Traumatic Brain Injury in Children under Age 24 Months : Analysis of Demographic Data, Risk Factors, and Outcomes of Post-traumatic Seizure

Sang-Youl Yoon, M.D., Yeon-Ju Choi, M.D., Seong-Hyun Park, M.D., Ph.D., Jeong-Hyun Hwang, M.D., Ph.D., Sung Kyoo Hwang, M.D., Ph.D.

Department of Neurosurgery, Kyungpook National University Hospital, Daegu, Korea

Objective : Traumatic brain injury (TBI) in children under age 24 months has characteristic features because the brain at this age is rapidly growing and sutures are opened. Moreover, children this age are completely dependent on their parents. We analyzed the demographic data and risk factors for outcomes in TBI patients in this age group to elucidate their clinical characteristics.

Methods : We retrospectively reviewed the medical records and radiological films of children under 24 months who were admitted to Kyungpook National University Hospital from January 2004 to December 2013 for TBI. Specifically, we analyzed age, cause of injury, initial Glasgow coma scale (GCS) score, radiological diagnosis, seizure, hydrocephalus, subdural hygroma, and Glasgow outcome scale (GOS) score, and we divided outcomes into good (GOS 4–5) or poor (GOS 1–3). We identified the risk factors for post-traumatic seizure (PTS) and outcomes using univariate and multivariate analyses.

Results : The total number of patients was 60, 39 males and 21 females. Most common age group was between 0 to 5 months, and the median age was 6 months. Falls were the most common cause of injury (n=29, 48.3%); among them, 15 were falls from household furniture such as beds and chairs. Ten patients (16.7%) developed PTS, nine in one week; thirty-seven patients (61.7%) had skull fractures. Forty-eight patients had initial GCS scores of 13–15, 8 had scores of 12–8, and 4 had scored 3–7. The diagnoses were as follows : 26 acute subdural hematomas, 8 acute epidural hematomas, 7 focal contusional hemorrhages, 13 subdural hygromas, and 4 traumatic intracerebral hematomas larger than 2 cm in diameter. Among them, two patients improved to good-to-moderate disability. Child abuse, acute subdural hematoma, and subdural hygroma were risk factors for PTS in univariate analyses. Multivariate analysis found that the salient risk factor for a poor outcome was initial GCS on admission.

Conclusion : The most common cause of traumatic head injury in individuals aged less than 24 months was falls, especially from household furniture. Child abuse, moderate to severe TBI, acute subdural hematoma, and subdural hygroma were risk factors for PTS. Most of the patients recovered with good outcomes, and the risk factor for a poor outcome was initial mental status.

Key Words : Craniocerebral trauma · Infants · Demography · Risk factors · Seizures · Glasgow outcome scale.

• Received : July 19, 2016 • Revised : December 15, 2016 • Accepted : March 14, 2017

• Address for reprints : Sung Kyoo Hwang, M.D., Ph.D.

Department of Neurosurgery, Kyungpook National University Hospital, 130 Dongdeok-ro, Jung-gu, Daegu 41944, Korea Tel : +82-53-200-5654, Fax : +82-53-423-0504, E-mail : shwang@knu.ac.kr

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INTRODUCTION

Traumatic brain injury (TBI) is one of the most important causes of disability and death in children²⁰⁾. The incidence of infantile injury is little reported and unclear, but it has been reported to be ~124 per 1000 children^{10,13)}. TBI in children under age 24 months has characteristic features that differ from those other age groups. Children this age are completely dependent on their caretakers; additionally, because their skulls are thin, sutures are opened and the brain is not fully maturated, their brain is more vulnerable to external forces. As a result, it is conceivable that pathophysiology and recovery process may be different from those of adults. The incidence of post-traumatic seizure (PTS) is reported to be ~10 to 20% among children with TBI^{1,3,14,20,24)}. Many authors have reported a higher incidence of PTS in young children than in other age groups^{2,7,20,27)}. PTS is a major cause of poor outcomes.

Most reports on pediatric TBI include children of wide age ranges, and few articles focus on the characteristic features of TBI in this infantile age group^{8,9,28)}. Therefore, we conducted this study to elucidate the demographic characteristics and risk factors of PTS and TBI outcomes in very young children.

MATERIALS AND METHODS

We retrospectively reviewed patients under age 24 months who were admitted to Kyungpook National University Hospital from January 2004 to December 2013 for TBI using their medical records and radiological findings. Specifically, we analyzed them for age, cause of injury, initial Glasgow coma scale (GCS) score, radiological diagnosis, development of seizure, hydrocephalus, and Glasgow outcome scale (GOS) score. The radiologic data we analyzed were skull fracture, acute subdural or epidural hematoma, subdural hygroma, traumatic intracerebral hematoma (defined as larger than 2 cm in diameter), focal contusional hemorrhage (defined as smaller than 2 cm), and diffuse axonal injury without significant hemorrhage.

We divided initial GCS scored into three groups: mild (GCS 13–15), moderate (GCS 8–12), or severe (GCS 3–7) head injury, and we divided outcomes into either good (GOS 4–5, good outcome or moderate disability) or poor (GOS 1–3, severe disability or death). We also statistically analyzed the risk factors for PTS and good or poor outcomes; we evaluated the chil-

dren's outcomes at the time of discharge from the hospital or at their last follow-up visits, which varied from one week to two years. The follow-up periods varied so markedly because most of the neurologically intact patients were not followed up for long periods, whereas patients with worse deficits stayed in the hospital or visited regularly for years.

Statistical methods

We calculated significance probability using the chisquared test; if the result was significant and the expected frequencies were less than five in more than 25% of the cells, we calculated p values using Fisher's exact test. We used multiple logistic regression analysis to test the partial effects of the variables that were significant in the univariate analysis and calculated the C-statistics. We conducted the statistical analyses using SAS software (SAS 9.4; SAS Institute Inc., Cary, NC, USA) and considered p values <0.05 to indicate statistical significance.

RESULTS

The total number of patients was 60, 39 males and 21 females; most common age group was between 0 and 5 months (26 cases, 43.3%), followed by 22 patients who were age 6–11 months; additionally, twelve patients were aged between 12 to 23 months. The mean age was 7.5 months and the median was six (Table 1). Regarding the etiology, falls were the most common cause of injuries at 29 (48.3%). Among these, 15 were falls from household furniture such as beds and chairs, comprising 25.0% of all cases (Table 2). On admission, 48 patients (80.0%) had GCS scores of 13 to 15, 8 (13.3%) had scores of 8 to 12, and 4 (6.7%) had scores of 3 to 7 (Table 3).

Skull fracture was found in 37 patients (61.7%) and was the

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Age (months)	Value
0-5	26 (43.4)
6—11	22 (36.7)
12–17	6 (10.0)
18–23	6 (10.0)
Total	60 (100.0)

Values are presented as number (%)

Table 2. Causes of traumatic brain injury

Cause	Value
Fall	29 (48.3)
Traffic accident	12 (20.0)
Abuse	4 (6.7)
Others*	15 (25.0)
Total	60 (100)

Values are presented as frequency (%). *Others included 5 hits by heavy object, 5 slip-downs, 2 hits by parent accidentally (not abuse), and 3 unknowns

Table 3. Distribution of initial GCS score on admission

GCS score	Value
13–15	48 (80.0)
8–12	8 (13.3)
3–7	4 (6.7)
Total	60 (100.0)

Values are presented as frequency (%). GCS : Glasgow coma scale

most common radiological finding. The intracranial diagnoses on computed tomography (CT) and magnetic resonance imaging (MRI) were as follows : 26 acute subdural hematomas, 8 acute epidural hematomas, 13 subdural hygromas, 7 focal contusional hemorrhages, and 4 traumatic intracerebral hematomas.

No patients had diffuse axonal injury without significant hemorrhage (Table 4). Two patients underwent craniotomy to remove hematomas, and 10 patients (16.7%) developed PTS; in nine of those ten, the PTS developed within one week after injury, and six of these patients showed generalized seizure. Two patients (3.3%) developed post-traumatic hydrocephalus, one of whom underwent ventriculo-peritoneal shunt. Four patients, 6.4% of all patients had been victims of child abuse; all of them had had PTS and poor outcomes. Regarding GOS scores, 55 patients showed good outcomes and 5 showed poor outcomes (Table 5).

Risk analysis

We conducted risk analysis regarding PTS and poor outcomes. According to univariate analyses, child abuse, subdural hygroma, acute subdural hematoma, and GOS score were related to a higher incidence of PTS (Table 6). Multivariate analysis revealed that subdural hygroma was a significant risk factor for seizure development (p=0.018, odds ratio [OR]=6.7,

Table 4. Frequencies of diagnosis according to CT or MRI findings

Diagnosis	Value
Skull fracture	37 (61.7)
Acute subdural hematoma	26 (43.3)
Acute epidural hematoma	8 (13.3)
Subdural hygroma	13 (21.7)
Traumatic ICH larger than 2 cm	4 (6.7)
Focal contusional hemorrhage less than 2 cm	7 (11.7)

Values are presented as number (%). CT : computed tomography, MRI : magnetic resonance imaging, ICH : intracerebral hematoma

Table 5. Distribution of GOS score on discharge

GOS outcome	Value
Died	2 (3.3)
Vegetative	1 (1.7)
Severe disability	2 (3.3)
Moderate disability	3 (5.0)
Good outcome	52 (86.7)
Total	60 (100)

Values are presented as frequency (%). GOS : Glasgow outcome scale

95% confidence interval [CI]=1.4–32.1), and patient outcomes were significantly related to traumatic intracerebral hematoma and initial GCS score on univariate analysis (Table 7). Multivariate analysis revealed that only initial GCS score was significantly related to outcomes (p=0.002, OR=26.5, 95% CI=3.2–216.2, c-statistics=0.97).

DISCUSSION

TBI in children is one of the main causes of disability and death. There are few reports of TBI in children aged less than 24 months, and studies most involve children of all age ranges^{8,10,12,13,15,18,26)}. However, children under age 24 months are completely dependent on caretakers, and it is conceivable that the etiology and mechanisms of TBI in these patients could differ from those in other age groups. The thin skull in these very young children is susceptible to fracture; moreover, because their brains grow rapidly, brain injury can result in serious permanent damage.

The incidence of TBI in children in this age range is unclear, but it has been reported to be \sim 124 per 1000 children^{10,13)}. Quayle et al.²⁶⁾ reported a clinical observational study con-

Diagnosis	Number of patients	Number of seizures	<i>p</i> -value
Cause of injury			
Abuse	4	4	
Traffic accident	12	1	0.003 ⁺
Fall	29	2	0.000 [†]
Others	15	3	0.009 [†]
Subdural hygroma	13	6	0.004
Acute subdural hematoma	26	8	0.015
Acute epidural hematoma	8	0	0.330
Skull fracture	37	4	0.161
Hydrocephalus	2	1	0.308
Glasgow outcome (poor) [‡]	5	0	0.578
Glasgow coma scale			
13—15	48	5	
8–12	12	4	0.042 [§]
3–7	4	1	

Table 6. Univariate analysis of risk factors for post-traumatic seizures*

*Factors with p-values larger than 0.5 were not included in the table, [†]Comparison between abuse and each other causes by Fisher's exact test, [‡]"Poor" meant severe disability or death, [§]Incidence was high in moderately to severely injured patients compared with the mild TBI group. TBI : traumatic brain injury

ducted in the United States through the Pediatric Emergency Care Applied Research Network. Among their cohort of 43399 patients, most were children aged one and two, and the most common injury mechanism was falls from heights, comprising 54%, followed by falling down stairs and falling to the ground from standing, walking, or running. Fakharian et al.¹⁰ also reported that falling was the most common cause of injury, and more than 65% of the injuries in that study had occurred at home. However, those authors stated that traffic accidents were an increasingly common cause of injury. In our series, the most common cause of TBI in this age group was falls, and approximately half of them were from household furniture such as beds and chairs. With this in mind, special care should be taken with infants, and safety regarding household furniture cannot be overemphasized. Injuries from falling from household objects were usually mild. Fall from low heights rarely cause significant brain injury, with the exception of epidural hematoma⁸⁾. Kim et al.¹⁸⁾ reported clinical features of TBI in children aged less than 24 months, and the most common mode of injury was stairs followed by in-car

 Table 7. Univariate analysis of risk factors for poor outcomes*

Diagnosis	Number of patients	Number of poor outcomes	<i>p</i> -value
Focal contusional hemorrhage	7	5	0.099
Traumatic ICH	4	2	0.032
Acute subdural hematoma	26	25	0.377
Hydrocephalus	2	1	0.161
Glasgow coma scale score			0.000
13—15	48	0	
8–12	8	2	
3–7	4	3	
Age			0.088
0—5	26	1	
6–11	22	1	
12—17	6	2	
18–23	6	1	

*Factors with p-values larger than 0.5 were not included in the table. ICH : intracerebral hematoma

accidents. The incidence of injury due to child abuse was 2.9%.

A number of authors have reported on non-accidental causes of TBI in this age group^{8,9,28)}. Duhaime et al.⁸⁾ reported that ~24% of head injuries in children aged less than two years were presumed inflicted, and 32% were suspicious for abuse, neglect, or social or family problems on careful interviewing by trauma social workers and ophthalmic evaluation. Children under age two years had worse outcomes than any other age groups. In our series, 4 of 60 cases (6.7%) were due to child abuse. Concern about child abuse is increasing in Korea, but the incidence is still considered low. We diagnosed child abuse based on history taken from caretakers or related persons and general physical examination. It is true that special consideration should be paid to diagnose non-accidental head injury in this age group, but this is problematic; subdural hematoma and retinal hemorrhage can suggest non-accidental injury, but not decisively so. Thus, careful history taking and whole body examinations are important, as is the role of the injury coordinator. It was not conclusive from our series that child abuse had resulted in poorer outcomes than other causes of injury due to the small number of abused patients. However, many papers have reported poor outcomes from abuse⁸⁾. We propose two reasons for the poor outcome in abused children; one is that most of the child abuse patients without mental deterioration did not visit the hospital, and the other is that rotational forces caused their injuries⁹.

Skull fracture was one of the most common features of head injury in our series, comprising 37 of 60 patients; it was associated with relatively mild TBI with good prognoses. It is conceivable that a thin skull is associated with susceptibility to skull fracture. A number of authors reported that the incidence of neurosurgical intervention in children with isolated skull fracture and no neurological deficit was so low that the patients could be discharged after initial evaluation^{12,22,25,29}. However, considering that PTS can result in devastating outcomes and that young children have a high incidence of these seizures, we recommend that patients with skull fracture be closely observed after admission. We routinely take brain CT scan in patients who visit our emergency room after head injury if they are not completely normal or if their parents want us to take CT scans in spite of our warning about possible radiation hazard.

Post-traumatic epilepsy (PTE) develops in 10-20% of children with TBI^{1,3,14,20,24)}. PTS can result in worse outcomes due to secondary brain injury from increased intracranial pressure, hypoxia, and increased metabolic requirements^{6,17)}. A number of authors report that children have a higher incidence of PTS than adults^{2,7,20,27)}. In particular, very young children are more prone to developing PTS. Kieslich and Jacobi¹⁶⁾ reported an increased incidence of early and late seizure (43.8%) in children under age two, who showed greater susceptibility to PTS. Several authors have also reported that subclinical epilepsy developed in a considerable number of patients^{4,22,25)}. Arndt et al.⁴⁾ monitored continuous EEG in pediatric head injury to detect subclinical early PTS and found seizures of any type in 43.7% of their cohort; additionally, 42.9% of the patients in that study, 6.9% of the total population, had subclinical seizures. Penetrating injury, subdural hematoma, and intracerebral hematoma showed a higher incidence than other types of injury. Liesemer et al.²⁰⁾ reported that age below two years, a GCS score below 8, and non-accidental trauma were risk factors for early PTS. In our study, the incidence of seizure was 16.7%, and risk factors were abuse, subdural hygroma, subdural hematoma, and GCS score. Severe TBI is a risk factor for a higher incidence of PTS^{1,6,14,20,27}. Our study also showed a higher rate of seizure development in

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moderate-to-severe than in mild TBI; however, there was no difference between moderate and severe injuries. Subdural hygroma was related to a high incidence of PTS in univariate and multivariate analyses in our series. The reason for the high frequency of PTS in subdural hygroma is unclear because there have been no similar reports regarding any relationship between PTS and subdural hygroma; its clinical courses and significance are usually neglected in studies of TBI. It is conceivable that subdural hygroma that develops as a result of minor laceration of the arachnoid membrane is associated with cortical injury; however, additional studies are needed to clarify this relationship. The incidence also varies according to the etiology of trauma; many authors have reported a higher incidence of PTS in non-accidental injury^{4,15,20,24)}. Our series also showed a higher incidence of PTS in abused children, despite their small number. Subdural hematoma has been reported to be a risk factor for PTS^{1,16,20)}. Considering the high incidence of seizure in infantile TBI, an aggressive approach is needed to prevent early seizure in TBI in infants. Role of antiepileptic drug (AED) in pediatric PTE is not definitely determined^{5,23)}. However, many authors report the effectiveness of AED to prevent early seizure^{11,19,21}. We routinely prescribed the preventive AED to patients admitted in our hospital; however, the duration was variable.

It is well-known that patient outcome is related to GCS on admission. In our series, initial GCS and intracerebral hematoma were significant risk factors on univariate analysis. However, multivariate analysis found that intracerebral hematoma did not contribute to the predictability of poor outcomes. We believe that we need more data to determine the importance of intracerebral hemorrhage as an independent risk factor.

One of the limitations of our study was the small number of patients despite our institution being a tertiary referral hospital with an active pediatric neurosurgical division. Although a high proportion of pediatric TBI patients are under age 24 months, the number of patients in our study was insufficient for reliable analysis. We believe that a cooperative inter-hospital study is required to provide a good reference database for managing and protecting against TBI in very young children. Second limitation was the wide range in follow-up periods; neurologically intact patients were followed for a short time, but patients with disability could be followed for years. Another limitation was diagnosis of subdural hygroma. It is not uncommon that some infant has wide subarachnoid space which reveals similar findings in CT scan and can be differentiated by MRI. However, we did not take MRI routinely and diagnosed the subdural hygroma when the subdural space was definitely wide initially or widened at the follow up CT scan.

CONCLUSION

We assessed the demographic data and risk factors for PTS and outcomes in TBI in children aged less than two years. Falls were the most common cause of injury, especially falls from household furniture. Child abuse, moderate-to-severe TBI, acute subdural hematoma, and subdural hygroma were risk factors for PTS, and injury severity and intracerebral hematoma were risk factors for poor prognoses.

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