Predicting the Role of Preoperative Intramedullary Lesion Length and Early Decompressive Surgery in ASIA Impairment Scale Grade Improvement Following Subaxial Traumatic Cervical Spinal Cord Injury

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Abstract

Background Traumatic cervical spinal cord injury (TCSCI) is a disabling condition with uncertain neurologic recovery. Clinical and preclinical studies have suggested early surgical decompression and other measures of neuroprotection improve neurologic outcome. We investigated the role of intramedullary lesion length (IMLL) on preoperative magnetic resonance imaging (MRI) and the effect of early cervical decompressive surgery on ASIA impairment scale (AIS) grade improvement following TCSCI.

Methods In this retrospective study, we investigated 34 TCSCI patients who were admitted over a 12-year period, from January 1, 2008 to January 31, 2020. We studied the patient demographics, mode of injury, IMLL and timing of surgical decompression. The IMLL is defined as the total length of edema and contusion/hemorrhage within the cord. Short tau inversion recovery (STIR) sequences or T2-weighted MR imaging with fat saturation increases the clarity of edema and depicts abnormalities in the spinal cord. All patients included had confirmed adequate spinal cord decompression with cervical fixation and a follow-up of at least 6 months.

Results Of the 34 patients, 16 patients were operated on within 24 hours (early surgery group) and 18 patients were operated on more than 24 hours after trauma (delayed surgery group). In the early surgery group, 13 (81.3%) patients had improvement of at least one AIS grade, whereas in the delayed surgery group, AIS grade improvement was seen in only 8 (44.5%) patients (early vs. late surgery; odds ratio [OR] = 1.828; 95% confidence interval [CI]: 1.036–3.225). In multivariate regression analysis coefficients, the timing of surgery and intramedullary edema length on MRI were the most significant factors in improving the AIS grade following cervical SCI. Timing of surgery as a unique variance predicted AIS grade improvement significantly.
Role of Preoperative IMLL and Early Decompressive Surgery in AIS Grade Improvement

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Introduction

Spinal cord injury (SCI) is a disabling neurologic condition with a significant socioeconomic impact on injured patients and their families. Global prevalence rates of traumatic SCI varied from approximately 250 per million to 906 per million and data on incidence varied from approximately 8 to 49.1 per million.1,2 The major cause of injury in the developed world is motor vehicle accidents as compared with two-wheeler accidents in Southeast Asia.3

Traumatic cervical spinal cord injury (TCSCI) affects the subaxial cervical spine with or without fracture and dislocations. TCSCI clinically can present as an incomplete injury with tetraparesis and sensory loss,4 whereas more severe injury results in tetraplegia with complete sensory loss below the level of injury. The American Spinal Injury Association (ASIA) Impairment Scale (AIS) is used commonly in grading the degree of sensory and motor impairment.5

The initial injury to the spinal cord at the time of an accident is known as a primary injury. Spinal cord injury is caused by fractured vertebral, herniated ruptured disk, and ligaments.6–9 The force of impact directly damages both ascending and descending pathways in the spinal cord and disrupts blood vessels and cell membranes10–13 causing spinal shock, hypertension, cord ischemia, ionic imbalance, and accumulation of neurotransmitter at the injured site.14,15 The secondary injury to the cord follows primary damage, causing further loss of remaining viable neurons in the vicinity. Secondary injury can be divided into acute, subacute, intermediate, and chronic phases.14 Within a few minutes of primary injury, the secondary injury begins causing local ischemia, vasospasm, and hypoperfusion.16–21 Ischemia increases local vascular permeability, compromising the integrity of the blood–spinal cord barrier (BSCB).21 This disruption of BSCB initiates a rapid influx of inflammatory cells like neutrophils. macrophages that release proinflammatory cytokines. The most common proinflammatory cytokines are tumor necrosis factor-α (TNF-α), interleukin-6 (IL-6), interleukin-1β (IL-1β).23–25

The release of proinflammatory cytokines, chemokines, nitric oxide, ions, and proteases, results in cell death through apoptosis and necrosis.

The intramedullary lesion on MRI represents the injury resulting from the molecular cascade, as a result of the trigger from primary injury. The intramedullary lesion includes the spinal cord edema and contusion/hemorrhage.26–29 The spinal cord edema rapidly increases in the first 48 hours after SCI, followed by a gradual decrease in the following 3 weeks.26,27 It collaborates with acute inflammatory changes in the first 48 hours.27 MRI can reliably show the extent of cord compression and signs of cord injury.28 Hematoma in the spinal cord has been associated with a poor neurologic prognosis and is one of the important predictors for neurologic outcome. The intramedullary blood itself leads to necrosis, apoptosis, and impaired regeneration.29,30

The primary injury cannot be influenced through treatment, but timely optimum management can reduce or limit secondary damage. Early surgical decompression of the spinal cord is neuroprotective.12,21–25 The neuroprotective effect seems to vary inversely with the time elapsed from injury to decompression.37–39 In this study, we analyzed the impact of preoperative intramedullary lesion length (IMLL), early surgical decompression, and other factors on AIS grade improvement after 6 months of follow-up. The null hypothesis was that in subaxial TCSCI preoperative IMLL on MRI and early surgical decompression do not affect long-term improvement in AIS grade.

Methods

This study is a retrospective analysis of over 12 years, from January 1, 2008 to January 31, 2020. We selected 34 TCSCI patients who were admitted to a level I trauma center. The inclusion criteria were older than 16 years and younger than 80 years, Glasgow Coma Scale (GCS) score greater than 14, patients with subaxial cervical spine fracture-dislocations, and MRI studies indicating adequate spinal cord

(p < 0.001). The mean IMLL was 41.47 mm (standard deviation [SD]: 18.35; range: 20–87 mm). IMLL was a predictor of AIS grade improvement on long-term outcome in bivariate analysis (p < 0.001). This study suggests that patients who had IMLL of less than 30 mm had a better chance of grade conversion irrespective of the timing of surgery. Patients with an IMLL of 31 to 60 mm had chances of better grade conversion after early surgery. A longer IMLL predicts lack of improvement (p < 0.05). If the IMLL is greater than 61 mm, the probability of nonconversion of AIS grade is higher, even if the patient is operated on within 24 hours of trauma.

Conclusion

Surgical decompression within 24 hours of trauma and shorter preoperative IMLL are significantly associated with improved neurologic outcome, reflected by better AIS grade improvement at 6 months’ follow-up. The IMLL on preoperative MRI can reliably predict outcome after 6 months. The present study suggests that patients have lesser chances of AIS grade improvement when the IMLL is ≥61 mm.
Role of Preoperative IMLL and Early Decompressive Surgery in AIS Grade Improvement

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Table 1 Morphology of injury based on AO classification (for detailed classification, see Vaccaro et al.40)

<table>
<thead>
<tr>
<th>AO: insignificant Injury</th>
<th>B1: posterior tension band injury (Bony)</th>
<th>C: translational injury in any axis displacement</th>
</tr>
</thead>
<tbody>
<tr>
<td>A1: wedge compression fracture</td>
<td>B2: posterior tension band injury (Ligamentous)</td>
<td></td>
</tr>
<tr>
<td>A2: split fracture</td>
<td>B3: anterior tension band injury</td>
<td></td>
</tr>
<tr>
<td>A3: incomplete burst fracture</td>
<td></td>
<td></td>
</tr>
<tr>
<td>A4: complete burst fracture</td>
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<td></td>
</tr>
</tbody>
</table>

Table 2 American Spinal Injury Association scale (ASIA) Impairment Scale (AIS) grade according to International Standards for Neurological Classification-1 (ISNC-1)

<table>
<thead>
<tr>
<th>AIS Grade</th>
<th>Description</th>
</tr>
</thead>
<tbody>
<tr>
<td>A0</td>
<td>Complete. No sensory or motor function is preserved below the level of injury</td>
</tr>
<tr>
<td>A1</td>
<td>Sensory incomplete. Sensory but no motor function is preserved below the motor level on either side of the body</td>
</tr>
<tr>
<td>A2</td>
<td>Motor incomplete. Motor function is preserved, but less than half of key muscle functions below the single neurologic level of injury (NLI) have a muscle grade &lt;3</td>
</tr>
<tr>
<td>A3</td>
<td>Motor incomplete status as defined above, with at least half (or more) of key muscle functions below the single NLI having a muscle grade ≥3</td>
</tr>
<tr>
<td>E</td>
<td>Normal</td>
</tr>
</tbody>
</table>

decompression following surgery and follow-up of at least 6 months after the trauma. Patients who had upper cervical SCI, inadequate spinal cord decompression on MRI, and patients on conservative treatment were excluded. We studied patient demographics, mode of injury, MRI findings, and time between trauma and surgical decompression. The IMLL is defined as the total length of edema and contusion/hemorrhage within the cervical cord. The morphology of injury was based on AO Spine classification. The injuries were classified into A0, A1–A4, B1–B3, and C (Table 1).40 After admission, the clinical parameters studied were ASIA motor score (AMS), ASIA sensory score (ASS), and the overall AIS grade.5 The improvement of AIS grade was defined as an improvement of at least one grade (Table 2).

MRI was performed using a 1.5-T MRI. The study included T1-weighted, T2-weighted, and short tau inversion recovery (STIR) sequences in sagittal and axial planes. Using STIR or T2-weighted imaging with fat saturation increases the clarity of edema in the bones and ligaments and depicts abnormalities in the spinal cord, disks, and epidural space (Figs. 1 and 2). Preoperative MRI was made in all the cases to delineate the degree of spinal cord compression, the extent of cord edema, presence of hematoma, and spinal stenosis. IMLL was measured in preoperative MRI for reference. Postoperative and follow-up MRI scans were obtained to evaluate the extent of spinal cord decompression and the course of the spinal cord lesion over time (Figs. 1 and 2). We divided IMLL into three groups41–43: (1) less than 30 mm, (2) 31 to 60 mm, and (3) greater than 61 mm. The rationale behind this grouping is that by dividing continuous data into three categories, it is more effective to correlate data without sacrificing the interpretability.43

All patients were medically stabilized before surgery37,44 intubated, and put on a ventilator if necessary. Preoperative cervical traction was used in 18 patients using Gardner-Wells tongs under fluoroscopy. We used 2.25 kg of weight per segment, not exceeding 20 kg. In most cases, a closed reduction of facet dislocation was achieved. Methylprednisolone was given if trauma was within 8 hours as per recommendations of the second National Acute Spinal Cord Injury Study (NASCIS-2) study.44 Patients who underwent decompressive surgery with fixation had either early surgery (<24 hours after trauma) or late surgery (≥24 hours after trauma). We operated as soon as possible after medical stabilization, but we classified our patients as per the Surgical Timing in Acute Spinal Cord Injury (STASCIS) study into early surgery and late surgery.37 The timing of surgery depended on patient arrival at a hospital emergency ward, the time required for medical stabilization, and the time required for necessary diagnostic investigation and documentation.37 Cervical cord decompression was done from either the anterior or the posterior approach. In a few cases, both approaches were used. Spinal fixation was done in all cases whether it was an anterior or posterior approach.45,46 The neurologic status was recorded at the 6-month follow-up. The 6 months of follow-up was based on recommendations used in the STASCIS, NASCIS, and Sygen trials.37–52 The AIS improvement of one grade represented grade conversion.13,27,45

Statistical analysis was performed using the IBM SPSS classic version. The descriptive analysis was done as a mean and standard deviation (SD) for continuous variables and frequencies for categorical variables. The correlation was analyzed using bivariate analysis calculating the Pearson coefficient and its significance (Figs. S1, S2, available online only). Multiregression analysis was done using grade improvement as a dependent variable, and grades on admission, mechanism of injury, IMLL, presence of hemorrhage/contusion, timing of surgery were independent variables.
Results

A total of 34 patients were included in this study. The mean age was 54.06 ± 15.44 years (range: 18–80 years). Twenty-nine (85.3%) patients were males and 5 (14.7%) were females (►Table 3). Of the 34 patients, 16 patients were operated on within 24 hours after SCI and were considered the early surgery group (►Table 3). Eighteen patients were operated on more than 24 hours after SCI and they were designated as the delayed surgery group. In the early surgery cohort, the mean age was 50.6 ± 19.6 with 13 males (81.3%) and 3 females (18.7%). In the late surgery cohort, the mean age was 57 ± 11.0 years with 16 males (88.9%) and 2 females (11.1%). Overall TCSCI was caused by a fall in 19 (55.9%) patients, motor vehicular accidents in 13 (38.2%) patients, and sports injury in 2 patients (►Table 3).

Among the patients, 30.8% of AIS grade A, 80.0% of AIS grade B, and 81.8% of AIS grade C at admission improved to a better AIS grade during 6 months of follow-up (p < 0.05; ►Table 4). The mean age of patients who improved to a better AIS grade was 52.48 ± 17.26 years, whereas the mean age of patients who did not show improvement was 56.62 ± 12.16 years. In the early surgery group, 13 (81.3%) patients had an AIS grade improvement of at least one grade, whereas 3 (18.7%) patients did not improve (►Table 3). In the delayed group (≥24 hours after trauma), AIS grade improvement was seen in 8 (44.5%) patients (early vs. late surgery: odds ratio [OR] = 1.828; 95% confidence interval [CI]: 1.036–3.225).

We performed multiregression analysis using grade improvement as a dependent variable, and timing of surgery, IMLL, presence of hemorrhage/contusion, mechanism of injury, grade on admission as independent variables. In the model summary (►Fig. S1, available online only) of multiregression analysis, R² is 0.55. This means that the contribution of these factors was 55% in grade improvement. Analysis of variance (ANOVA) of the regression model suggests the overall contribution being significant (p < 0.001; R² = 0.55). In the final step of regression analysis coefficients, IMLL and timing of surgery were the most significant factors related to AIS grade improvement (►Fig. S1, available online only). The timing of surgery as a unique variance predicted AIS grade improvement better (p < 0.001).

In our study, the mean IMLL was 41.47 mm (SD: 18.35; range: 20–87 mm). IMLL on preoperative MRI was a significant predictor of grade conversion in bivariate analysis (p < 0.001) (►Fig. S2). Longer IMLL on preoperative MRI

Fig. 1 A 55-year-old woman sustained a fall and was brought to the emergency room 6 hours after trauma. At the time of admission, she was hypotensive and on a ventilator. Her American Spinal Injury Association scale (ASIA) Impairment Scale (AIS) grade was B. (A) Preoperative magnetic resonance imaging (MRI) of the cervical spine showed C6 vertebral body fracture with retropulsed fragment causing significant cervical cord compressive injury with bleeding at the epicenter and spinal cord edema extending from C3 to C8. It also shows a bright T2-weighted signal in the surrounding ligaments that is suggestive of disruption. The intramedullary lesion length (IMLL) was 78 mm. She underwent anterior cervical fusion within 24 hours resulting in proper realignment of the spinal column. (B) Postoperative MRI taken after 1 week of surgery indicated good decompression with significant improvement of intramedullary edema (IMLL = 60 mm). (C) MRI after 4 weeks showed a further decrease in IMLL to 8 mm. Six months later, her AIS grade was D.
predicted poor AIS grade conversion after 6 months as depicted in Figs. 3 and 4. AIS grade A patients had a mean IMLL of 57.54 ± 19.4 mm, AIS grade B of 35.2 ± 8.30 mm, and AIS grade C of 28.2 ± 5.25 mm.

There were 12 patients in whom IMLL was ≤30 mm, 15 patients had IMLL of 31 to 60 mm, and 7 patients had IMLL ≥61 mm. In the first group (IMLL ≤30 mm), AIS grade improvement of at least one grade was seen in 10 patients (83.3%). Four patients had early surgery and all four patients improved (→ Table 5). Eight patients underwent late surgery, out of which six patients (75%) improved (p = 0.317; 95% CI: −0.778 to 0.278). In the second group (IMLL: 31–60 mm), 10 patients (66.6%) showed grade improvement (p < 0.05). Out of nine patients who had early surgery (≤24 hours of admission), eight patients improved, whereas two patients who had late surgery improved. In the third group with IMLL ≥61 mm, three patients had early surgery and four patients underwent late surgery. Only one patient who was operated on within 24 hours had grade conversion (p = 0.236).

The probability of grade improvement is higher with shorter IMLL (→ Fig. S1, available online only). A longer IMLL predicts lack of improvement (p < 0.05). If the IMLL is more than 61 mm, the probability of nonconversion of AIS grade is higher, irrespective of the timing of surgery. These findings rejected our null hypothesis.

The AMS on admission was 28.32 ± 15 for the whole cohort. The AMS of patients who converted to a better AIS grade was 33.67 ± 12.48 (p < 0.05). Patients who did not show any grade improvement had a mean AMS score of 19.69 ± 15.14. The ASS of the patients on admission was 56.7 ± 27.27. The ASS of patients who had grade conversion was 61.43 ± 23.6 and the ASS of those who did not convert to a better grade was 49 ± 31.9 (p > 0.05).

An intramedullary hemorrhage was present on preoperative MRI at the epicenter of injury in 13 (38.2%) patients. Nine (69.2%) patients with intramedullary hemorrhage did not improve after 6 months of follow-up. This was significant on univariate analysis (p = 0.005), but not on multiregression analysis (→ Table 3).

Discussion

This study analyzed factors such as admission AMS, AIS grade, injury morphology, the timing of surgery, surgical approach, presence of intramedullary hemorrhage, and IMLL on AIS grade conversion at 6 months after TCSCI. On multiregression analysis, using grade improvement as a dependent variable, preoperative IMLL and early surgery were the two most significant factors that predicted AIS grade improvement.

In this series, there was a significant association between preoperative IMLL and AIS grade conversion (p = 0.03). Our AIS grade conversion rates were similar to results reported by Marino et al46 and Aarabi et al42 (→ Table 4).

Patients who had a mean preoperative IMLL of 33.71 ± 9.96 mm had grade conversion, whereas patients who had no grade conversion had a mean IMLL of 54 ± 22.05 mm (p < 0.001). In the present study, patients with IMLL less than 30 mm showed better grade improvement irrespective of the timing of surgery. This is because shorter IMLL indicated less severe SCI (→ Figs. 3 and 4). Patients with IMLL between 31 and 60 mm had more often grade conversion with early surgery. Probably, the observed effect was related to a prevention of secondary cord damage by early adequate decompressive surgery. Patients with IMLL greater than 61 mm had a lesser probability of AIS grade improvement irrespective of early or late surgery (→ Fig. 2). In these patients, delayed surgical decompression is justified.

The IMLL on MRI is proportional to the severity of the primary and secondary injuries.13,15,27,36,53–59

Parashari et al41 studied the prognostic role of MRI and its association with the clinical outcome and concluded that chances of improvement were more in patients with cord edema involving less than 3 cm of the spinal cord than patients with cord edema involving greater than 3 cm of the spinal cord.

According to Aarabi et al,42 patients who had IMLL close to 62.4 mm and adequate decompression had a better chance of conversion (p = 0.001) than patients with inadequate decompression and an IMLL of 100.3 mm.

Is there an association between preoperative IMLL on MRI and the severity of SCI? In our study, longer IMLL was...
associated with more severe AIS grade on admission. Edema and necrosis on MRI represent the injury resulting from molecular cascade as suggested by a preclinical high-field (9.4 T) MRI study on rats.53,54 This hyperintense signal and lesion length on high-field MRI were the most valuable parameters and were highly correlated to histopathologic changes.53,54 Several studies have indicated a significant relationship between the severity of SCI, neurologic outcome, and the finding of contusion hemorrhage on MRI.55–63

Table 3  Patient characteristics and relationship with AIS grade conversion

<table>
<thead>
<tr>
<th>Characteristics</th>
<th>Frequency</th>
<th>AIS grade conversion: yes</th>
<th>AIS grade conversion: no</th>
<th>Comments</th>
</tr>
</thead>
<tbody>
<tr>
<td>Age (y)</td>
<td>54.06 ± 15.44</td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Gender (%)</td>
<td></td>
<td></td>
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<td></td>
</tr>
<tr>
<td>Male</td>
<td>29 (85.3)</td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Female</td>
<td>5 (14.7)</td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Mode of accident (%)</td>
<td></td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>MVC</td>
<td>13 (38.2)</td>
<td>8 (38.1)</td>
<td>5 (38.5)</td>
<td>p = 0.904</td>
</tr>
<tr>
<td>Fall</td>
<td>19 (55.9)</td>
<td>12 (57.1)</td>
<td>7 (53.8)</td>
<td></td>
</tr>
<tr>
<td>Others</td>
<td>2 (5.9)</td>
<td>1 (4.8)</td>
<td>1 (7.7)</td>
<td></td>
</tr>
<tr>
<td>AIS at admission (%)</td>
<td></td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>AIS grade A</td>
<td>13 (38.2)</td>
<td>4 (30.8)</td>
<td>9 (69.2)</td>
<td>p = 0.08</td>
</tr>
<tr>
<td>AIS grade B</td>
<td>10 (29.4)</td>
<td>8 (80)</td>
<td>2 (20)</td>
<td></td>
</tr>
<tr>
<td>AIS grade C</td>
<td>11 (32.4)</td>
<td>9 (81.8)</td>
<td>2 (18.2)</td>
<td></td>
</tr>
<tr>
<td>ASIA Motor Score (AMS)</td>
<td></td>
<td>28.32 ± 15.0</td>
<td>33.67 ± 12.48</td>
<td>p = 0.011</td>
</tr>
<tr>
<td>ASIA Sensory Score (ASS)</td>
<td></td>
<td>56.68 ± 27.2</td>
<td>61.43 ± 23.6</td>
<td>p &gt; 0.05</td>
</tr>
<tr>
<td>Injury morphology (%)</td>
<td></td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>A0</td>
<td>10 (23.5)</td>
<td>8 (80)</td>
<td>2 (20)</td>
<td>p = 0.14</td>
</tr>
<tr>
<td>A1–A4</td>
<td>19 (55.9)</td>
<td>12 (63.2)</td>
<td>7 (36.8)</td>
<td></td>
</tr>
<tr>
<td>B1–B3</td>
<td>2 (8.8)</td>
<td>1 (50)</td>
<td>1 (50)</td>
<td></td>
</tr>
<tr>
<td>C</td>
<td>3 (11.8)</td>
<td>0</td>
<td>3 (100)</td>
<td></td>
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<tr>
<td>Timing of surgery (%)</td>
<td></td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Early surgery (&lt;24 h)</td>
<td></td>
<td>16 (47.1)</td>
<td>13 (81.2)</td>
<td>3 (18.8)</td>
</tr>
<tr>
<td>Delayed surgery (&gt;24 h)</td>
<td></td>
<td>18 (52.9)</td>
<td>8 (44.4)</td>
<td>10 (55.6)</td>
</tr>
<tr>
<td>Surgical approach (%)</td>
<td></td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Anterior</td>
<td>15 (44.1)</td>
<td>11 (73.3)</td>
<td>4 (26.7)</td>
<td>p = 0.578</td>
</tr>
<tr>
<td>Posterior</td>
<td>17 (50)</td>
<td>8 (47)</td>
<td>9 (53)</td>
<td></td>
</tr>
<tr>
<td>Combined</td>
<td>2 (5.9)</td>
<td>2 (100)</td>
<td>[ndash]</td>
<td></td>
</tr>
<tr>
<td>Preoperative IMLL (%)</td>
<td></td>
<td></td>
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<td></td>
</tr>
<tr>
<td>group 1; &lt;30 mm</td>
<td>12 (35.3)</td>
<td>10 (83.3)</td>
<td>2 (16.7)</td>
<td>p = 0.317; OR: 0.6; 95% CI: 0.36–0.99</td>
</tr>
<tr>
<td>Group 2; 31–60 mm</td>
<td>15 (44.1)</td>
<td>10 (66.7)</td>
<td>5 (33.3)</td>
<td>p &lt; 0.05; OR: 16; 95% CI: 1.093–234.2</td>
</tr>
<tr>
<td>Group 3; &gt;61 mm</td>
<td>7 (20.6)</td>
<td>1 (14.2)</td>
<td>6 (85.8)</td>
<td>p = 0.236; OR: 3; 95% CI: 0.968–9.302</td>
</tr>
<tr>
<td>Presence of hemorrhage (%)</td>
<td></td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Yes</td>
<td>13 (38.2)</td>
<td>4 (30.8)</td>
<td>9 (69.2)</td>
<td>p = 0.005; 95% CI: 0.190–0.813</td>
</tr>
<tr>
<td>No</td>
<td>21 (61.8)</td>
<td>17 (81)</td>
<td>4 (19)</td>
<td></td>
</tr>
</tbody>
</table>

Abbreviations: AIS: American Spinal Injury Association impairment scale; ASIA, American Spinal Injury Association scale; ACI, confidence interval; IMLL, intramedullary lesion length; MVC, motor vehicle collision; OR, odds ratio.

In this study, IMLL was delineated in both STIR and T2 images pre- and postoperatively (Fig. 1 and 2).64,65 Our study infers that preoperative IMLL was a significant predictor of long-term outcome after TTCSCI.

The other aspect of IMLL is its expansion after trauma. According to Aarabi et al27 and Le et al,66 the rate of expansion of the IMLL differs with the severity of subaxial cervical injury. The speed of expansion measures from 200 µm/h in AIS grade C patients to 900 µm/h in AIS grade C.
A and B patients. After a cervical injury, progressive edema and hemorrhage cause severe progressive compression of the cervical cord circumferentially against the dura and the bony structures at multiple levels.\textsuperscript{67–69} The spinal cord edema can be reduced through timely and adequate surgical decompression and restoration of alignment\textsuperscript{69,70} as shown in $\textbullet$ Figs. 1, 2, 5, and 6. Interestingly, it was found that adequate decompression was done in only 66% of AIS grade A and B patients,\textsuperscript{71} which resulted in a poor outcome.

In our series, the presence of intramedullary hemorrhage on preoperative MRI was a bad sign for AIS grade conversion. AIS grade conversion occurred only in 30.8% of patients with intramedullary blood ($\textbullet$ Fig. 7). On multivariate analysis, the IMLL was a stronger predictor of AIS grade conversion than the presence of intramedullary hemorrhage. The possible explanation could be that IMLL includes intramedullary hematoma.\textsuperscript{72} Flanders et al\textsuperscript{62} reported that hemorrhage is almost always present concurrently with edema, and on MRI it is surrounded by a hyperintensity normally associated with edema.

The greatest lengths of spinal cord edema and hematoma were found in those with the most severe (AIS grades A and B) injuries.\textsuperscript{26} Hematoma in the spinal cord was seen in all AIS A and B patients and in 50% of the AIS C patients.\textsuperscript{26} The presence of intramedullary hemorrhage was predictive of worse neurologic recovery. The extent of hemorrhage

\begin{table}
\centering
\caption{Correlation between IMLL and AIS grade conversion}
\begin{tabular}{|c|c|c|c|c|c|}
\hline
IMLL & No. of patients & Early surgery & AIS grade conversion (%) & Delayed surgery & AIS grade conversion (%) & Significance of early surgery vs. delayed surgery and 1 grade improvement \\
& & & after early surgery & & after delayed surgery & \\
\hline
$\leq$30 mm & 12 & 4 & 4 (100) & 8 & 6 (75) & $p = 0.317$ OR: 0.6; 95% CI: 0.362–0.995 Not significant \\
31–60 mm & 15 & 9 & 8 (88.9) & 6 & 2 (33.3) & $p < 0.05$ OR: 16; 95% CI: 1.093–234.2 Significant \\
$\geq$61 mm & 7 & 3 & None & 4 & 1 (25) & $p = 0.286$ OR: 3; 95% CI: 0.968–9.302 Not significant \\
\hline
\end{tabular}
\end{table}

Abbreviations: AIS: American Spinal Injury Association impairment scale; CI, confidence interval; IMLL, intramedullary lesion length; OR, odds ratio.

Fig. 3 Bar graph showing the relation between the three groups of intramedullary lesion length (IMLL), grade improvement and timing of surgery.
within the cord has been correlated with outcome, with a small focus on hemorrhage of less than 4 mm more likely to be associated with radiologic and clinical improvement on follow-up, compared with a larger spinal cord hematoma. 

The cervical compression should be relieved early, but the question is how early? In our study, early decompression (<24 hours) has shown a significant impact on improvement in the AIS scale of at least one grade (Fig. 3). In the early surgery group, 81.3% of patients had an AIS grade improvement, whereas in the delayed group, AIS grade improvement was seen in 44.5% patients only (p = 0.027). Early surgical decompression relieves cord compression, preventing further secondary injury. The STASCIS trial, published in 2012, was a prospective study of 222 patients undergoing early (<24 hours) versus late (>24 hours) decompression. Patients receiving early surgical intervention were twice as likely to improve by 2 or more AIS grades at 6 months. A prospective Canadian cohort study by Wilson et al suggested that a significant number of patients who underwent early decompression improved by two or more AIS grades. Dvorak et al reported shorter hospital stay after early decompression for patients with ASIA A or B injuries. Studies that performed postoperative MRI indicate that up to 25% of patients may need expansive dura plasty for adequate spinal cord decompression to improve functional outcome. Early decompression is recommended in clinical management guidelines by the American Association of Neurological Surgeons (AANS) and the Congress of Neurological Surgeons. The recent AO Spine guideline also recommends decompression within 24 hours of SCI. There are advocates of surgery within 12 hours (ultra-early) with better outcome after cervical SCI. There are studies that suggest there is no benefit of early decompression. Aarabi et al concluded that complete decompression of cervical cord, confirmed by postoperative MRI, determines the long-term neurologic outcome and not the timing of surgery. In their study, decompression had been performed either ultra-early (<12 hours) or early (12-24 hours). A study by Vaccaro et al revealed no significant neurologic benefit when cervical spinal cord decompression was performed less than 72 hours after injury as compared with waiting longer than 5 days, whereas a systematic review suggested that decompression within 24 hours resulted in improved outcomes compared with both delayed decompression and conservative treatment.

Neuroprotective measures within the acute phase (48 hours) could also be beneficial. The efficacy of many therapies during the acute phase is still under evaluation like riluzole and minocycline. Both are presently being tested for efficacy and safety in phase III randomized controlled trials. Riluzole is a sodium channel blocking drug that

Table 5  AIS grade conversion (%) in the present series and other series

<table>
<thead>
<tr>
<th>AIS grade conversion</th>
<th>Spiess et al 2009 (%)</th>
<th>Marino et al (%)</th>
<th>Aarabi et al (%)</th>
<th>Current study (%)</th>
</tr>
</thead>
<tbody>
<tr>
<td>A</td>
<td>28.3</td>
<td>29.8</td>
<td>26.9</td>
<td>30.8</td>
</tr>
<tr>
<td>B</td>
<td>73.3</td>
<td>75.2</td>
<td>65.5</td>
<td>80</td>
</tr>
<tr>
<td>C</td>
<td>79.6</td>
<td>86.2</td>
<td>78.9</td>
<td>81.8</td>
</tr>
</tbody>
</table>

Abbreviations: AIS: American Spinal Injury Association impairment scale; IMLL, intramedullary lesion length.
decreases the influx of calcium and prevents the trigger of increased $\text{Na}^+$ influx from continuous activation of voltage-gated $\text{Na}^+$ channels,\textsuperscript{96} thus preventing further neuronal damage. Minocycline is an antibiotic that has neuroprotective properties in preclinical studies.\textsuperscript{98} It has been shown to reduce apoptosis of oligodendrocytes and microglia and improve neurologic recovery in rodent models of SCI.\textsuperscript{99} To date, the most effective clinical treatment to limit tissue damage following primary injury is early surgical decompression ($<24$ hours postinjury) of the injured spinal cord.\textsuperscript{70–72} Overall, the extent of the primary injury determines the severity and outcome of SCI.\textsuperscript{53,84}

### Limitations

This study has a limitation of being a retrospective, single-center study. Also, the number of patients were low due strict inclusion criteria. Another limitation was not having MRI
prior to trauma to document already preexisting changes in the cervical spine.

**Conclusion**

Primary injury and early stages of secondary injury of the cervical spinal cord are very well visualized on MRI as an intramedullary lesion. The IMLL on preoperative MRI can reliably predict outcomes after 6 months. Patients who had IMLL of less than 30 mm had a better chance of grade conversion irrespective of the timing of surgery. Patients with an IMLL length of 31 to 60 mm had a higher chance of grade conversion if early surgery is performed. Longer IMLLs (> 61 mm) are associated with a higher rate of lacking improvement. Surgical decompression within 24 hours of trauma significantly improves neurologic outcome, reflected by better AIS grade improvement at 6 months of follow-up. Other factors like patient demographics, mechanism of injury, grade on admission, and fracture morphology had no significant effect on long-term prognosis.
Conflict of Interest
None declared.

References
20 Furlan JC, Fehlings MG. Cardiovascular complications after acute spinal cord injury; pathophysiology, diagnosis, and management. Neurosurg Focus 2008;25(05):E13
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Swank ML, Sutterlin CE III, Bossons CR, Dias BE. Rigid internal fixation with lateral mass plates in multilevel anterior and posterior reconstruction of the cervical spine. Spine 1997;22(03):274–282


Resnick DK. Updated guidelines for the management of acute cervical spine and spinal cord injury. Neurosurgery 2013;72(Suppl 2):1


Kim M, Hong SK, Jeon SR, Roh SW, Lee S. Early (<48 hours) versus late (>48 hours) surgery in spinal cord injury: treatment outcomes and risk factors for spinal cord injury. World Neurosurg 2018;118:e513–e525


Wells JE, Hurlbert RJ, Fehlings MG, Yong VW. Neuroprotection by minocycline facilitates significant recovery from spinal cord injury in mice. Brain 2003;126(Pt 7):1628–1637

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