohol or drugs that would affect the ability to drive the vehicle. The legal blood alcohol limits and breath content limits in several countries around the world are typically 0.08 g/dl.

Alcohol test is generally performed by measuring the alcohol levels in blood, breath, or urine. Till date, various types of tools have been developed to detect alcohol levels, ranging from simple to complex ones. Vukovic et al. stated that factual BACs and related expert toxicological decision based solely on the results obtained from breath analyzers might not be correct [13]; however, in an emergency setting, it is difficult to perform a breath analyzer test because most of the patients with TBI cannot obey the command and/or have a decreased level of consciousness. Individuals with a low activity of ADH (alcohol-metabolizing enzymes) have low tolerance to alcohol and are therefore rapidly intoxicated [9]; in other words, they are more vulnerable to more serious injuries from motor vehicle crashes. In the present study, the average ADH concentration in the control group was 41.57 ± 7.66 ng/ml, whereas it was 17.17 ± 8.45 ng/ml in the intoxication group. This result is consistent with a previous study showing that the normal ADH concentration in plasma or serum is 59 um ng/ml based on ELISA [14] and ADH has been reported to be low in alcoholics, it low activity contribute to the reduced first pass ethanol metabolism associated with raised ethanol blood concentrations in these people. [15].

Alcohol contributes significantly to mortality and morbidity in trauma patients, despite the type of the injury [4]. Most of our patients with TBI (64.5%) were under alcohol intoxication, whereas only falls contributed to the mild head injury category. However, we speculated that there is a difference in alcohol consumption due to the different culture of Asian and Western youth, as the levels of ethanol have been found to be high in victims of road traffic crashes in Western countries, associated with high mortality rates, compared with Asian youth who could not afford legal alcohol consumption and often turn to bootleg alcohol or local moonshine or primarily alcohol containing methanol that is locally known as that locally rily alcohol.
Our results demonstrated significantly negative correlation between ADH concentration and initial GCS score; thus, we conclude that patients with TBI with a history of alcohol intoxication and an ADH concentration ≤152 centng/ml have a relative influence of lower initial GCS scores by as much as 6.53 times (p = 0.000). However, in fact, this prediction might vary for each case, as there are several influencing factors such as the type of alcohol (true ethanol or rmiras oplasannaploasantc, this , this prediction might v the period between alcohol consumption and admission to the hospital. There was also a significantly negative correlation between ADH concentration and the type of lesion; the lower the ADH concentration, the more likely the patient has a 3.98 times probability of multiple lesions (p = 0.029). In addition, the correlation between ADH concentration and the outcome was significantly negative, namely, improvement, no improvement, or death. The lower the ADH concentration, the more likely the patient has a 5.27 times probability of worse outcomes (p = 0.003). In this study, we did not find a significant correlation between ADH concentration and length of stay, that is, ADH concentration did not affect the patient’s length of stay (p = 0.392).

However, this study has several limitations, namely, we don’t rates, data that intoxicated TBI patients who come to RSHS (whether they were chronic alcoholics, regular drinker or just a beginner), making it difficult for us to conclude the lower level of ADH. There were also “protection against alcoholism” phenomenon in Asia people (Mongoloid); the final outcome assessment of this study did not completely describe the characteristics because most of the patients decided for forced discharged due to financial problems. Follow-up is required after the patient was forced discharged. A study is already ongoing to increase the number of samples and variations in the type of lesions, the existence of racial differences in alcohol sensitivity and also to compare with more practical BAC detection devices, such as a rapid alcohol test.

Conclusions

ADH concentrations ≤152 centng/ml patients with TBI with a history of alcohol intoxication statistically correlated with lower initial GCS scores on admission by about 6.53 times (p = 0.000), which increased the risk of multiple lesions by 3.98 times (p = 0.029) and worse final outcomes by 5.27 times (p = 0.003). ADH concentrations in intoxicated TBI patients can be taken into consideration to decide about the therapeutic action and further informed consent to the family or as a legal information to the law-enforcement authority.

Ethical Statement

This study was approved by Faculty of Medicine, Universitas Padjadjaran ethics committee and has been performed in according ethical standards laid down Helsinki Declaration 1964.

Authors Contributions

All authors had examined, treated, observed, and follow-up the patients. All authors participated in writing the manuscript, had read, and approved the final manuscript.

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