Neurodevelopmental Prognostic Factors in 73 Neonates with the Birth Head Injury

Kyoung Mo Kim¹, Sung Min Cho², Soo Han Yoon¹, Yong Cheol Lim¹, Moon Sung Park³, and Mi Ran Kim⁴

¹Department of Neurosurgery, Ajou University Hospital, Ajou University School of Medicine, Suwon, Korea

²Department of Neurosurgery, Wonju Severance Christian Hospital, Yonsei University Wonju College of Medicine, Wonju, Korea ³Department of Pediatrics, Ajou University Hospital, Ajou University School of Medicine, Suwon, Korea

⁴Department of Obstetrics and Gynecology, Ajou University Hospital, Ajou University School of Medicine, Suwon, Korea

Objective: The objective of this study was to reinterpret the neurodevelopmental prognostic factors that are associated with birth head injury by performing a long-term follow-up.

Methods: Seventy-three neonates with head injuries were retrospectively analyzed after a duration of 10.0 ± 7.3 years to determine the correlations between perinatal factors, including gender, head circumference, gestational age, body weight, and mode of delivery, and head injury factors from radiologic imaging with social, fine motor, language, and motor developmental quotients.

Results: There was a statistically significant difference between perinatal factors and head injury factors with respect to head circumference, body weight, gestational age, mode of delivery, Apgar scores at 1 min, cephalohematoma, subdural hemorrhage, subarachnoid hemorrhage, and hypoxic injury, but no direct correlation by regression analysis was observed between perinatal factors and developmental quotients. Of the head injury factors, falx hemorrhage showed a significant indirect relationship with the language and motor developmental quotients. Mode of delivery, subgaleal hematoma, cephalohematoma, greenstick skull fracture, epidural hemorrhage (EDH), tentorial hemorrhage, brain swelling, and hypoxic injury showed an indirect relationship with social development.

Conclusion: In terms of perinatal factors and head injury factors, mode of delivery, subgaleal hematoma, cephalohematoma, greenstick skull fracture, EDH, tentorial hemorrhage, falx hemorrhage, brain swelling, and hypoxic injury displayed an indirect relationship with long-term development, and therefore these factors require particular attention for perinatal care. (Korean J Neurotrauma 2018;14(2):80-85)

KEY WORDS: Birth injury · Falx hemorrhage · Greenstick skull fracture · Neonate · Neurodevelopment · Prognostic factor.

Introduction

The incidence of perinatal traumatic head injuries ranges 2% to 7% of all pregnancies, is responsible for 2% of

Department of Neurosurgery, Ajou University Hospital, Ajou University School of Medicine, 164 Worldcup-ro, Yeongtong-gu, Suwon 16499, Korea

Tel: +82-31-219-5233, Fax: +82-31-219-5238

neonatal deaths, and is reported to be increased in dystocic presentation or instrumental delivery.^{12,19)} In macrosomic infants birth injury increases six-fold, and it has been shown that there is a relationship between Apgar scores and neonatal or infant mortality.^{9,11)} Furthermore, there have been studies which observed that in the long-term subgaleal hematomas leads to increased neonatal or infant mortality rates, long-term observation of birth injury showed increased rates of epilepsy and neurological disabilities.^{6,10)} It is also suggested that there may be a relationship with adolescent attention deficit hyperactivity disorder (ADHD) and mental disease in adults.^{4,17)} Additionally, recent advancement of ultrasound, magnetic resonance imaging (MRI) and computed tomograpy (CT) has enhanced diagnostic efficien-

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Address for correspondence: Soo Han Yoon

E-mail: ee80@hanmail.net

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cies.^{2,3,7,18)} Therefore, it is deemed that there is necessity for statistical investigation into the long-term neurodevelopmental prognostic factors such as perinatal factors and birth head injury factors.

The authors of the current study ascertained the relationship between perinatal factors and birth head injury factors among birth head injury neonates, and compared the longterm follow-up outcomes of developmental quotients which may lead to a reinterpretation of neurodevelopmental prognostic factors.

Materials and Methods

Study design

We included 73 infants who were born at the Ajou University Hospital and who were diagnosed with cephalhematoma, subgaleal hematoma, large cranium, and an Apgar score of less than 6 by CT or MRI for a period of 12 months from July 1, 2006 to June 31, 2007 for this study. We excluded those birth head injuries neonates with congenital anomalies, and hemophiliac disorders from study. A retrospective study of the above neonates with regard to perinatal factors and cranial injury factors, and post-birth development assessment by the Denver test was performed.

Patient data

Perinatal factors included gender, head circumference, gestational age, body weight, mode of delivery, Apgar score at 1 minute, Apgar score at 5 minutes, frontal horn ratio and frontal-parietal width ratio. Among the 73 neonates there were 41 boys and 32 girls. The mean head circumference was 58.3 ± 37.9 percentile, mean gestational age 38.4 ± 2.4 weeks, mode of delivery was vaginal delivery in 49, Cesarean section in 22, and vacuum delivery in 2. The mean body weight was 3034 ± 575 g, and the Apgar score at 1 minute was 7.1 ± 1.8 , and at 5 minutes 8.1 ± 1.7 .

Head injury factors which included cephalohematoma, subgaleal hematoma, linear skull fracture, depressed skull fracture, greenstick skull fracture, epidural hemorrhage (EDH), subarachnoid hemorrhage, tentorial hemorrhage, falx hemorrhage, intracerebral hemorrhage, intraventricular hemorrhage, brain swelling, hypoxic injury were assessed by CT and MRI. Greenstick skull fracture analysis was according to the definitions and classification of Cho et al.⁷¹ Among the 73 neonates, the numbers of patients with linear skull fracture, depressed skull fracture, greenstick skull fracture were assessed. In those neonates with cephalohematoma, subgaleal hematoma, EDH, subarachnoid hemorrhage, tentorial hemorrhage, falx hemorrhage, hypoxic in-

TABLE 1. Frequency of head injury factors in 73 neonates

Head injuries	Lesion (%)
Cephalohematoma	26 (35.6)
Subgaleal hematoma	19 (26.0)
Linear skull fracture	2 (2.7)
Depressed fracture	4 (5.5)
Greenstick fracture	41 (56.2)
Epidural hemorrhage	4 (5.5)
Subdural hemorrhage	17 (23.3)
Subarachnoid hemorrhage	8 (11.0)
Tentorial hemorrhage	36 (49.3)
Falx hemorrhage	27 (37.0)
Intracerebral hemorrhage	4 (5.5)
Intraventricular hemorrhage	2 (2.7)
Brain swelling	47 (64.4)
Hypoxic injury	35 (47.9)

jury, lesions less than 0.5 cm were defined as Grade 0, lesions 0.5 to 1.0 cm as Grade 1, lesions 1.0 to 1.5 cm as Grade 2, and lesions greater than 1.5 cm as Grade 3. The presence of intracerebral hemorrhage or intraventricular hemorrhage that was less than 5 mL was defined as Grade 0, more than 5 mL as Grade 1, 5 to 10 mL as Grade 2, and greater than 10 mL as Grade 3. When the distance of the cranial sutural distance in brain swelling was less than 0.5 cm it was defined as Grade 2, and greater than 1.5 cm as Grade 1, 1.0 to 1.5 cm as Grade 2, and greater than 1.5 cm as Grade 3. The frequency of head injury factors are shown in Table 1.

Follow-up developmental examinations were performed by the Denver test at the outpatient clinic visit, and consisted of social, fine motor, language, and motor functions which were compared with the normal developmental age that were calculated for relative developmental quotients as the percentage. The mean values for social, fine motor, language, and motor relative developmental quotients were 0.685 ± 0.273 , 0.716 ± 0.255 , 0.623 ± 0.260 , 0.709 ± 0.247 , respectively, the mean follow-up period was 10.0 ± 7.3 years, and the ratio of developmental delay that was less than 0.7 was 34.2%, 41.1%, 57.5%, and 37.5%, respectively.

Statistical analysis

Statistical analyses were performed using a Logistic regression correlation, χ^2 Fisher's exact test, and *p*-values were calculated. Apgar scores less than 7 were considered to be abnormal. When the social, fine motor, language, and motor relative developmental quotients less than 0.6, 0.7, and 1.0 were defined as abnormal and delayed, correlation analysis showed no statistical difference between cut off values of 0.6, 0.7, and 1.0, and we only displayed the cut off value of relative developmental quotients 0.7.

Results

Correlation of perinatal factors and birth head injuries Logistic regression correlation among the perinatal factors including gender, head circumference, gestational age, body weight, mode of delivery, did not show statistical significant relationship with 1 minute Apgar scores, nor with the 5 minute Apgar scores. However, the χ^2 analyses of 1 minute Apgar scores demonstrated that there was statistically significant difference when perinatal and head injury

TABLE	E 2.	Loaistic rearessic	n correlations of	perinatal fac	ctors with head	l iniurv factors
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Head injury factors	Perinatal factors	Estimate*	se	p-value	OR (95% CI)
Cephalohematoma	Gestational age	0.1271	0.1079	0.239	1.133 (0.919-1.403)
	Head circumference	0.0108	0.0068	0.115	1.010 (0.997-1.025)
	Body weight	0.0004	0.0004	0.282	1.000 (1.000-1.001)
Subgaleal hematoma	Gestational age	-0.0322	0.1081	0.766	0.968 (0.783-1.197)
	Head circumference	0.005	0.0072	0.486	1.005 (0.991-1.019)
	Body weight	0.0002	0.0004	0.604	1.000 (0.999-1.001)
Linear fracture	Gestational age	0.066	0.316	0.832	1.069 (0.575–1.987)
	Head circumference	0.0538	0.058	0.357	1.055 (0.941-1.183)
	Body weight	-0.0004	0.0012	0.708	0.999 (0.997-1.002)
Depressed fracture	Gestational age	0.460	0.347	0.185	1.584 (0.802-3.128)
	Head circumference	0.005	0.014	0.727	1.005 (0.977-1.033)
	Body weight	0.001	0.001	0.388	1.001 (0.999-1.003)
Greenstick fracture	Gestational age	0.146	0.100	0.145	1.157 (0.951-1.408)
	Head circumference	0.008	0.006	0.228	1.008 (0.995-1.020)
	Body weight	0.000	0.000	0.321	1.001 (1.000-1.001)
Epidural hemorrhage	Gestational age	0.448	0.343	0.192	1.565 (0.799-3.066)
	Head circumference	0.0022	0.0138	0.873	1.002 (0.975-1.030)
	Body weight	0.00168	0.0011	0.128	1.002 (1.000-1.004)
Subdural hemorrhage	Gestational age	0.0977	0.1222	0.424	1.103 (0.868-1.401)
	Head circumference	0.0178	0.0087	0.041	1.018 (1.001-1.036)
	Body weight	0.0007	0.0005	0.149	1.001 (1.000-1.002)
Subarachnoid hemorrhage	Gestational age	-0.1924	0.1405	0.171	0.824 (0.626-1.087)
	Head circumference	-0.0046	0.0098	0.635	0.995 (0.976-1.015)
	Body weight	-0.0014	0.0006	0.032	0.998 (0.997-1.000)
Tentorial hemorrhage	Gestational age	0.0638	0.0972	0.511	1.065 (0.881-1.290)
	Head circumference	0.0019	0.0062	0.757	1.002 (0.990-1.014)
	Body weight	-0.0003	0.0004	0.433	0.999 (0.999-1.000)
Falx hemorrhage	Gestational age	0.0837	0.1035	0.418	1.087 (0.888-1.332)
	Head circumference	0.0112	0.0068	0.098	1.011 (0.998-1.025)
	Body weight	-0.0002	0.0004	0.537	0.999 (0.999-1.001)
Intracerebral hemorrhage	Gestational age	0.0341	0.2185	0.876	1.034 (0.674–1.588)
	Head circumference	0.0025	0.0139	0.852	1.002 (0.976-1.030)
	Body weight	-0.0011	0.0008	0.184	0.998 (0.997-1.001)
Intraventricular hemorrhage	Gestational age	-0.2352	0.2517	0.350	0.790 (0.483-1.295)
	Head circumference	-0.0236	0.0222	0.287	0.976 (0.935-1.020)
	Body weight	-0.0029	0.0016	0.071	0.997 (0.994-1.000)
Brain swelling	Gestational age	0.119	0.1002	0.235	1.126 (0.926-1.371)
	Head circumference	-0.0028	0.0065	0.663	0.997 (0.984-1.010)
	Body weight	0.0002	0.0004	0.630	1.0002 (0.999-1.001)
Hypoxic injury	Gestational age	-0.434	0.1306	0.001	0.647 (0.502–0.837)
	Head circumference	-0.0216	0.00696	0.002	0.978 (0.965-0.992)
	Body weight	-0.0011	0.0004	0.013	0.994 (0.998-1.000)

*Regression coefficient. se: standard errors, OR: odds ratio, CI: confidence interval

factors were compared with respect to greenstick skull fracture, tentorial hemorrhage, brain swelling, and hypoxic injury. When the 5 minutes Apgar scores were analyzed by χ^2 /analysis of variance, the comparison of perinatal and head injury factors in terms of mode of delivery, brain swelling, and hypoxic injury showed significant difference. cumference, subarachnoid hemorrhage and body weight, hypoxic injury and gestational age, hypoxic injury and head circumference, and hypoxic injury and body weight (Table 2).

Correlation of perinatal factors and birth head injuries with developmental quotients

There was also significant logistic regression correlation difference between perinatal factors and head injury factors with respect to subdural hemorrhage and head cir-

There was no significant association of perinatal factors such as gender, head circumference, gestational age, body

Developmental quotients	Perinatal factors	Estimate*	se	p-value	OR (95% CI)
Social	Sex	1.743	1.145	0.127	5.714 (0.606-53.904)
	Mode of delivery	-0.725	1.09	0.506	0.484 (0.057-4.100)
	Gestational age	-0.043	0.183	0.812	0.957 (0.668–1.371)
	Head circumference	0.004	0.013	0.737	1.004 (0.980-1.030)
	Body weight	-0.000	0.000	0.548	0.999 (0.998-1.001)
Fine motor	Sex	-0.182	0.693	0.793	0.833 (0.214-3.244)
	Mode of delivery	-0.242	0.676	0.720	0.784 (0.208-2.958)
	Gestational age	0.077	0.149	0.606	1.080 (0.806-1.448)
	Head circumference	0.005	0.009	0.566	1.005 (0.987-1.024)
	Body weight	0.000	0.000	0.542	1.000 (0.999-1.002)
Language	Sex	1.42	1.18	0.228	4.137 (0.410-41.796)
	Mode of delivery	-0.449	1.105	0.684	0.638 (0.073-5.567)
	Gestational age	0.165	0.252	0.511	1.179 (0.720-1.934)
	Head circumference	0.001	0.013	0.938	1.001 (0.974-1.028)
	Body weight	0.000	0.000	0.528	1.000 (0.999-1.003)
Motor	Sex	0.852	0.772	0.270	2.345 (0.516-10.662)
	Mode of delivery	0.073	0.691	0.915	1.075 (0.278-4.170)
	Gestational age	0.038	0.159	0.808	1.039 (0.761-1.420)
	Head circumference	-0.008	0.009	0.414	0.991 (0.973-1.011)
	Body weight	0.000	0.001	0.847	1.000 (0.999-1.001)

*Regression coefficient. se: standard errors, OR: odds ratio, CI: confidence interval

TABLE 4. Fisher's exact test for correlations of head injury factors with developmental quotients

	p-values on developmental quotients				
Head injury lactors	Social	Fine motor	Language	Motor	
Cephalohematoma	1.000	1.000	0.290	0.703	
Subgaleal hematoma	0.316	0.437	0.567	1.000	
Linear fracture	1.000	1.000	1.000	1.000	
Depressed fracture	1.000	0.641	0.206	1.000	
Greenstick fracture	1.000	1.000	0.626	0.453	
Epidural hemorrhage	1.000	1.000	1.000	1.000	
Subdural hemorrhage	1.000	0.435	1.000	0.672	
Subarachnoid hemorrhage	1.000	1.000	1.000	0.586	
Tentorial hemorrhage	1.000	0.085	1.000	0.056	
Falx hemorrhage	1.000	0.080	0.033	0.023	
Intracerebral hemorrhage	0.252	1.000	1.000	0.378	
Intraventricular hemorrhage	0.133	1.000	1.000	1.000	
Brain swelling	0.649	0.737	1.000	0.703	
Hypoxic injury	1.000	0.738	0.616	0.713	

Perinatal factors and	p-values on developmental quotients					
head injury factors	Social	Fine motor	Language	Motor		
Mode of delivery	0.039	0.638	0.240	0.417		
Subgaleal hematoma	0.036	0.218	0.842	0.511		
Cephalohematoma	0.026	0.202	0.919	0.971		
Greenstick skull fracture	0.033	0.169	0.704	0.666		
Epidural hemorrhage	0.036	0.237	0.829	0.818		
Tentorial hemorrhage	0.028	0.308	0.783	0.856		
Brain swelling	0.025	0.055	0.328	0.089		
Hypoxic injury	0.001	0.001	0.116	0.002		

TABLE 5. Significant correlations of perinatal factors and head injury factors with developmental quotients χ^2

weight, mode of delivery, with social, fine motor, language, motor developmental delay (Table 3). Fisher's exact test of head injury factors such as social, fine motor, language, and motor development demonstrated significant relationship with falx hemorrhage and language development, and falx hemorrhage and motor development (Table 4). When the 4 developmental quotients were compared by χ^2 for perinatal factors and head injury factors, mode of delivery, subgaleal hematoma, cephalohematoma, greenstick skull fracture, EDH, tentorial hemorrhage, brain swelling, and hypoxic injury were significant factors for social development. Fine motor and motor development was only significant in hypoxic injury, while language development was nor significant in any fields (Table 5).

Discussion

Correlation of perinatal factors and birth head injuries

It has been shown in previous studies among the perinatal factors - mode of delivery, time of birth, low Apgar score at 5 minutes, gestational age at birth are associated with the neonatal death rate.^{9,15)} Proposed predictors of birth trauma has been suggested to decreased fetal heart rate, shoulder dystocia, male gender, neonatal weight, neonatal head circumference, prolonged gestation, macrosomia, low Apgar scores, delivery during risk hours, instrumental delivery, fundal pressure, vaginal delivery, delivery by resident, induction of labor, second stage labor exceeding 60 minutes, epidural anesthesia, parity, and maternal age and delivery in a teaching hospital.^{1,13,14)} In contrast, maternal age, ethnicity, diabetes, and operative vaginal delivery have not been related to birth trauma.¹⁶⁾

According to the results of our current research, the correlation analysis between perinatal factors and head injury factors and Apgar scores showed statistical relationships of Apgar scores with mode of delivery, greenstick skull fracture, tentorial hemorrhage, brain swelling, or hypoxic injury. In addition, the statistical analysis between perinatal factors and head injury factors revealed that head circumference, body weight, gestational age, and mode of delivery were associated with perinatal factors such as subdural hemorrhage, subarachnoid hemorrhage, hypoxic injury, and cephalohematoma. Therefore, while there is no direct relationship between most perinatal factors and Apgar scores, it is postulated that there may be an indirect association between Apgar scores and perinatal factors because relationships exist between perinatal factors and Apgar scores and between many head injury factors and Apgar scores.

It has been suggested in many previous researches that a large fetal head will result in increased compression during passage through the birth canal, subsequently leading to more frequent head injuries. Likewise, our observation in the present study shows that increase of the head circumference by 1 cm will increase subdural hemorrhage by 1.018 fold, increase of newborn body weight by 1 kg will worsen subarachnoid hemorrhage by 0.9985 fold, longer gestational age by 1 week and increase of head circumference by 1 cm and increase of newborn body weight by 1 kg resulted in increased hypoxic injury by 0.65, 0.98, and 0.99 fold, respectively. It is therefore envisaged that newborn head size governing perinatal factors such as gestational age, body weight, head circumference all contribute partially to birth head injuries of newborns.

Correlation of perinatal factors and birth head injuries with developmental quotients

Perinatal factors or injury such as low birth weight, reduced head circumference, incubator, hypoxic-ischemic brain injury or intraventricular hemorrhage not only increases neonatal mortality rates but also may attribute to severe neurological impairment including cerebral palsy, seizures, mental retardation, and learning disabilities, or may be related to mental diseases such as ADHD, autism, and schizophrenia in the long-term period.^{3,5,8,17)} Among the 4 development fields, social development showed significant differences in many perinatal factors and head injury factors and motor and language development showed difference only in the falx hemorrhage factor, while fine motor did not show such difference over any factors. Therefore, detailed caution should be given to birth injuries such as mode of delivery, subgaleal hematoma, cephalohematoma, greenstick skull fracture, EDH, tentorial hemorrhage, falx hemorrhage, brain swelling, and hypoxic injury as long-term development differences may arise. Access from other perspectives are envisaged to be necessary pertaining to social development.

Conclusion

The perinatal and cranial injury factors including mode of delivery, subgaleal hematoma, cephalohematoma, greenstick skull fracture, EDH, tentorial hemorrhage, falx hemorrhage, brain swelling, hydrocephalus, and hypoxic injury showed indirect relationship with long-term development in this study. Therefore, particular attention requires especially these perinatal and head injury factors in perinatal care.

The authors have no financial conflicts of interest.

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