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***Observational Study***

**Factors related to improved** **American Spinal Injury Association grade of acute traumatic spinal cord injury**

Tian C *et al*. ATSCI and current treatments

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**Abstract**

BACKGROUND

Acute traumatic spinal cord injury (ATSCI) usually results in disability, yet data on contemporary national trends of ATSCI incidence are limited.

AIM

To provide a systematic and basic theoretical basis for improving the treatment of acute spinal cord injury.

METHODS

Data from the Peking University Third Hospital Inpatient Sample databases were analyzed. A total of 304 patients with ATSCI were included from 2012 to 2017. The epidemiological data, treatment, complications and clinical outcomes of these patients were reviewed.

RESULTS

Of the 304 patients, 257 (84.5%) were male, and 75% of the patients were 55 years old or younger. 135 patients had improved follow-up American Spinal Injury Association (ASIA) grades (44.4%). Only 14 patients with ASIA grade A improved. A statistically significant difference in prognosis between patients who underwent surgery within 72 h and those who underwent surgery after 72 h was observed (*P* < 0.05). Surgery within 72 h resulted in better prognosis. The Steroid group and the Non-Steroid group showed a significant difference in outcome among patients with ASIA grades A and B (*P* < 0.05). Patients with pneumonia had a poorer prognosis than patients without pneumonia (*P* < 0.05). Surgery within 72 h resulted in better prognosis.

CONCLUSION

This study found that there was no significant difference in hospitalization time and prognosis between the Steroid group and the Non-Steroid group, but the patients with severe spinal cord injury (ASIA grades A and B) who underwent surgery combined with steroid therapy had a better prognosis than those who underwent surgery alone. The disastrous consequences of ATSCI and lack of consensus on the management strategy are obvious. Further improvements in treatment planns are needed in order to obtain more reliable functional outcomes.

**Key Words:** Acute spinal cord injury; Trauma; Surgery; American Spinal Injury Association; Steroids; Prognosis

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**Core** **Tip:** Our study describes the present treatment of acute traumatic spinal cord injury. This study found that surgery within 72 h resulted in a better prognosis. Patients with American Spinal Injury Association (ASIA) grades A and B had a significantly better prognosis when treated with steroids. The present study demonstrates that patients with severe spinal cord injury (ASIA grades A and B) may benefit from early surgery and steroid administration. These findings have implications for future clinical practice.

**INTRODUCTION**

Acute spinal cord injury (ASCI) is a common type of severe trauma in the emergency department, which results in catastrophic consequences for patients. The vast majority of patients are disabled due to severe neurological damage, or can even result in fatalities. This places an enormous financial burden and emotional stress on families and society.

The most recent model systems estimate of 12500 new cases per year was revised recently from 11000 new cases per year based on the increase in the United States population. The number of individuals with acute traumatic spinal cord injury (ATSCI) is increasing, mostly in young and middle-aged men[1]. Patients are paralyzed and even die, causing huge medical and economic burdens on families and society.

Research in this area is extensive, including the continuous improvement in surgical techniques and protocols, the clinical application of hormones, clinical research on drugs to protect and promote nerve growth, and the exploration of emerging stem cell transplantation treatments, *etc.*[2-7] However, there is currently no mature and effective measures that can improve the patient's functional outcome[8,9].

This article reviews the epidemiology and treatment of patients with ATSCI in our hospital over the past 5 years, in order to provide a systematic and basic theoretical basis for improving the treatment of ASCI.

**MATERIALS AND METHODS**

A single-center retrospective cohort study was conducted in an urban tertiary hospital. Patients with ATSCI were recruited from 2012 to 2017 using the following inclusion and exclusion criteria: (1) age > 18 years old; (2) ASCI diagnosed based on American Spinal Injury Association (ASIA) criteria; (3) ASIA grade A to D; and (4) Surgical intervention during hospital stay. Patients with acute cerebrovascular disease were excluded. The patients were followed-up at one year after discharge, the ASIA score was recalculated, and the recovery of neurological function was evaluated according to the ASIA criteria.

Statistical data processing was performed using SPSS 22.0 (IBM Inc*.*, Armonk, NY, United States). Analysis of variance (ANOVA) was used for measurement data. The t-test was used for quantitative data, and the Chi-square test was used for categorical data. *P* < 0.05 was considered statistically significant.

**RESULTS**

***Patient demographics and clinical characteristics***

There were 440 ATSCI patients enrolled in the study. A total of 304 cases were included in the final analysis (Figure 1). Patient demographics are shown in Table 1.

ATSCI was more common among young and middle-aged men. The most common injury site was the cervical spinal cord. Falls and motor vehicle crashes were the two most common causes. The study found that patients with fracture had a worse prognosis than patients without fracture (*P* < 0.05). A shorter time from onset to treatment resulted in a better prognosis (*P* < 0.05).

In addition to preventive antibiotic therapy, other treatments such as proton pump inhibitors (PPIs), nutritional supplements and analgesia were administered. All patients underwent surgical treatment with 133 patients undergoing surgery within 72 h. 208 patients were treated with steroids (Table 2), and 44 patients were treated with high-dose steroids. 243 patients were treated with mannitol.

There was no significant difference in mean hospitalization days or length of intensive care unit stay between the Steroid group and the Non-Steroid group (*P* > 0.05). There were no significant differences in the average number of hospitalization days and prognosis between the high-dose steroid group and the low-dose steroid group (*P* > 0.05).

***Complications***

There was a difference in the prognosis between patients with complications and those without complications (*P* < 0.05). The presence of complications during hospitalization was documented, and included 53 cases of pneumonia, 37 cases of electrolyte disorder, 30 cases of liver injury, 19 cases of tracheotomy, 17 cases of lower extremity deep venous thrombosis, 7 cases of urinary tract infection, 6 cases of stress ulcer, 4 cases of pulmonary embolism, 3 cases of hematoma, 1 case of acute myocardial infarction, 1 case of heart failure, and 1 case of cerebrospinal fluid leakage (Figure 2). The presence of a significant association between common complications and steroid use is shown in Table 3.

Of the 304 patients, 135 patients had improved follow-up ASIA grades (44.4%). Only 14 patients with ASIA grade A improved. There was a statistically significant difference in prognosis between patients who underwent surgery within 72 h and those who underwent surgery after 72 h (*P* < 0.05, Figure 3A). There were significant differences between the Steroid group and the Non-Steroid group only among patients with ASIA grades A and B (*P* < 0.05, Figure 3B). However, the incidence of pneumonia increased with steroid treatment (*P* < 0.05, Table 3). Patients with pneumonia had a worse prognosis than patients without pneumonia (*P* < 0.05, Figure 3C). Additionally, patients with an ASIA grade of A had the worst prognosis compared with other patients (*P* < 0.05, Figure 3D).

**DISCUSSION**

ATSCI severely affects daily life. The incidence of ATSCI is more common in young and middle-aged males. The cervical spine is the most commonly injured site. Frequent causes of injury are falls and traffic accidents. Beck *et al*[10] found that there were 706 cases of ATSCI, most of which were the result of transportation accidents or falls. The incidence of ATSCI resulting from falls increased by 9%/year (95%CI: 4%-15%). In the present study, of 304 patients, 44.4% had improved neurological function. Only 14 patients with ASIA grade A improved. The improvement rate was 17.9%. Of these patients, 5 improved from grade A to grade B, 6 improved from grade A to grade C, and 3 improved from grade A to grade D. None of these patients had normal neurological function at one-year follow up. At present, the guidelines and consensus indicate that surgery is an effective means of treating ASCI and that surgery can alleviate the symptoms of nerve compression. However, there are no standard or effective treatments for improving functional outcomes in these patients.

Our study found that patients with fracture and dislocation had a poorer prognosis than patients without fracture or dislocation, as these patients usually had complications such as spinal cord compression and thus the injury was severe. The shorter the time from onset to treatment (including surgery within 72 h and medication), the better the neurological prognosis might be. Yang *et al*[11] found that high-dose steroid therapy within 8 h and adequate spinal decompression after injury can effectively improve spinal cord function in thoracolumbar spinal injury. Prognosis was closely related to the timely treatment of patients in the early stage of the injury and the reduction of spinal cord compression. In addition, fewer complications indicated a better neurological prognosis. Patients with pneumonia had a worse prognosis. Hou *et al*[12,13] found that ASIA and moderate to severe respiratory complications were variables significantly correlated with tracheostomy. Jaja *et al*[14] found that pneumonia, wound infections, and sepsis (PWS) were the leading causes of acute mortality after ATSCI. Furthermore, as steroids can inhibit inflammation and reduce edema, they were often used as the first-line treatment in ATSCI[15]. However, the American Association of Neurological Surgeons has proposed guidelines issued in 2013 opposing the use of methylprednisolone as a routine treatment for ATSCI[16]. In recent years, an increasing number of clinical studies have shown that steroids have limited effects and benefits in ATSCI, and the use of high doses of steroids may cause serious complications, including gastrointestinal hemorrhage and respiratory tract infection[17-20]. Our study found that the prognosis of patients treated with surgery combined with steroid therapy was not significantly different to that in those treated with surgery alone, while patients with severe spinal cord injury (ASIA grade B patients) had an improved prognosis. However, the incidence of infections caused by steroids may increase.

This study was limited by the long course of spinal cord injury, slow recovery of neurological function, low improvement rate of the disastrous consequences of ATSCI and the current lack of mature treatments. Further improvements in the treatment plans are needed in order to preserve neurological function in patients. In addition, we also need to actively carry out randomized multicenter studies with improved sample size to provide more clinical evidence for the optimal management strategy in ATSCI patients.

**CONCLUSION**

Our study described the current clinical management of ATSCI patients with long-term follow-up and the findings of this study have important implications for future clinical practice. Surgery within 72 h resulted in a better prognosis. It was found that there was no significant difference in hospitalization time and prognosis between the Steroid group and the Non-Steroid group. However, patients with severe spinal cord injury (ASIA grades A and B) who underwent surgery combined with steroid therapy had a better prognosis than those who underwent surgery alone. The disastrous consequences of ATSCI and the lack of consensus on treatment require further study to improve the treatment strategy in order to obtain more reliable functional improvement.

**ARTICLE HIGHLIGHTS**

***Research background***

Acute spinal cord injury is a common type of severe trauma in the emergency department, and can have catastrophic consequences in patients.

***Research motivation***

To describe the present treatment of acute traumatic spinal cord injury (ATSCI) and provide important suggestions for future clinical practice.

***Research objectives***

The objective of our study was to provide a systematic and basic theoretical basis for improving the treatment of ATSCI.

***Research methods***

This retrospective study enrolled patients with ATSCI from 2012 to 2017, and recorded their demographic and clinical characteristics. Patients were followed-up at one year after discharge. The American Spinal Injury Association (ASIA) score was re-calculated, and the recovery of neurological function was evaluated according to ASIA guidelines.

***Research results***

A total of 440 ATSCI patients were enrolled in our study, and 304 of these patients were included in the final analysis. The results of the analysis indicated that ATSCI was more likely to occur in young and middle-aged males and the cervical spinal cord was the most commonly injured site. A statistically significant difference in prognosis was observed between patients who underwent surgery within 72 h and those who underwent surgery after 72 h (*P* < 0.05). A significant difference between the Steroid group and the Non-Steroid group in terms of outcome was noted in the ASIA grade A and B subgroups (*P* < 0.05). In addition, patients with pneumonia had a worse prognosis compared with those without pneumonia (*P* < 0.05).

***Research conclusions***

Our study described the current situation in ATSCI patients with long-term follow-up and the findings in our study have important implications for future clinical practice. Surgery within 72 h results in a better prognosis. This study found that there was no significant difference in hospitalization time and prognosis between the Steroid group and the Non-Steroid group. However, patients with severe spinal cord injury (ASIA grades A and B) who underwent surgery combined with steroid therapy had a better prognosis than those who underwent surgery alone. The consequences of ATSCI are disastrous and there is still a lack of consensus on treatment. We need to further improve the treatment strategy in order to obtain more reliable functional improvement.

***Research perspectives***

This research area requires randomized multicenter studies with a larger sample size to provide more clinical evidence for the optimal management strategy in ATSCI patients to achieve a better prognosis.

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**Footnotes**

**Institutional review board statement:** This study was approved by the Ethics Committee of Peking University Third Hospital, China (No. M2020232).

**Informed consent statement:** Patients were not required to give informed consent for this study because the analysis used anonymous clinical data.

**Conflict-of-interest statement:** The authors declare that there is no conflict of interest.

**Data sharing statement:** Technical appendix, statistical code, and dataset available from the corresponding author at maqingbain@bjmu.edu.cn.

**STROBE statement:** The authors have read the STROBE Statement—checklist of items, and the manuscript was prepared and revised according to the STROBE Statement—checklist of items.

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**Figure Legends**



**Figure 1 Enrollment and randomization of patients.** ASIA: American Spinal Injury Association.



**Figure 2 Complications in the study population.**



A

B



D

C

**Figure 3 Analysis of prognostic follow-up.** A: Prognosis between surgery within 72 h and surgery after 72 h (a*P* < 0.05); B: Prognosis between the Steroid group and the Non-steroid group in American Spinal Injury Association (ASIA) A and B patients (a*P* < 0.05); C: Prognosis between the non-pneumonia group and the pneumonia group (a*P* < 0.05); D: Prognosis of patients with different ASIA grades on admission (*P* < 0.05). ASIA: American Spinal Injury Association.

**Table 1 Demographics and clinical characteristics of patients**

|  |  |  |  |  |
| --- | --- | --- | --- | --- |
|  | **Total (*n* = 304)** | **Improved (*n* = 135)** | **Not-improved (*n* = 169)** | ***P* value** |
| Demographics |  |  |  |  |
| male, *n* (%) | 257 (84.5) | 111 (82.2) | 146 (86.4) | 0.318 |
| Age, yr (mean ± SD) | 49.81 ± 13.03 | 50.47 ± 12.78 | 49.28 ± 13.24 | 0.67 |
| Co-morbidity, *n* (%) | 92 (32.2) | 38 (28.8) | 53 (31.4) | 0.455 |
| ASCI characteristics |  |  |  |  |
| Etiology, *n* (%) |  |  |  | 0.456 |
| Motor vehicle crashes | 90 (29.6) | 41 (30.4) | 49 (29.0) |  |
| High-altitude fall injuries | 37 (12.2) | 13 (9.6) | 24 (14.2) |  |
| Unintentional falls | 136 (44.7) | 66 (48.9) | 70 (41.4) |  |
| Bruise by heavy object | 19 (6.3) | 6 (4.4) | 13 (7.7) |  |
| other | 22 (7.2) | 9 (6.7) | 13 (7.7) |  |
| Multiple trauma, *n* (%) | 41 (13.5) | 13 (9.6) | 28 (16.6) | 0.078 |
| Fracture, *n* (%) | 137 (45.1) | 52 (38.5) | 85 (50.3) | 0.040a |
| Region of fracture, *n* (%) |  |  |  | 0.042a |
| Cervical  | 269 (88.5) | 126 (93.3) | 143 (84.6) |  |
| Lumbar  | 24 (7.9) | 5 (84.6) | 19 (11.2) |  |
| Thoracic | 11 (3.6) | 4 (3.0) | 7 (4.1) |  |
| ASIA grade at admission |  |  |  | 0.0 |
| A | 78 (25.7) | 14 (10.4) | 64 (37.9) |  |
| B | 33 (10.9) | 16 (11.9) | 17 (10.1) |  |
| C | 66 (21.7) | 47 (34.8) | 19 (11.2) |  |
| D | 127 (41.8) | 58 (43.0) | 69 (40.8) |  |
| Time from event to visit | 3.06 ± 2.13 | 2.7 ± 1.96 |  32.9 ± 2.20 | 0.019a |
| Treatment (h) |  |  |  |  |
| Time from event to surgery |  |  |  | 0.005c |
| Within 72 h  | 133 (43.8) | 71 (52.6) | 62 (36.7) |  |
| After 72 h | 171 (56.3) | 64 (47.4) | 107 (63.3) |  |
| Steroid usage | 208 (68.4) | 87 (64.4) | 169 (71.6) | 0.182 |
| Complications | 100 (32.9) | 33 (24.4) | 67 (39.6) | 0.005c |
| Hospital stay (d) | 9.89 ± 10.11 | 7.79 ± 5.92 | 11.55 ± 12.25 | 0.002c |
| ICU admission, *n* (%) | 95 (31.3) | 34 (25.2) | 61 (36.1) | 0.041a |

a*P* < 0.05; b*P* < 0.01; c*P* < 0.005. ASCI: Acute spinal cord injury; ASIA: American Spinal Injury Association; ICU: Intensive care unit.

**Table 2 Steroid treatment**

|  |  |  |  |
| --- | --- | --- | --- |
|  | **Steroid** | **Non-steroid**  | ***P* value** |
| Average hospitalization days |  | 10.32 ± 11.4 | 8.94 ± 6.4  | 0.117 |
| Average hospitalization days in ICU |  | 10.75 ± 11.9 | 7.68 ± 6.3  | 0.095 |
| Mean dose of steroids (mg/d)DexamethasoneMethylprednisolone |  | 15.20 ± 5.0286.02 ± 213.4 |  |  |
| Total dose of steroids (mg) |  |  |  |  |
| Dexamethasone |  | 42.66 ± 23.5 |  |  |
| Methylprednisolone |  | 833.96 ± 717.8 |  |  |

ICU: Intensive care unit.

**Table 3 Complications in the Steroid group and Non-Steroid group.**

|  |  |  |  |  |
| --- | --- | --- | --- | --- |
|  |  | **Steroid** | **Non-steroid** | ***P* value** |
| Pneumonia (*n* = 53) |  | 46 | 7 | 0.002c |
| Electrolyte disturbances (*n* = 37)DVT (*n* = 17)  |  | 2911 | 86 | 0.1640.734 |

c*P* < 0.005; DVT: Deep venous thrombosis.