Research Article

Correlation between Coagulation Fibrinolysis Function and Outcomes during Hospitalization in Patients with Severe Traumatic Hemorrhagic Shock

Louwei Zhang, Maosheng Lin, Xuhua Tang, and Yejiang Tang

Department of Emergency Medicine, Zhuji Affiliated Hospital of Shaoxing University, Zhuji 311800, Zhejiang, China

Correspondence should be addressed to Louwei Zhang; zhanglouwei@163.com

Received 25 April 2022; Accepted 9 June 2022; Published 30 June 2022

Objective. To analyze the correlation between coagulation fibrinolysis function and outcomes during hospitalization in patients with severe traumatic hemorrhagic shock.

Methods. A retrospective collection was performed on the clinical data of 106 patients with severe traumatic shock admitted to the hospital between January 2020 and January 2022. According to the injury severity score (ISS), they were divided into the S1 group (ISS < 25 points, n = 70) and the S2 group (ISS ≥25 points, n = 36). Prothrombin time (PT), fibrinogen (Fib), thrombin time (TT), and activated partial thromboplastin time (APTT) were detected by the coagulation assay. D-dimer (D-D) was detected by an enzyme-linked immunosorbent assay. Antithrombin activity (AT: A) and plasminogen activity (PLG: A) were detected by the chromogenic substrate method. The relationship between coagulation fibrinolysis indexes and injury severity was analyzed by Spearman’s correlation analysis. The predictive value of coagulation fibrinolysis indexes for outcomes of patients with severe traumatic hemorrhagic shock was evaluated by receiver operating characteristic (ROC) curves.

Results. The levels of PT, APTT, D-D, TT, AT: A and PLG: A in the S2 group were higher than those in S1 group, while the Fib level was lower than that in the S1 group (P < 0.05). A Spearman’s analysis showed that PT, APTT, TT, D-D, AT: A, and PLG: A were positively correlated with injury severity (P < 0.05), while Fib was negatively correlated with it (P < 0.05). Among the 106 patients, there were 89 survived cases and 17 died cases. The levels of PT, APTT, D-D, AT: A and PLG: A in the death group were lower than those in the survival group, while the Fib level was higher than that in the survival group. The results of ROC curve analysis showed that serum PT, APTT, Fib, TT and D-D were of predictive value for outcomes (AUC = 0.713, AUC = 0.763, AUC = 0.712, AUC = 0.761, AUC = 0.730, AUC = 0.765, AUC = 0.673, P < 0.05), and cutoff values were 20.29 s, 34.79 s, 3.54 g/L, 20.97 s, 1.42 μg/L, 73.53% and 63.97%, respectively. Conclusion. There is coagulation and fibrinolysis dysfunction in patients with severe traumatic hemorrhagic shock, which is related to injury severity. The coagulation fibrinolysis indexes have a certain predictive value for outcomes of patients.

1. Introduction

Traumatic shock is a combination of factors such as severe violent blows on the body, major organ damage, and massive hemorrhage. The body’s decompensation syndrome is formed. Therefore, the etiology and pathology of traumatic shock are more complicated than simple hemorrhagic shock. Severe traumatic hemorrhagic shock occurs when the patient bleeds a lot, blood volume decreases, and local tissue necrosis occurs due to severe impact by external force in a short time [1]. The main clinical presentations are severe pain, shortness of breath, pale face, weakened jugular vein pulse, and confusion. Due to insufficient systemic tissue perfusion, metabolic disorders, impaired cell ischemia, hypoxia, and severe traumatic hemorrhagic shock, it is prone to cause damage to multiple organs [2, 3]. The condition of patients is complex and in danger, with many early complications and high mortality, which makes it a severe illness in the hospital. Effective treatment should be given to reduce the complications and improve the patient’s prognosis. The pathophysiological process of traumatic ischemic shock is complex, and the coagulation chain reaction that can be
triggered by shock plays an important role in the disease [4]. Results of some studies have shown [5] that when the patient suffers severe liver damage, the incidence of consumptive coagulopathy will reach as high as 50%. However, the impact of the shock on the coagulation and fibrinolysis systems of patients with traumatic blood loss and the relationship between shock and patient outcomes remains controversial [6, 7]. The bodies of patients with shock are in a hypercoagulable state and are prone to suffering thrombosis as the high-intensity injury damages the repair function of the body seriously and the coagulation and fibrinolysis function are gradually disturbed [8]. If severe traumatic hemorrhagic shock onsets rapidly and the treatment is not timely, the risk of death will be greatly increased. This study analyzed the changes in coagulation and fibrinolysis in patients with severe traumatic shock and the association between the changes and the outcome in order to provide guidance for the clinical.

2. Clinical Data and Methods

2.1. General Information. The clinical data of 106 patients with severe trauma shock admitted to our hospital from January 2020 to January 2022 were retrospectively selected. Patients were divided into S1 group (ISS score < 25, n = 70) and S2 group (ISS score ≥ 25, n = 36) based on the severity of trauma (ISS). Group S1 had 38 males and 32 females, aged 12–70 years, with an average age of 49.64 ± 9.78 years. Causes of trauma were traffic accident injury in 15 cases, fall injury in 22 cases, traffic accident injury in 16 cases, and knife stabbing injury in 17 cases; trauma site: brain injury in 12 cases, chest injury in 23 cases, and abdominal injury was the main injury in 17 cases and 18 cases were mainly spinal and limb injuries. In the S2 group, there were 20 males and 16 females, aged 15–70 years, with an average age of 51.21 ± 9.36 years. Causes of trauma: 9 cases of traffic accidents, 12 cases of falling from height, 7 cases of traffic accidents, and 8 cases of knife stabbing; trauma site: brain injury in 5 cases, chest injury in 14 cases, and abdominal injury in 14 cases out of which 9 cases were mainly sprains and 8 cases were mainly spinal and limb injuries. Inclusion criteria were meeting the diagnostic criteria for severe traumatic hemorrhagic shock [9]; ISS score ≥16 points on admission; survival time after rescue ≥24 hours. Exclusion criteria were patients with liver and kidney dysfunction; patients with mental disorders; and patients with missing clinical data. There was no significant difference in age, gender, trauma cause, and trauma site between the S1 and S2 groups (P > 0.05). This study has been approved by the Medical Ethics Committee of our hospital.

2.2. Detection of Serum Coagulation and Fibrinolysis Indexes. The next day, after the patients were enrolled, 5 mL of peripheral venous blood was collected in the fasting state, and the supernatant was collected after centrifugation (1200 g, 8 cm, 5 min). Prothrombin time (PT), fibrinogen (Fib), thrombin time (TT), and activated partial thromboplastin time (APTT) were detected by coagulation method. D-dimer (D-dimer, D-D) was detected by an enzyme-linked immunofluorescence assay. The instruments and related kits were purchased from Beijing Boaosen Biotechnology Co., Ltd. The antithrombin activity (AT : A) and plasminogen activity (PLG : A) were detected by the chromogenic substrate method. The kits were purchased from Wuhan Purity Biotechnology Co., Ltd., and the operations were carried out in strict accordance with the kit instructions.

2.3. Statistical Processing. The SPSS 22.0 software was used for statistical analysis. All experimental data conformed to normal distribution, and measurement data were presented as mean ± standard deviation (X ± S), and the two-sample independent t-test was used to compare the differences between groups. The enumeration data were expressed as [cases (%)], and the difference between the two groups was compared by the χ2 test. Spearman’s correlation was used to analyze the relationship between the coagulation and fibrinolysis indexes and the severity of the injury. The receiver operator characteristic curve (ROC) was used to measure the predictive effect of coagulation and fibrinolysis indexes in patients with severe traumatic hemorrhagic shock. The test level was α = 0.05, and P < 0.05 indicated that the data had a significant statistical difference.

3. Result

3.1. Comparison of Serum Coagulation and Fibrinolysis Indexes in Each Group. The levels of PT, APTT, D-D, TT, AT : A, and PLG : A in group S2 were higher than those in group S1, the level of Fib was lower than that in group S1, and the difference was statistically significant (P < 0.05), as shown in Table 1 and Figure 1.

3.2. Correlation Analysis between the Severity of Injury and Coagulation and Fibrinolysis Indexes in Patients. The result of Spearman analysis showed that PT, APTT, TT, D-D, AT : A, and PLG : A were positively correlated with injury severity (P < 0.05), and Fib was negatively correlated with injury severity (P < 0.05). As shown in Table 2 and Figure 2.

3.3. Comparison of Serum Coagulation and Fibrinolysis Indexes between Death Group and Survival Group. The patient’s survival time within three months was recorded. Among the 106 patients, 89 survived and 17 died. The levels of PT, APTT, D-D, AT : A, and PLG : A in the death group were lower than those in the survival group, and the Fib level was higher than that in the survival group (P < 0.05). As shown in Table 3.

3.4. Predictive Value of Coagulation and Fibrinolysis Indexes on Patient Outcome. The ROC results show that serum PT, APTT, Fib, TT, D-D, AT : A and PLG : A could all predict the outcome of patients (AUC = 0.713, AUC = 0.683, AUC = 0.712, AUC = 0.761, AUC = 0.730, AUC = 0.765, AUC = 0.673, P < 0.05). At this time, the cutoff values
The coagulation and fibrinolysis systems are in a state of dynamic equilibrium under physiological conditions. When the body is injured, the balance system is disrupted, and the coagulation and fibrinolysis systems are activated [12]. In this study, the levels of PT, APTT, D-D, TT, AT: A, and PLG: A in group S2 were higher than those in group S1, and the level of Fib was lower than that in group S1. D-D is a specific product after the degradation of cross-linked fibrin, which mainly reflects the function of fibrinolysis. The increase in its concentration indicates that the fibrinolytic system and the coagulation system are activated [13]. Fib is a protein with a coagulation function that is synthesized by the liver [14]. PT mainly reflects whether the extrinsic coagulation is normal or not and reflects the content of plasma factors II, V, VII, and X, and its prolongation is seen in congenital coagulation factor deficiency [15]. TT refers to the blood coagulation time after adding standardized prothrombin to plasma, which reflects the anticoagulant substances in the body, and its prolongation indicates hyperfibrinolysis [5]. During traumatic hemorrhagic shock, the coagulation and fibrinolysis systems, as the main systems involved in stress and hemostasis, are activated urgently and play an important role in the occurrence and development of shock. Its changes within a certain range have a positive effect on hemostasis and antishock [16]. However, if the factors that induce the coagulation and fibrinolysis system are excessive or continuously stimulated, it may increase the degree of shock. This study shows that the balance of coagulation and

### Table 1: Comparison of serum coagulation and fibrinolysis indexes in each group (X ± S).

| Group     | Number of cases | PT (s)       | APTT (s)     | Fib (g/L) | TT (s)       | D-D (mg/L) | AT: A (%) | PLG: A (%) |
|-----------|-----------------|--------------|--------------|-----------|--------------|------------|-----------|------------|
| Group S1  | 70              | 17.54 ± 1.82 | 29.81 ± 3.04 | 4.41 ± 0.57 | 17.25 ± 1.87 | 1.03 ± 0.12 | 76.13 ± 7.74 | 77.33 ± 7.82 |
| Group S2  | 36              | 21.46 ± 2.33 | 35.72 ± 3.61 | 3.28 ± 0.39 | 23.86 ± 2.46 | 1.44 ± 0.15 | 72.58 ± 7.36 | 61.85 ± 6.32 |

| t value   | P value          |
|-----------|------------------|
| 9.527     | <0.001           |
| 8.886     | <0.001           |
| 10.668    | <0.001           |
| 15.441    | <0.001           |
| 15.276    | <0.001           |
| 2.273     | <0.001           |
| 10.270    | <0.001           |

**Figure 1:** Comparison of serum coagulation and fibrinolysis indexes in each group. PT: prothrombin time, APTT: activated partial thromboplastin time, Fib: fibrinogen, TT: thrombin time, D-D: D-dimer, AT: A: antithrombin activity, and PLG: A: plasminogen activity; *P < 0.05 compared with the S1 group.

**Figure 2:** Comparison of serum coagulation and fibrinolysis indexes in each group. PT: prothrombin time, APTT: activated partial thromboplastin time, Fib: fibrinogen, TT: thrombin time, D-D: D-dimer, AT: A: antithrombin activity, and PLG: A: plasminogen activity; *P < 0.05 compared with the survival group.

### Table 2: Correlation analysis between the severity of injury and coagulation and fibrinolysis indexes in patients.

| Indicator | r     | P     |
|-----------|-------|-------|
| PT        | 0.542 | 0.036 |
| APTT      | 0.611 | 0.027 |
| TT        | 0.654 | 0.041 |
| Fib       | −0.586| 0.003 |
| D-D       | 0.637 | 0.017 |
| AT: A     | 0.589 | 0.042 |
| PLG: A    | 0.609 | 0.038 |

**4. Discussion**

The main pathophysiological change of traumatic hemorrhagic shock is the mismatch between blood volume and vascular volume, resulting in insufficient perfusion of peripheral tissues and inducing changes in microcirculation, abnormal oxygen metabolism, inflammatory response, coagulation disorders, and secondary damage to organs [10, 11]. The disease has the characteristics of a short-term onset and high mortality. Therefore, clinicians should take timely intervention measures to save the patient’s life as much as possible.
fibrinolysis in patients with traumatic hemorrhagic modification is disrupted, and the dysfunction of coagulation and fibrinolysis becomes more serious with the aggravation of the disease. The results of Spearman’s correlation analysis showed that PT, APTT, TT, D-D, AT: A, and PLG: A were positively correlated with injury severity, and Fib was negatively correlated with injury severity. Previous studies have shown [17] that the more severe the injury is in patients with traumatic hemorrhage, the more likely they are to have coagulation and fibrinolysis dysfunction, which coincides with the results of this study. The results indicate that the coagulation system is continuously activated in the acute stage of severe trauma, the patient’s blood is in a state of hypercoagulation and high viscosity, and the risk of intra-vascular coagulation and thrombosis is greatly increased. Therefore, at this time, the blood volume of the patient should be replenished in a timely and effective manner, which can help the patient return to a normal physiological state and gradually seek medical treatment for the pathological process. At the same time, during the treatment process, attention should be paid to the coagulation and fibrinolysis function status of patients, and interventions such as early coagulation substrates should be given. At the same time, this study also found that among the 106 patients,

### Table 3: Serum coagulation and fibrinolysis indexes of death group and survival group (X±S).

| Group          | Number of cases | PT (s)        | APTT (s)      | Fib (g/L) | TT (s)        | D- (μg/L) | AT: A (%)     | PLG: A (%)     |
|----------------|-----------------|---------------|---------------|-----------|---------------|-----------|---------------|---------------|
| Survival group | 89              | 18.06±1.96    | 30.86±3.12    | 4.59±0.51 | 18.63±2.03    | 1.08±0.12 | 75.89±7.62    | 74.58±7.68    |
| Death group    | 17              | 22.85±2.35    | 36.87±3.74    | 3.20±0.34 | 24.18±2.62    | 1.59±0.16 | 69.84±7.14    | 58.64±6.23    |
| t value        |                 | 8.927         | 7.045         | 10.767    | 9.837         | 15.174    | 3.028         | 8.056         |
| P value        |                 | <0.001        | <0.001        | <0.001    | <0.001        | 0.003     | <0.001        |

![ROC curve](image.png)

**Figure 3:** Predictive effect of coagulation and fibrinolysis indexes on patient outcomes.

### Table 4: Predictive effect of coagulation and fibrinolysis indexes on patient outcome.

| Indicator | AUC   | Standard error | Sensitivity (%) | Specificity (%) | Cutoff value | 95% CI          | P     |
|-----------|-------|----------------|-----------------|-----------------|--------------|----------------|-------|
| PT        | 0.713 | 0.094          | 70.59           | 79.78           | 20.29 (s)    | 0.617–0.797    | <0.05 |
| APTT      | 0.683 | 0.081          | 64.71           | 80.90           | 34.79 (s)    | 0.586–0.700    | <0.05 |
| Fib       | 0.712 | 0.088          | 76.47           | 78.65           | 3.54 (g/L)   | 0.616–0.796    | <0.05 |
| TT        | 0.761 | 0.069          | 76.47           | 69.66           | 20.97 (s)    | 0.669–0.839    | <0.05 |
| D-D       | 0.730 | 0.080          | 70.59           | 84.27           | 1.42 (μg/L)  | 0.635–0.811    | <0.05 |
| AT: A     | 0.765 | 0.074          | 76.47           | 75.28           | 73.53 (%)    | 0.672–0.842    | <0.05 |
| PLG: A    | 0.673 | 0.072          | 70.59           | 65.17           | 63.97 (%)    | 0.575–0.761    | <0.05 |
89 survived and 17 died. The levels of PT, APTT, D-D, AT: A, and PLG: A in the death group were lower than those in the survival group, and the Fib level was higher than that in the survival group, indicating that the coagulation and fibrinolysis indexes of the patients in the death group were significantly abnormal. Analysis of ROC results showed that serum PT, APTT, D-D, AT: A, and PLG: A all had a certain value in predicting patient outcomes. Previous studies have shown [18] that the abnormality of coagulation and fibrinolysis in patients with traumatic brain injury in the early post-injury period is related to the severity of the traumatic brain injury and has a certain evaluation value for the patient’s condition and prognosis, which is basically consistent with this study. Therefore, coagulation and fibrinolysis dysfunction after traumatic blood loss is an important reason for the aggravation and death of shock patients. Paying attention to the changes in coagulation and fibrinolysis indexes in shock patients caused by traumatic blood loss and early intervention are of great significance in reducing the mortality of patients.

In conclusion, the blood of patients with severe traumatic hemorrhagic shock is in a hypercoagulable state, with continuous activation of the coagulation system and hyperfibrinolysis, increasing the risk of secondary thrombosis and bleeding, and the patient’s state increases with the severity of the injury. Coagulation and fibrinolysis indexes are of great significance for assessing the severity of injury and predicting the prognosis of patients. This study has a certain guiding significance for clinical treatment of patients with severe traumatic shock and improvement of prognosis, but there are still shortcomings: the research is still in the preliminary stage, and the specific mechanism of action on the coagulation and fibrinolysis system after traumatic blood loss has not been deeply analyzed. Due to the small number of research samples, there may be errors in the analysis results, and a large number of subsequent clinical data are required for verification.

Data Availability
The data can be obtained from the author upon reasonable request.

Conflicts of Interest
The authors declare that the research was conducted in the absence of any commercial or financial relationships that could be construed as potential conflicts of interest.

References
[1] Y. Jiang, Q. Liang, and H. E. Jin, “Clinical research on dysfunction of coagulation and fibrinolysis in delayed traumatic intracranial hematoma,” Chinese Journal of Neuromedicine, vol. 40, no. 6, pp. 501–506, 2004.
[2] H. B. Moore, E. E. Moore, E. Gonzalez et al., “Hemolysis exacerbates hyperfibrinolysis, whereas plateolysis shuts down fibrinolysis,” Shock, vol. 43, no. 1, pp. 39–46, 2015.
[3] U. Amara, D. Rittirsch, M. Flierl et al., “Interaction between the coagulation and complement system,” Advances in Experimental Medicine and Biology, vol. 632, 2008.
[4] S. Iwamoto, A. Takasu, and T. Sakamoto, “Therapeutic mild hypothermia: effects on coagulopathy and survival in a rat hemorrhagic shock model,” The Journal of Trauma, Injury, Infection, and Critical Care, vol. 68, no. 3, pp. 669–675, 2010.
[5] C. Zentai, D. van der Meijden, T. Braunschweig et al., “Hemostatic therapy using tranexamic acid and coagulation factor concentrates in a model of traumatic liver injury,” Anesthesia and Analgesia, vol. 123, no. 1, pp. 38–48, 2016.
[6] S. Miniello, M. Testini, M. G. Balzanelli, and G. Cristallo, “Turbe della coagulazione nel trauma severo: il ruolo del chirurgo nella prevenzione,” Annali Italiani di Chirurgia, vol. 75, no. 3, pp. 293–297, 2004.
[7] L. S. Gall, K. Brohi, and R. A. Davenport, “Diagnosis and treatment of hyperfibrinolysis in trauma a European perspective,” Seminars in Thrombosis and Hemostasis, vol. 43, no. 2, pp. 224–234, 2017.
[8] Y. C. Guan, “Relationship between coagulation-fibrinolysis system and DIC in patients with traumatic shock,” Applied Journal of General Practice, vol. 33, no. 1, pp. 61–66, 2008.
[9] W. S. Tong, X. U. Jun-Fa, and L. I. Gao-Yi, “Relationship between abnormal blood coagulation-fibrinolysis and progressive intracranial hemorrhage in patients with traumatic brain injury,” Chinese Journal of Clinical Neurosurgery, vol. 26, no. 012, pp. 1029–1038, 2009.
[10] B. J. Eastridge, J. B. Holcomb, and S. Shackelford, “Outcomes of traumatic hemorrhagic shock and the epidemiology of preventable death from injury,” Transfusion, vol. 59, no. S2, pp. 1423–1428, 2019.
[11] C. Pitotti and J. David, “An evidence-based approach to nonoperative management of traumatic hemorrhagic shock in the emergency department,” Emergency Medicine Practice, vol. 22, no. 11, pp. 1–24, 2020.
[12] O. Kloeters, D. Vasilic, P. Hupkens, and D. Ulrich, “Markers of blood coagulation and fibrinolysis in patients with early and delayed microsurgical reconstructions in the lower extremities,” Journal of Plastic Surgery and Hand Surgery, vol. 51, no. 6, pp. 420–426, 2017.
[13] H. Robert-Ebadi and M. Righini, “D-dimer: well beyond diagnosis!” JMV-Journal de Médecine Vasculaire, vol. 45, no. 5, pp. 239-240, 2020.
[14] O. Grottke, S. Mallaiah, K. Karkouti, F. Saner, and T. Haas, “Fibrinogen supplementation and its indications,” Seminars in Thrombosis and Hemostasis, vol. 46, no. 1, pp. 38–49, 2020.
[15] A. Dorgalaleh, E. J. Favaloro, M. Bahraini, and F. Rad, “Standardization of prothrombin time/international normalized ratio (PT/INR),” The International Journal of Hemostasis, vol. 43, no. 1, pp. 21–28, 2021.
[16] H. B. Moore, E. E. Moore, P. J. Lawson et al., “Fibrinolysis shutdown phenotype masks changes in rodent coagulation in tissue injury versus hemorrhagic shock,” Surgery, vol. 158, no. 2, pp. 386–392, 2015.
[17] M. J. Delano, S. B. Rizoli, S. G. Rhind et al., “Prehospital resuscitation of traumatic hemorrhagic shock with hypertonic solutions worsens hypocoagulation and hyperfibrinolysis,” Shock, vol. 44, no. 1, pp. 25–31, 2015.
[18] T. Wada, S. Gando, K. Maekawa et al., “Disseminated intravascular coagulation with increased fibrinolysis during the early phase of isolated traumatic brain injury,” Critical Care, vol. 21, no. 1, pp. 219–235, 2017.