Association of Tranexamic Acid with GCS in Patients with Traumatic Brain Injury

Kamil Ziad a*, Inayat Ali Khan b*, Lal Shehbaz a*, Santosh Kumar Sidhwani c*, Bina Fawad d* and Ariba Fahim e,

a Emergency Medicine, Ziauddin Hospital North Nazimabad, Pakistan.
b Department of Neurosurgery, Ziauddin Hospital North Nazimabad, Pakistan.
c Department of Pathology, Ziauddin University, Pakistan.
d Community Health Sciences, Ziauddin University, Pakistan.
e Ziauddin Hospital North Nazimabad, Pakistan.

Authors’ contributions

This work was carried out in collaboration among all authors. All authors read and approved the final manuscript.

Article Information

DOI: 10.9734/JPRI/2022/v34i37A36200

Open Peer Review History:

This journal follows the Advanced Open Peer Review policy. Identity of the Reviewers, Editor(s) and additional Reviewers, peer review comments, different versions of the manuscript, comments of the editors, etc are available here: https://www.sdiarticle5.com/review-history/78549

Received 03 October 2021
Accepted 08 December 2021
Published 10 May 2022

ABSTRACT

Background: Traumatic Brain Injury (TBI) occurs due to head injury ranging from simple blow to penetrating injury. TBI refers to a violently produced brain damage that is either blunting or piercing that accelerates or decelerates, resulting in the development of various clinical signs and symptoms such as loss of consciousness or lowered awareness level, entire or partial loss of memory, and various neurological or neuropsychological abnormalities, all of which can lead to disability or death. As a result, antifibrinolytic drugs like tranexamic acid (TXA) may help to prevent traumatic intra-cranial hemorrhage. The aim of current study was to determine the effect of Tranexamic acid on Glasgow Coma Scale score in TBI patients

Methodology: In this Quasi experimental type of study the 126 patients were recruited from the emergency department of Ziauddin University and Hospital North site who were presented with the traumatic brain injury. Confirmed patients of TBI were distributed into two groups; Tranexamic acid Group: A group of TBI patient received 1 gm. of tranexamic acid infused over ten minutes within

*Corresponding author: E-mail: rajperkamil@gmail.com;
three hours of injury along with standard treatment and Placebo Group: A group of TBI patient only received standard treatment.

Results: The mean age of the study subjects was found to be 45.6±18.7 ranged between 18 to 89 years. While assessing the clinical parameters mean temperature recorded is 36.6 °C, with mean respiratory rate and pulse rate of 20.3 and 91.6 respectively. The comparative analysis of the two treatment groups in which one group got the TXA and other had the usual recommended drugs only with the effectiveness of TXA in term of improvement of GCS score showed highly significant statistical association.

Conclusion: Use of Trenaxamic acid (TXA) helps in improvement of GCS at many intervals of 6 hours, 12 hours and 24 hours as compared to those who received conventional treatment.

Keywords: Tranexamic acid; GCS; traumatic; hematoma; injury.

1. INTRODUCTION

Traumatic Brain Injury (TBI) occurs due to head injury ranging from simple blow to penetrating injury [1]. TBI refers to a violently produced brain damage that is either blunt or piercing, or accelerates or decelerates, resulting in the development of various clinical signs and symptoms such as loss of consciousness or lowered awareness level, entire or partial loss of memory, and various neurological or neuropsychological abnormalities, all of which can lead to disability or death. TBI is regarded as a major public health issue that not only affects the global population but also causes socioeconomic issues [2-5]. TBI incidence is increasing annually, according to the World Health Organization (WHO), as the rate of disability and mortality. TBI can result in permanent or temporary brain impairment, limiting social and practical tasks and lowering quality of life [6]. TBI that lasts for a long time might lead to depression and other chronic illnesses. A recent study on global incidence of traumatic brain injury by Dewan MC, et al. reported that approximately 69 million cases of TBI are reported annually around the world of which road traffic accidents (RTAs) and falls are the most commonly reported [7]. Prevalence of RTAs was high in South East Asia (56%) and Africa (56%) and RTAs was low in North America (25%) [8]. Incidence of road traffic injuries (RTI) were high 1.5% in Southeast Asia followed by Europe with incidence of 1.2%. Overall global incidence of TBI was 939 cases in 0.1 million people (69 million/year), whereas incidence of mild TBI was 740 cases in 0.1 million people (55.9 million/year) and the incidence of severe TBI was 73 cases in 0.1 million people (5.48 million/year) [9]. 274,436 patients in emergency department of different hospital of Pakistan and reported traumatic brain injury in 12,125 (4.4%) patients [10]. Intracranial bleeding is one of the most common and severe complication of brain injury that not only increases the risk of disability but also increases the risk of mortality [11]. Bleeding can start immediately after injury and can persists for several hours. Intracranial bleeding increases the intracranial pressure leading towards brain herniation and ultimately death [12]. An initial coma score evaluation is done using the Glasgow Coma Scale (GCS) which was developed in the year 1974 by two neurosurgeons at the University of Glasgow (Teasdale and Jannet) [13]. It is the most commonly used tool for assessment of consciousness. GCS was categorized into three parameters for evaluation of consciousness including; eye, verbal and motor response. GCS has a score ranging from 3-15, three for worst and 15 for best. TBI is categorized into following three types on the basis of GCS score; mild TBI with GCS score 14-15, moderate TBI with GCS score 9-13 and severe TBI with GCS score 3-8 [14-16].

2. METHODOLOGY

The aim of current study was to determine the effect of Tranexamic acid on GCS score in TBI patients. In this Quasi experimental type of study the 126 patients were recruited from the emergency department of Ziauddin University Hospital North site who were presented with the traumatic brain injury. All the adult patients aged 18 and above between GCS 3 and 12 with evident of intracranial hemorrhage were included by using consecutive sampling technique. Demographic details (such as name, gender, age), presenting complains and medical history of each patient was obtained from patient’s family. Each patient was triaged, evaluated for vital signs including temperature, pulse, BP and RR. In other words a primary survey followed by a secondary survey was performed. The GCS was evaluated, patient was stabilized. The
patient underwent an urgent CT scan brain.

Confirmed patients of TBI were distributed into following two groups:

1. **Tranexamic acid Group**: A group of TBI patient received 1 gm. of tranexamic acid infused over ten minutes within three hours of injury along with standard treatment.

2. **Placebo Group**: A group of TBI patient only received standard treatment.

Patients of both groups were evaluated for GCS at 6 hours, 12 hours and then twenty-four hours after administration of tranexamic acid and standard treatment. There was no risk involved in this study as well as no any risk of drug reactions. Adverse effects and as far as management of allergic responses of the drug is concerned, such patients were treated with immediate discontinuation of the drug, followed by applying the anaphylaxis protocol like starting steroids, anti-allergic medications, strict monitoring of vitals, securing the airway if needed, fluids to build up the patient’s hydration.

### 3. RESULTS

The mean age of the study subjects was found to be 45.6±18.7 ranged between 18 to 89 years. While assessing the clinical parameters mean temp recorded is 36.6 °C, with mean respiratory rate and pulse rate of 20.3 and 91.6 respectively. Mean systolic blood pressure recorded was 143.3mmHg and systolic blood pressure was 84.9mmHg. Mean GCS at time of presentation in emergency department, after 6 hour, 12 hours and 24 hours are calculated as 10.71, 10.78, 10.95 and 11.06 respectively that showed the significant improvement of GCS with time Table 1.

The study included 91 (72.2 percent) males and 35 (27.8%) females. For statistical purposes, we split the patients into three age groups: 18 to 40 years, 41 to 60 years, and > 60 years, and found that the majority of the patients were in the 41 to 60 year age group (52/126: 41.3 percent), followed by the 18 to 40 year age group. We also discovered that a high number of patients (70/126: 55.6%) had TBI as a result of road traffic accidents (RTA) and unintentional falls (54/126: 42.9%), while the rest had TBI as a result of other causes such as hit or conflict. We also graded all of the TBI severity based on their clinical characteristics and radiological results. The majority of patients (93/126: 73.8%) had a moderate TBI, while 33/126: 26.2%) had a severe damage. None of them had shown up with a minor injury. Figs. 1, 2 and 3.

The comparative analysis of the two treatment groups in which one group got the txa and other had the usual recommended drugs only with the effectiveness of txa in term of improvement of gcs score showed highly significant statistical association with p value: 0.00 Table 2. Furthermore we also determined the link between the association of the effectiveness with improvement of gcs score after some intervals of 6 hours, 12 hours and 24 hours. Each and every time interval had highly significant association p-values showed in Table 3.

### Table 1. Means of study parameters in all study subjects

<table>
<thead>
<tr>
<th>Parameter</th>
<th>Mean</th>
<th>Minimum</th>
<th>Maximum</th>
</tr>
</thead>
<tbody>
<tr>
<td>Age</td>
<td>45.6 Years</td>
<td>18 Years</td>
<td>89 Years</td>
</tr>
<tr>
<td>Temperature</td>
<td>36.6 °C</td>
<td>36 °C</td>
<td>39 °C</td>
</tr>
<tr>
<td>Systolic BP</td>
<td>143.3mmHg</td>
<td>60mmHg</td>
<td>260mmHg</td>
</tr>
<tr>
<td>Diastolic BP</td>
<td>84.9mmHg</td>
<td>40mmHg</td>
<td>197mmHg</td>
</tr>
<tr>
<td>Respiratory Rate</td>
<td>20.3/Min</td>
<td>12/Min</td>
<td>24/Min</td>
</tr>
<tr>
<td>Pulse Rate</td>
<td>91.6/Min</td>
<td>58/Min</td>
<td>156/Min</td>
</tr>
</tbody>
</table>

### Table 2. Association of effectiveness with treatment groups

<table>
<thead>
<tr>
<th>Effectiveness</th>
<th>Txa</th>
<th>Placebo</th>
<th>P-value</th>
</tr>
</thead>
<tbody>
<tr>
<td>Gcs improved</td>
<td>42</td>
<td>10</td>
<td>0.00*</td>
</tr>
<tr>
<td>Gcs not improved</td>
<td>7</td>
<td>67</td>
<td></td>
</tr>
</tbody>
</table>

*Fisher’s Exact Test
Fig. 1. Frequency of study subjects according to gender

Fig. 2. Frequency of cause of injury

Fig. 3. Severity of traumatic brain injuries
Table 3. Statistical link of effectiveness with GCS at different intervals

<table>
<thead>
<tr>
<th>GCS improvement</th>
<th>GCS Interval</th>
<th>P-value</th>
</tr>
</thead>
<tbody>
<tr>
<td>Effectiveness</td>
<td>At arrival</td>
<td>0.002*</td>
</tr>
<tr>
<td></td>
<td>After 6 hours</td>
<td>0.007*</td>
</tr>
<tr>
<td></td>
<td>After 12 hours</td>
<td>0.006*</td>
</tr>
<tr>
<td></td>
<td>After 24 hours</td>
<td>0.000*</td>
</tr>
</tbody>
</table>

*p independent sample t-test

4. DISCUSSION

TBI is the leading cause of death and disability in the World. TBI-related intracranial hemorrhage (ICH) is linked to a high risk of coagulopathy, which increases the risk of hemorrhage growth and increases the mortality rate [17]. As a result, antifibrinolytic drugs like tranexamic acid (TXA) may help to prevent traumatic ICH. The purpose of this study was to see how much ICH reduced after TXA was administered to TBI patients. Short courses of tranexamic acid (TXA) have been shown to minimize rebleeding in spontaneous intracranial haemorrhage and to reduce bleeding in elective surgery [2]. In TBI patients, the safety of early short-course TXA care was consistent with no increased risk of non-fatal vascular occlusive incidents with early short-course TXA treatment in traumatic bleeding patients. [18]. In present study we also found a significant improvement of GCS score measured at different of 6 hour, 12 hours and 24 hours after presenting in the emergency department. Same findings were also found by Abolfazl et al in their randomized clinical trial in 2017 and others. In present research the mean age was found 45.6 years that is in line with the other studies that found the same results. [10,19] Age remained a mainstay of discussion for long time as some primary traumatic brain injuries are common in old ages and some are more common in youngers. Primary brain injury in our study population is common in middle aged persons followed by young adults due to different causative agents as road traffic accident (RTA) are more evident in this age group and in our analysis majority of the study participants had TBI secondary to RTA and fall. Male were predominant in present study as compared to female due to different exposure of risk factors and trends in our population. RTAs are more evident and common in males in our areas on account of less number of female drivers [20]. In current study we found that most of patients presented with TBI have GCS between 10 to 11 irrespective of cause and age. In many researches have different GCS scores that may be due to many factors like old age have poor GCS because of low understanding of language and obeying command [21]. The Glasgow Coma Scale (GCS), a scoring system never intended to classify brain injury per se but rather level of consciousness, has historically been used to classify traumatic brain injury as mild, moderate, or extreme [13] While the GCS can be helpful in the clinical management and prognosis of TBI, it "does not include clear details about the pathophysiologic processes that are responsible for the neurologic defects and targeted by therapies," according to a TBI consensus workgroup. We also found that use of TXA or fibrinolytics in TBI helps in improvement of patient in term of consciousness (GCS). Many researches support this as they have positive results after use of fibrinolytic agents [22,23]. While fibrinolytic assays are not commonly used to diagnose posttraumatic coagulopathy, circulating biomarkers such as D-dimer and fibrin degradation products have shown promise in predicting outcome. Early tranexamic acid administration in patients with TBI and reported hyperfibrinolysis has been suggested as a way to enhance clinical results even further [22]. The current research supports the use of tranexamic acid as an empiric treatment for severe trauma haemorrhage.

5. CONCLUSION

Traumatic brain injury is more evident in male at second decade of life. RTA followed by fall is the more common cause. Patients presented in emergency department have moderate traumatic brain injury with mean GCS of 10. Severe traumatic brain injury is also reported with low GCS. All of the patients have associated intracerebral hemorrhage in both cases. Use of Trenaxamic acid (TXA) helps in improvement of GCS at many intervals of 6 hours, 12 hours and 24 hours as compared to those who received conventional treatment.

CONSENT

Verbal and written informed consent was obtained from all patients.
ETHICAL APPROVAL

Ethical approval was taken from Ethics Review Committee (ERC) of Ziauddin University Karachi accordance with institutional guidelines (3070121KZEM).

COMPETING INTERESTS

Authors have declared that no competing interests exist.

REFERENCES


Peer-review history:
The peer review history for this paper can be accessed here:
https://www.sdiarticle5.com/review-history/78549