Introduction

Endotracheal intubation in patients with traumatic brain injury (TBI) and damage to the cervical spine is an important factor in determining life and death. Endotracheal intubation failures create the possibility of respiratory arrest, secondary brain damage caused by hypoxia or hypercapnia, and aspiration pneumonia [1]. In addition, intubation itself may also cause brain herniation in patients with elevated intracranial pressure (ICP), such as brain hematoma. Therefore, the purpose of airway management in patients with neurological damage is the maintenance of adequate oxygenation and respiration, as well as the prevention of aspiration. However, the procedure should be performed while considering the possibility of increasing intracranial pressure and cervical spine injury.

Endotracheal Intubation in Patients with TBI

Intubation is common in the emergency room when traumatic brain surgery occurs due to an emergency in the operating room. This review discusses the airway management of patients with head trauma upon arrival to the emergency room.
Intubation should be performed without delay when respiratory arrest or cardiac arrest is imminent. Intubation must be performed even when the patient is unconscious, and aspiration due to vomiting is expected. The indications for intubation are inadequate oxygenation, improper respiration, loss of laryngeal reflex, and when neurological dysfunction and cardiopulmonary dysfunction are expected [2]. However, intubation itself can cause severe hemodynamic changes, and this risk should be considered in risk-benefit analyses. Rapid sequence intubation should be performed while considering increased ICP to prevent secondary brain injury [3-5]. Increased ICP can be expected particularly if the patient is unconscious, has one or both pupils dilated, or assumes a decerebrate posture [6]. Additionally, ICP can arise as a reflex sympathetic response due to intubation and direct laryngeal reflex. Comatose patients appear to have no reaction, but the stimulation of intubation may increase ICP [7].

Therefore, intubation following only muscle relaxant administration should be discouraged in patients with increased ICP. Intubation should be performed gently by experienced technicians to minimize irritation. The use of lidocaine (1.5 mg/kg, intravenous [IV]), esmolol (1–2 mg/kg, IV), fentanyl (2–3 μg/kg, IV), propofol (2 mg/kg, IV), ketamine (2 mg/kg, IV), and muscle relaxants is also required [8-14].

Can Ketamine be used in TBI Patients?

Recent studies have reported the features of ketamine [15, 16]. Studies of ketamine since the 1970s have shown that its administration alone increases ICP in brain injury patients [17-19]. Therefore, ketamine should not be used in these patients. However, sedative and analgesic medications, such as barbiturates, opioids, and propofol, can decrease blood pressure in TBI patients, consequently decreasing cerebral perfusion pressure (CPP). Contrary to previous reports, ketamine was shown to decrease ICP during interventions that can increase ICP. Bar-Joseph et al. [12] monitored blood pressure, ICP, and CPP for 10 min after ketamine (1–1.5 mg/kg) administration to lower markedly elevated ICP and reported that ketamine was a very effective and safe medication for TBI patients because it decreased ICP during interventions that can increase ICP, without lowering CPP or mean arterial pressure (MAP). Jabre et al. [20] compared etomidate and ketamine use during rapid sequence intubation in acutely ill patients and demonstrated that the ventilator weaning time, vasopressor weaning time, and 28-day survival rates were not different between these medications; however, adrenal insufficiency was higher in the etomidate group. Comparison of the sequential organ failure assessment score revealed that ketamine was superior to etomidate in septic patients and showed better 28-day survival rates.

Prevention of ICP Elevation after Intubation

Two techniques to prevent an additional increase in ICP after intubation are an immediate head elevation after intubation, or tracheal intubation in the head-elevated position, if possible. MAP should be always maintained over 80–100 mmHg, and CPP should be maintained over 60 mmHg, if ICP monitoring is possible, with adequate analgesic and sedative administration for pain control. Notably, hypoxemia must be prevented. It is important to use 100% oxygen to maintain adequate pre-oxygenation before induction and maintain peripheral oxygen saturation over 94%. However, the oxygen concentration should be lowered to maintain FIO2, at 0.5 or PaO2, within 110–300 mmHg when the patient is hemodynamically stabilized and oxygenated adequately after intubation (Table 1) [21].

Risks and Benefits of Pre-hospital Endotracheal Intubation

Whether paramedics should perform endotracheal intubation in TBI patients or it should be performed in the emergency room is controversial. The use of sedatives and muscle relaxants is also controversial [22-29]. Wang et al. [30] reported a retrospective study that compared two groups of 4,098 TBI patients with similar disease and similar age who received pre-hospital intubation or hospital intubation. Patients who received pre-hospital intubation had a 3.99-fold higher mortality rate than hospital intubation patients. Stiell et al. [31] compared major trauma patients who received pre-hospital intubation and patients who did not undergo this procedure. There was no difference in the overall survival rate, but major trauma patients with a GCS below 9 who received out-of-hospital intubation had a lower survival rate. A study of 2,135 severe TBI patients with a GCS of 8 or below demonstrated that patients with out-of-hospital advanced airway management had a higher mortality rate on day 28 and a worse 6-month neurological outcome than patients who received advanced airway management at...
the emergency department [32]. The following reasons for the poor results for out-of-hospital advanced airway management are suspected: 1) suboptimal paramedic training; 2) lack of intubation skills (such as holding a laryngoscope for a long time); 3) iatrogenic hypotension and bradycardia; and 4) intubation failure due to insufficient use of muscle relaxants [23,33-36]. Unintended hyperventilation after intubation can decrease brain oxygen delivery and perfusion, which are the most important causes of poor outcomes [37]. By contrast, Bernard et al. [27] reported that brain trauma patients older than 15 years with a GCS of 9 or below with rapid sequence intubation (3 min of pre-oxygenation and intubation under monitoring by pulse oximetry, end-tidal CO2 measurement, and electrocardiography, and medications, such as fentanyl, midazolam, and succinylcholine) had a better neurological outcome 6 months after the injury. The paramedic rapid sequence intubation group exhibited better results because this group received 100% oxygen via an endotracheal tube during transfer to the hospital, which increased the oxygen partial pressure in the blood and lessened neurological injury. Both groups showed similar oxygen saturation. Therefore, hyperoxia may reduce neurological injury. Davis et al. [38] reported that TBI patients who had PaO2 values between 110 and 487 mmHg with mild to moderate hyperoxemia on hospital arrival exhibited decreased mortality rates. However, extreme hyperoxemia increases mortality, likely due to hyperoxic cerebral vasoconstriction and oxygen free-radical formation [39-42]. The optimal oxygen tension is not known, but normoventilation and mild hyperoxemia using mechanical ventilation are ideal because hypocapnia (end-tidal CO2 < 35 mmHg) can cause cerebral vasoconstriction that leads to secondary brain injury, and hypercapnia can increase ICP via cerebral vasodilation.

Assessment of Cervical Spine Injury

The possibility of a cervical spine injury in a blunt trauma patient is approximately 1.8% [43]. The most common site of injury is the cervical spine 2 (C-spine 2, C2), and C6 and C7 are vulnerable [44]. The probability of a C-spine injury increases in the elderly, because of degenerative changes in the spine [45]. However, head injury patients are approximately fourfold more likely to have additional C-spine injuries than those without head injury. Patients with a GCS below 8 also have a higher likelihood of additional C-spine injury [46]. Demetriades et al. [47] examined 14,755 patients who visited a major trauma center over a 5-year period and reported that the incidence of C-spine injury was 2%. However, the incidence of C-spine injury rose to 10.2% in patients with a GCS of 8 or below. The possibility of damage to the cervical spine should always be considered in patients with head trauma until it can be excluded. However, the following circumstances must be considered to exclude the possibility of C-spine injury: 1) the absence of tenderness in the central portions of the C-spine, 2) no focal neurological deficit, 3) clear consciousness, 4) no evidence of intoxication, and 5) no distracting damage [48].

C-spine Anatomy and Injury Mechanisms

To understand the damage caused by flexion or extension of the C-spine during intubation, we will review the anatomy of the C-spine and its damage mechanism. The atlanto-occipital joint is responsible for 50% of the neck flexion and extension, and the atlanto-axial joint is responsible for 50% of the neck rotation. Because there is no intervertebral disc at these two joints, there is a high likelihood of inflammatory arthritis [49]. Anterior longitudinal ligament (ALL) prevents hyperextension, while posterior longitudinal ligament (PLL) prevents hyperflexion. The widest portion of the spinal canal is C1-C3, with a mid-sagittal diameter of approximately 16–30 mm. The narrowest part is C4-C7, with a mid-sagittal diameter of 14–23 mm. In this region, hyperextension can reduce the canal diameter by approximately 2–3 mm; thus, caution is needed in patients with hyperextension injury [50]. Axial compression injury is often caused by activities such as diving, and bursting fracture of the atlas is also called a “Jefferson fracture”. This site has a wide canal diameter, and neurological injury is rare. However, when the transverse ligament is damaged as well, it can become very unstable and requires caution [51]. An odontoid fracture occurs when flexion, extension, and rotation mechanisms are mixed. There are three types. In a type 1 odontoid fracture, only the tip of the dens is detached with no significant influence on instability. In a type 2 odontoid fracture, the base of the dens is broken. When fractures are associated with transverse ligament injury, it makes very unstable with a high mortality rate. In a type 3 odontoid fracture, the C2 body breaks off with the dens, with the atlas and occiput moving as a single mass, making it highly unstable. A C5 flexion teardrop fracture commonly occurs by a flexion mechanism [51]. It is usually caused by car accidents or a diving injury, with the body of the vertebra displacing posteriorly, often causing neurologic impairment. Bilateral facet dislocation is the most serious damage, with the complete destruction of both the ALL and PLL commonly accompanying neurologic injury. An example of an extension mechanism fracture would be a hangman’s fracture—i.e., fracture of the pedicle of an axis. It can be caused by severe hyperextension caused by diving or a car accident, and spinal cord injury is not severe because the spinal canal is wide at the C2 level. Of the fractures caused by extension injury, the teardrop fracture of the C2 is termed an ‘unstable injury’. It is caused by the tension of the ALL, which leads to the breakage of the anterior body of the C2. Although the neurological deficit is not severe, extension should be performed carefully [51]. Common
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Clinical Notes of C-spine Injury and Manual In-line Stabilization (MILS)

Airway management of patients with C-spine injury should be performed using the MILS technique to minimize cervical movement, and rapid sequence intubation should be applied. To perform the MILS exactly, the assistant must maintain the patient's head in a neutral position while fixing it against the force exerted during endotracheal intubation [51]. The MILS method is the best way to effectively limit the movement of the C-spine at the lowest cost. In a study using magnetic resonance imaging, De Lorenzo et al. [52] reported placing a pillow approximately 2-cm high beneath the head increases the spinal canal/spinal cord ratio at the C5-6 levels, a region that is injured frequently. The application of the MILS maneuver decreased C-spine extension when exposing the arytenoid under a direct laryngoscope [53]. However, the MILS technique does not completely eliminate C-spine movement. Lennarson et al. [54] compared C-spine movement when MILS was performed, and it was not eliminated while applying 7–10 pounds of traction in a cadaver with C4-5 transection. C-spine extension was less in the case of traction. However, cervical subluxation was increased by applying the MILS technique. Distraction was reported to increase during traction and decrease during MILS. Therefore, compared with emergency situations in which the location of damage is unknown because the site of injury has not been assessed, the application of the MILS technique reduces cervical extension.

Laryngoscopic View according to Neck Stabilization Technique

Patients with cervical spine injuries have limitations when opening their mouth for intubation or to position an airway because they are wearing cervical collars [55]. The anterior portion of the cervical collar must be removed, and another anesthetic assistant must hold the patient's head on the bedside to minimize patient movement during intubation [51]. It is known that MILS reduces mouth opening and worsens the laryngoscopic view [56]. By performing MILS, the rate of Cormack-Lehane grade worsening of >1 grade is 35.6%, while the possibility of it worsening >2 grades is approximately 10% [57]. In addition, mouth opening decreases by approximately 10 mm depending on the type of cervical collar [55].

C-spine Movement during Intubation

Sawin et al. [58] used fluoroscopy to record a movie of C-spine movement in a normal patient during intubation. At the time of blade insertion, the spine moved little, but when the blade was elevated for vocal cord exposure, extension occurred from the occiput to the C1 level and flexion occurred at the C2-C5 level. However, using a video laryngoscope for intubation without direct visualization and aligning the oral, laryngeal, and tracheal axis could minimize the movement of the cervical spine.

Comparison by Airway Maintenance Equipment in Cervical Spine Patients

Completely transecting cadavers at the C5-6 level, and compared to the spinal movement using the Miller straight blade, Macintosh curved blade, and Corazelli-London-Mccoy hinged blade, the MILS technique showed significantly lower anteroposterior (AP) displacement than the cervical collar [59]. In terms of AP and axial movement with MILS, use of the Miller straight blade showed better results than the Macintosh curved blade and Corazelli-London-Mccoy hinged blade. Watts et al. [60] conducted a study on 29 patients undergoing general anesthesia using the Bullard laryngoscope and Macintosh laryngoscope, and compared C-spine extension, time to intubation, and grade of laryngeal view between patients using and not using MILS.

### Table 2. Common Cervical Spine Fractures according to Injury Mechanism

| Injury mechanism       | Common cervical spine fracture site                                                                 |
|-----------------------|-------------------------------------------------------------------------------------------------------|
| Axial compression mechanism | Jefferson fracture (unilateral or bilateral fractures of the anterior and posterior arches of C1) |
| Multiple mechanisms   | Odontoid fractures                                                                                 |
|                       | - Type 1: avulsion fracture of the tip of the dens                                                   |
|                       | - Type 2: localized fracture of the base of the dens                                                   |
|                       | - Type 3: extended fracture into the C2 body                                                           |
| Flexion mechanism     | C5 flexion tear-drop fracture (anterior inferior aspect of the C5 body)                               |
|                       | Bilateral facet dislocation                                                                          |
|                       | Clay shoveler fracture (an avulsion fracture of the C7 spinous process)                               |
| Extension mechanism   | Hangman’s fracture (pedicle fractures of the C1 or C2)                                                |
|                       | C2 extension tear-drop fracture (avulsion fracture of the anterior inferior aspect) of the body      |
Under MILS, use of the Bullard laryngoscope reduced C-spine extension at all spine levels during intubation but increased the intubation time to about twice that using the Macintosh blade. Using a new type of video-laryngoscope called the AirWay Scope® (AWS, Pentax, ToKýo, Japan), the extension movement from the occiput to C4 decreased greatly compared with the Macintosh laryngoscope [61]. Another type, the AirTraq laryngoscope (King Medical Systems, Newark, DE, USA), also decreased extension of the C2-C5 spine and C5-thoracic segment spine by up to 66% compared with the Macintosh laryngoscope, while the time to intubation was similar, leading to the conclusion that the AirTraq is better for patients with cervical spine injury [62]. Comparing cervical spine motion during intubation using the Glidescope® (Verathon Inc., Bothell, WA, USA), the Macintosh no. 3 blade, and the lightwand (Trachlight®, Laerdal, Armonk, NY, USA) under MILS, the Glidescope® decreased C2-C5 motion, and the lightwand decreased extension motion at the occiput-C1, C1-C2, C2-C5, and C5-thoracic levels [63]. Comparing direct laryngoscopy and the Glidescope® under the MILS technique in 20 patients without C-spine pathology, 100% of patients using the Glidescope® were viewed as Cormack–Lehane grade 1, 65% of patients using direct laryngoscopy were viewed as Cormack–Lehane grade 2, and 35% of patients using direct laryngoscopy were viewed as Cormack–Lehane grade 3. The Glidescope® offered better glottis visualization than the direct laryngoscope, with the same C-spine movement [64]. Malik et al. [65] performed a study to identify a novel laryngoscope technique using MILS; after comparison, the Pentax AWS showed higher probability of grade 1 on Cormack–Lehane laryngoscope view than the Glidescope®, the Truview EVO2, and the Macintosh blade, indicating that the AWS was the most useful. Additionally, Liu et al. [66] compared the Glidescope® and AWS; the AWS required half the time to complete the tracheal intubation process, with easier intubation. However, in terms of cervical movement, fiber-optic bronchoscope-guided nasal intubation showed the least cervical movement, although it is difficult for novices to perform [67].

C-spine Movement during Pre-intubation Maneuvers

In a cadaver study, we have learned that pre-intubation techniques such as the chin lift or jaw thrust can narrow the space available for a cord. However, the cricoid pressure does not further increase the movement of the cranio-cervical junction [68]. Hauswald et al. [69] reported that mask ventilation requires more attention than any other procedure because it worsens cervical spine displacement.

Summary

In TBI patients, mild hyperoxemia and normocapnia are helpful in preventing secondary brain injury, and use of appropriate sedatives and muscle relaxants is recommended. The management of patients with head trauma should always consider C-spine motion restriction because there is a possibility of head trauma accompanied by cervical spine injury. Performing MILS ensures safe endotracheal intubation, but the procedure itself makes intubation difficult. Endotracheal intubation with a video-laryngoscope is easy to learn and has a good field of view, so its use is increasing. However, the Glidescope® does not decrease cervical movement compared with a direct laryngoscope, and a video laryngoscope has better vocal cord visualization, therefore ensuring a higher rate of successful endotracheal intubation.

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