### ORIGINAL ARTICLE / ОРИГИНАЛНИ РАД

# Cranial ultrasound as a complementary method to the general movements assessment in preterm infants for predicting the neurological outcome – a single center experience

THE TRAPCKOR

Dragan Zlatanović<sup>1,2</sup>, Hristina Čolović<sup>1,2</sup>, Vesna Živković<sup>1,2</sup>, Anita Stanković<sup>2</sup>, Milica Kostić<sup>2</sup>, Vuk Pejčić<sup>2</sup>, Vukota Radovanović<sup>2</sup>

<sup>1</sup>University of Niš, Faculty of Medicine, Department for Physical Medicine and Rehabilitation, Niš, Serbia; <sup>2</sup>Niš University Clinical Centre, Physical Medicine and Rehabilitation Clinic, Niš, Serbia

#### SUMMARY

**Introduction/Objective** Implementing cranial ultrasound (CUS) into daily clinical practice represents a major advance in the diagnosis and treatment of newborns. Preterm birth is considered a risk factor for abnormal neurological development. The study aimed to evaluate the significance of CUS in preterm infants as a complementary method to the General Movements Assessment for predicting neurological outcomes. The study focused on a cohort of infants without significant neonatal morbidity.

**Methods** The study included 160 preterm infants and was designed as a prospective clinical study. Statistical analysis included cranial ultrasound findings and the assessment of spontaneous motor activity in the first five days after birth ("Writhing" period), perinatal data, and pregnancy data.

**Results** There was a statistically significant association between abnormal CUS findings and cerebral palsy in the final neurological outcome (p < 0.001). Pathological CUS findings were significantly more frequent in preterm infants born before 30 weeks of gestation (p < 0.001), those delivered by cesarean section (p < 0.001), and infants with an Apgar score < 8 at one and five minutes (p < 0.001). The specificity of a normal CUS was 86% but increased to 100% when combined with a normal General Movements Assessment. **Conclusion** This research confirms that CUS can be a valuable tool for predicting neurological outcomes in preterm infants. It can provide data that can guide the judicious use of different monitoring methods

and rationalize their examinations. **Keywords:** preterm infants; cranial ultrasound; General Movements; Prechtl's method; neurodevelopmental outcome

#### INTRODUCTION

Motor development in newborns, infants, and young children relies on the health of the central nervous system (CNS), and is influenced by genetic patterns and external stimuli. Mental and motor development are closely connected, with significant neurological changes occurring within the first days and months of life. To assess these changes accurately, repeated evaluations, or developmental monitoring, are essential, as a single assessment may not detect certain neurological issues due to the immature CNS's variable responses [1, 2, 3].

Special clinical attention is drawn to newborns who have some risk of developing disorders of the CNS. The most common perinatal factors are prematurity, low birth weight of the newborn, low Apgar score, multiple pregnancies, birth trauma, inadequate presentation of the fetus, and termination of delivery by cesarean section (C-section) [3–6]. Premature newborns (born before the gestational age of 37 weeks) have a higher risk of sudden death syndrome and complications in general, compared to full-term infants [7, 8, 9].

Apart from the neurological and neurokinesiological examination of newborns and infants, additional diagnostic methods used in clinical practice to detect neurological abnormalities are cranial ultrasound (CUS) examination and magnetic resonance imaging. Implementing CUS into daily clinical practice represents a major advance in the diagnosis and treatment of newborns [10–13].

The general motor assessment (GMA) in premature infants is defined as a high-certainty method for predicting neurological outcomes. Still, this type of examination should only be performed by a trained and certified physician, who is not available in every hospital [14, 15, 16]. For that reason, this study aimed to evaluate the significance of the initial CUS in preterm infants as a widely available, complementary method of examination to GMA for predicting neurological outcomes.

#### **METHODS**

This study was designed as a prospective clinical study and included preterm infants (gestational age  $\leq$  37 weeks), born in the Gynecology and Obstetrics Clinic of the University Clinical Center Niš between 2012 and 2014. During this study period, 7 142 children were born

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#### Correspondence to:

Dragan ZLATANOVIĆ
Department of Physical Medicine
and Rehabilitation
Faculty of Medicine
81 Dr Zoran Đinđić Blvd.
18000 Niš, Serbia

draganzlatanovic1@gmail.com

266 Zlatanović D. et al.

at the University Clinical Center Niš. Among them, 629 (8.8%) were born prematurely. All children who had any serious perinatal complications such as sepsis, necrotizing enterocolitis, and lung disease were excluded from the study. Additionally, some of the preterm infants were excluded from the study due to the presence of deformities or congenital anomalies, as well as genetic syndromes of a newborn, invalid video recordings of neonatal motions, parental refusal to participate in the study, or nonattendance in the follow-up in our institution. A two-year follow-up was completed for 160 preterm infants, and they were analyzed in this study.

For each infant included in the research, detailed perinatal data were taken: sex, gestational age / gestational weeks (GW) (< 30 weeks; 30–37 weeks), body weight at birth, body length at birth, head circumference, Apgar score value at the first and fifth minutes, data on method of delivery (C-section or not), data on multiple (twin) pregnancy and CUS findings.

GMA was carried out according to the basic principles of the Prechtl method within five days after the birth ("Writhing" period) and was based on video analysis and performed by a licensed person for GMA expertise [14, 15, 16]. To get good video recordings (lasting up to 25 minutes), the baby needed to be awake, calm, not crying, with open eyes, without irregular breathing, and moving ("State 4"). For premature babies younger than 36 GW, recordings were made when they started moving, even if they were asleep.

General movements (GMs) were classified into four types [14, 15, 16]:

- 1. Normal Writhing Movements (N): smooth, twisting movements with low to moderate strength and slow to moderate speed, oval or twisting in shape;
- Poor Repertoire (PR): limited variety and less complex than normal (baby starts a movement but doesn't finish it, making the sequence look incomplete or broken);
- Cramped Synchronized Movements (CS): abnormal movements where the muscles of the body and limbs tighten and relax at the same time; movements are stiff and lack the smooth flow of normal writhing movements;
- 4. Chaotic Movements (CH): sudden, jerky movements with very large and random motions of the arms and legs, uncoordinated with a lack of smoothness or pattern

Definitive neurological outcome was assessed based on a detailed neurological examination at the age of 24 months (corrected calendar age). The examination was performed by a certified neurologist specializing in pediatric neurology. Neurological outcome was classified as normal findings (completely normal neurological findings); minimal neurological dysfunction (MND), according to Touwen Infant Neurological Examination criteria or nonspecific signs without clear and definitive signs of cerebral palsy [17, 18]; cerebral palsy (CP) was classified according to Surveillance of Cerebral Palsy in Europe criteria [19].

CUS examination was performed with LOGIQ™ (GE Healthcare, Chicago, Il, USA) machine with a high-frequency linear probe (7–11 MHz) within five days after the birth. Repeat CUS examination was performed two weeks after the birth, but statistical analysis included only the findings of the first examination. All CUS examinations were performed by the same person and categorized into five groups of interest (according to the guidelines from the ELGAN study [13] and Prechtl's recommendations [14, 15, 16]):

- CUS 1 normal finding;
- CUS 2 hyperechogenicity of the brain parenchyma lasting up to 14 days;
- CUS 3 hyperechogenicity of the brain parenchyma that lasts longer than 14 days;
- CUS 4 intraventricular hemorrhage;
- CUS 5 periventricular leukomalacia.

The medical doctor who analyzed spontaneous motor activity in newborns and the medical doctor who analyzed CUS findings did not have access to any results or data on the newborns.

Statistical analysis was conducted using IBM SPSS Statistics, Version 20.0 (IBM Corp., Armonk, NY, USA). The normality of continuous variables in defined groups was determined. Anthropometric mode values were presented as percentiles based on child growth standards. Continuous variable comparisons between groups utilized the Mann-Whitney test, while qualitative variables were analyzed with Pearson's  $\chi^2$  test. For category variables with samples fewer than 5, Pearson  $\chi^2$  or Fisher's exact tests were used. To evaluate the CUS method, sensitivity, specificity, positive predictive value (PPV), negative predictive value, and diagnostic odds ratio (DOR) were calculated based on normal CUS findings and final neurological outcomes. A two-tailed p-value of < 0.05 was considered statistically significant.

**Ethics:** The study was performed in line with the Declaration of Helsinki and approved by the Ethics Board of the University Clinical Center Niš (Decision No. 5718/1).

#### **RESULTS**

A statistically significant difference in neurological outcome after 24 months existed between infants born before and after 30 weeks (p < 0.001). Lower body weight of newborns (p < 0.001), body length (p < 0.05), and Apgar score at the first and fifth minutes (p < 0.001) were statistically significantly associated with a worse neurological outcome. The prevalence of different types of GMs in the observation period of up to five days differed statistically significantly between the different outcomes (p < 0.001). The outcome after a follow-up of 24 months was normal in 124, qualified as MND in 22, and diagnosed as CP in 14 preterm infants (Table 1).

Pathological CUS findings were statistically significantly more common in preterm infants born before 30 GW (p < 0.001), delivered via C-section (p < 0.001), and those with an Apgar score < 8 at both the first and fifth minutes after birth (p < 0.001) (Table 2).

Table 1. Clinical characteristics of subjects according to outcome after 24 months

Clinical characteristics		0	utcome after 24 mont	C			
		Normal (n = 124)	MND (n = 22)	CP (n = 14)	Summarized (n = 160)	р	
<sup>a,b</sup> Gestational age mode (min–max)		35 (29–36) GW	35 (33–35) GW	29 (27–29) GW	35 (27–36) GW	0.014	
≥ 30 weeks		122 (98.39%)	22 (100%)	0 (0%)	144 (90%)	< 0.001	
< 30 weeks		2 (1.61%)	0 (0%)	14 (100%)	16 (10%)	V 0.001	
<sup>a</sup> Sex	Female	60 (48.39%)	10 (45.45%)	8 (57.14%)	78 (48.75%)	0.78	
	Male	64 (51.61%)	12 (54.55%)	6 (42.86%)	82 (51.25%)	0.78	
<sup>a</sup> Twins	No	92 (74.19%)	18 (81.82%)	14 (100%)	124 (77.5%)	0.079	
TWITIS	Yes	32 (25.81%)	4 (18.18%)	0 (0%)	36 (22.5%)	0.079	
<sup>a</sup> Caesarean section	No	80 (64.52%)	12 (54.55%)	6 (42.86%)	98 (61.25%)	0.227	
Caesaleali section	Yes	44 (35.48%)	10 (45.45%)	8 (57.14%)	62 (38.75%)	0.227	
<sup>b</sup> Birth weight (g) mode (min–max) mode in percentiles		2150 (2000–2350) 18.9%	1750 (1350–2400) 3.4%	1320 (1250–1350) 57.1%	2125 (1250–2400) 18.9%	< 0.001	
Birth body length (cm) mode (min–max) mode in percentiles		45 (42–47) 24.5%	44 (43–45) 14.9%	39 (35–40) 55.2%	44 (42–47) 14.9%	0.021	
bHead circumference (cm) mode (min–max) mode in percentiles		30 (29–32) 7.5%	30 (28–32) 7.5%	28 (26–29) 75.2%	30 (26–32) 7.5%	0.114	
bApgar score (1 – minute) mode (min–max)		8 (8–9)	8 (8–8)	6 (1–7)	8 (1–9)	< 0.001	
<sup>b</sup> Apgar score (5– minute) mode (min–max)		9 (8–9)	8 (8–8)	7 (5–8)	9 (5–9)	< 0.001	
<sup>a</sup> GMs within 5 days							
N		94 (75.81%)	0 (0%)	0 (0%)	94 (58.75%)		
PR		30 (24.19%)	22 (100%)	2 (14.29%)	54 (33.75%)	< 0.001	
CS		0 (0%)	0 (0%)	12 (85.71%)	12 (7.5%)		
СМ		0 (0%)	0 (0%)	0 (0%)	0 (0%)		

MND – minimal neurological dysfunction; CP – cerebral palsy; GW – gestational weeks; min – minimum; max – maximum; GM – general movements; N – normal writhing movements; PR – poor repertoire; CS – cramped synchronized movements; CM – chaotic movements;  $\alpha$  etast;  $\alpha$  etast;

**Table 2.** Distribution of normal and abnormal cranial ultrasound findings in relation to clinical characteristics of preterm newborns

		Cranial ultras			
Clinical c	haracteristics	Normal (n = 96)	Abnormal (n = 64)	р	
Cov	Female (n = 78)	52 (66.7%)	26 (33.3%)	0.108	
Sex	Male (n = 82)	44 (53.7%)	38 (46.3%)		
Costationalago	≥ 30 weeks (n = 144)	94 (65.3%)	50 (34.7%)	< 0.001	
Gestational age	< 30 weeks (n = 16)	2 (12.5%)	14 (87.5%)		
Turing	No (n = 124)	76 (61.3%)	48 (38.7%)	0.566	
Twins	Yes (n = 36)	20 (55.6%)	16 (44.4%)	0.566	
Cesarean section	No (n = 98)	70 (71.4%)	28 (28.6%)	< 0.001	
Cesarean section	Yes (n = 62)	26 (41.9%)	36 (58.1%)		
	9 (n = 52)	38 (73.1%)	14 (29.6%)		
Apgar score (1-minute)	8 (n = 82)	52 (63.4%)	30 (36.6%)	< 0.001	
(1-iiiiidte)	< 8 (n = 26)	6 (23.1%)	20 (76.9%)		
	9 (n = 90)	64 (71.7%)	26 (28.9%)		
Apgar score (5-minute)	8 (n = 52)	30 (57.7%)	22 (42.3%)	< 0.001	
(5 minute)	< 8 (n = 18)	2 (11.1%)	16 (88.9%)		

Pearson's χ<sup>2</sup> test

Among six preterm infants with an Apgar score of 1 in the first minute, four scored 2 and two scored 5 in the fifth minute. Four developed CP, with two having scores of 1 and 2, and two with scores of 1 and 5. All CP cases showed hyperechogenicity of brain parenchyma detectable

for up to 14 days. Additionally, two preterm infants diagnosed with MND had scores of 1 in the first minute and 2 in the fifth minute, with hyperechogenicity lasting longer than 14 days. Due to the small sample size, more research is necessary to draw definitive conclusions about this subgroup.

Previously determined statistically significant clinical characteristics of preterm infants for the final outcome also showed a statistically significant relation with CUS findings (Table 3).

Table 4 indicates that deviations from normal CUS findings are smallest in subjects with a normal final neurological outcome. A significant association was observed between CP in the final outcome and abnormal CUS findings (p < 0.001). Pathological CUS findings varied significantly among the groups based on final neurological outcomes (normal, MND, CP) (p < 0.05). Altered findings were most prevalent in subjects with CP

and least frequent in those with a normal outcome.

Table 5 indicates that a normal CUS finding has a high specificity of 72.2% for a normal final neurological outcome, though its sensitivity is lower at 67.8%. The specificity of normal CUS findings improves when assessing

bMann–Whitney test

268 Zlatanović D. et al.

**Table 3.** Distribution of different cranial ultrasound findings in relation to previously significant clinical characteristics of preterm newborns

Clinical characteristics								
		CUS 1 (n = 96)	CUS 2 (n = 36)	CUS 3 (n = 20)	CUS 4 (n = 6)	CUS 5 (n = 2)	р	
Gestational age	≥ 30 weeks (n = 144)	94 (65.3%)	30 (20.8%)	18 (12.5%)	2 (1.4%)	0 (0%)	< 0.001	
	< 30 weeks (n = 16)	2 (12.5%)	6 (37.5%)	2 (12.5%)	4 (25%)	2 (12.5%)		
Cesarean section	No (n = 98)	70 (71.4%)	16 (16.3%)	8 (8.2%)	2 (2%)	2 (2%)	0.002	
	Yes (n = 62)	26 (41.9%)	20 (32.3%)	12 (19.4%)	4 (6.5%)	0 (0%)		
	9 (n = 52)	38 (73.1%)	12 (23.1%)	2 (3.8%)	0 (0%)	0 (0%)	< 0.001	
Apgar score (1-minute)	8 (n = 82)	52 (63.4%)	16 (19.5%)	12 (14.6%)	2 (2.4%)	0 (0%)		
(1-minute)	< 8 (n = 26)	6 (23.1%)	8 (30.8%)	6 (23.1%)	4 (15.4%)	2 (7.7%)		
Apgar score (5-minute)	9 (n = 90)	64 (71.1%)	18 (20%)	8 (8.9%)	0 (0%)	0 (0%)	< 0.001	
	8 (n = 52)	30 (57.7%)	10 (19.2%)	8 (15.4%)	8 (8.9%)	2 (3.8%)		
	< 8 (n = 18)	2 (11.4%)	8 (44.4%)	4 (22.2%)	4 (22.2%)	0 (0%)		

CUS 1 – normal finding; CUS 2 – hyperechogenicity of the brain parenchyma lasting up to 14 days;

CUS 3 – hyperechogenicity of the brain parenchyma that lasts longer than 14 days;

CUS 4 – intraventricular hemorrhage; CUS 5 – periventricular leukomalacia;

Pearson's x2 test

**Table 4.** Distribution of different cranial ultrasound findings in relation to different neurological outcomes

Navwalasiaal	Cranial ultrasound finding						
Neurological outcome	Normal (n = 96)		р				
Normal (n = 124)	84 40 (67.74%) (32.26%)						
MND (n = 22)	10 12 (45.45%) (54.55%)				< 0.001		
CP (n = 14)	2 (14.29%)		12 (85.71%)				
	CUS 1 (n = 96)	CUS 2 (n = 36)	CUS 3 (n = 20)	CUS 4 (n = 6)	CUS 5 (n = 2)		
Normal (n = 124)	84 (67.74%)	28 (22.58%)	12 (9.68%)	0 (0%)	0 (0%)		
MND (n = 22)	10 (45.45%)	4 (18.18%)	6 (27.27%)	2 (9.09%)	0 (0%)	0.003	
CP (n = 14)	2 (14.29%)	4 (28.57%)	2 (14.29%)	4 (28.57%)	2 (14.29%)		

MND – minimal neurological dysfunction; CP – cerebral palsy; CUS 1 – normal finding; CUS 2 – hyperechogenicity of the brain parenchyma lasting up to 14 days; CUS 3 – hyperechogenicity of the brain parenchyma that lasts longer than 14 days; CUS 4 – intraventricular hemorrhage; CUS 5 – periventricular leukomalacia Pearson's  $\chi^2$  test

outcomes without CP or with CP. Combining CUS and GMA achieved 100% specificity and PPV in ruling out deviations from normal neurological outcomes.

#### **DISCUSSION**

Periventricular subependymal hemorrhage and intraventricular hemorrhage are complications that usually occur in the first days after birth and are characteristics of premature babies. Frequency of these hemorrhages in newborns who weigh < 1500 g and are < 32 GW of age is

up to 25%. In newborns who weigh < 1000 g, this frequency is up to 40%. Extremely low birth weight and extremely low gestational age represent a strong predisposition for long-term complications, including CP/MND [9, 20].

Prechtl found that increased brain tissue echodensity is temporary and has limited prognostic value when lasting less than two weeks [12]. Other studies recommend the first CUS on the third day after birth and a follow-up before the end of two weeks [21]. In our study, 96 subjects (60%) had normal CUS findings at the first examination. Among them, 10 had MND, and two were diagnosed with CP. Both infants were born before 30 GW, suggesting that their early preterm birth affected the maturation of their CNS.

Our study on CUS findings shows differences compared to existing literature, likely due to the smaller sample sizes often used in those studies [21]. For instance, a large study of premature infants born after 33 weeks of gestation noted pathological CUS findings in 13%. The authors highlighted that even slight differences in gestational age can affect CUS specificity, complication rates, and neurological outcomes [22]. While some studies report that C-section is not significantly linked to poor neurological outcomes, factors like Apgar scores and head circumference are associated with these outcomes [22, 23]. In contrast, our study found a significant association between C-section delivery and unfavorable neurological outcomes, likely influenced by the overlap of low Apgar scores and head circumferences in C-section cases.

The availability of CUS as a diagnostic method and its application in daily clinical practice represents a major diagnostic advance. In a recent study, normal CUS findings have a PPV for the final normal neurological outcome of 89.4%, and PPV increases to 97.9% for a final neurological outcome that excludes CP. These results indicate that the subjects with a normal finding on the CUS may have a pathological neurological outcome and it correlates with the already described studies. Recent research sup-

ports the fact that the pathological CUS finding has a predictive value for later neurological deviations. The sensitivity of these findings was not high, which indicates the need to follow up on the development of children who, in the first five days after birth, had abnormal findings, either during the examination by CUS or during the GMA. Pathological CUS findings are statistically significantly more frequent in subjects who had CP in the final outcome (p < 0.05).

The diagnostic odds ratio of CUS findings without pathology in all examined combinations is > 1, which implies the ability of the single method to determine the final outcome. DOR doubles in the case of determination of

**Table 5.** Distribution of different cranial ultrasound findings in relation to final neurological outcomes and evaluation of the method significance

		CUS finding		Sensitivity = 67.8%	
		Normal	Abnormal	Specificity = 72.2%	
Neurological	Normal	84 (67.7%)	40 (32.3%)	PPV = 89.4%	p = 0.005
outcome	Abnormal	10 (27.8%)	26 (72.2%)	NPV = 39.4%	
				DOR = 5.42	
		CUS finding		Sensitivity = 63%	
		Normal	Abnormal	Specificity = 85.7%	
Neurological	Non-CP	92 (63%)	54 (37%)	PPV = 97.9%	p = 0.018
outcome	СР	2 (14.3%)	12 (85.7%)	NPV = 18.2%	
				DOR = 10.26	
		CUS and GMA finding		Sensitivity = 51.6%	
		Normal	Abnormal	Specificity = 100%	
Neurological	Normal	64 (48.4%)	60 (51.6%)	PPV = 100%	p < 0.001
outcome Abnormal		0 (0%)	36 (100%)	NPV = 37.5%	
				DOR = infinity	
		CUS and G	MA finding	Sensitivity = 43.9%	
		Normal	Abnormal	Specificity = 100%	
Neurological	Non CP	64 (43.8%)	82 (56.2%)	PPV = 100%	p = 0.038
outcome	СР	0 (0%)	14 (100%)	NPV = 14.6%	
				DOR = infinity	

CUS – cranial ultrasound; CP – cerebral palsy; GMA – general movements assessment; PPV – positive predictive value; NPV – negative predictive value; DOR – diagnostic odds ratio; Pearson's  $\chi^2$  test and determination of the sensitivity, specificity, PPV, NPV, and DOR

non-CP/CP in the final outcome (DOR = 10.26) compared to normal/abnormal determination (DOR = 5.42). A DOR value > 10 indicates excellent diagnostic value. Results combining CUS and GMA are impressive. In the case of determining DOR about non-CP/CP in the final outcome as well as normal/abnormal final outcome, DOR tends to infinity because in no case of both normal CUS findings and normal GMA findings did MND or CP develop.

In this study, 100% of preterm infants with periventricular leukomalacia developed CP, which is in agreement with the research of other authors who indicate the high predictive value of this finding for the later development of CP [24]. All subjects with a finding of intraventricular hemorrhage had one of the pathological outcomes (MND/CP) in the final outcome. Periventricular echodensity in the frontal white matter, which disappeared for up to 14 days, did not affect spontaneous motor activity, but periventricular echodensity in the same zone lasting longer than 14 days was associated with abnormal development of spontaneous motor activity.

A recent study included a limited number of preterm infants due to various exclusion criteria, primarily focusing on those born at or above 30 GW, which comprised 90% of participants. Most infants were not delivered via C-section or from twin pregnancies. After a 24-month follow-up, most subjects showed normal neurological findings; however, the majority of those who developed CP were born before 30 GW (14 out of 16). This suggests that insufficient maturity of the CNS may be a contributing factor. Given that preterm infants with serious complications were excluded from this study, it should be emphasized that results show the prognostic values of CUS and GMA for relatively "low-risk" preterm infants. This fact represents a limiting factor for the generalizability of our results.

Future research should involve more infants (both "low-risk" and "high-risk"), especially those under 30 GW, and a more detailed categorization of CUS and neurological outcomes could enhance the findings' applicability. Studies indicate that preterm infants can show abnormal GM in the first week after birth, often due to factors like electrolyte imbalances or changes in cerebral blood flow, despite later having normal neurological outcomes. To prevent misleading results, it is recommended that the first GM evaluation be conducted after the first week [25]. In our study, we performed the first GMA earlier due to some newborns leaving the maternity ward before this period, following Prechtl's recommendations [14, 15] while excluding those with significant complications. All newborns with normal GM findings at the initial evaluation had a normal neurological outcome at 24 months.

GMA has better predictive value in later periods of development, with the best predictive value in the "fidgety" period (50–54 GW) [14, 15, 16]. Recognizing the impor-

tance of initial examinations and effectively incorporating them into practice is crucial for timely therapy application. This study suggests that developing a prognostic model for predicting neurological outcomes in a larger, multicentric study could be beneficial. This model should integrate GMA and CUS findings along with statistically significant perinatal clinical characteristics and pregnancy data from preterm cases.

#### **CONCLUSION**

This study highlights the strong link between clinical characteristics, CUS findings, and neurological outcomes in preterm infants. Pathological CUS findings were more prevalent in those born before 30 weeks of gestation, delivered by C-section, and with lower Apgar scores. Normal CUS results were highly specific for normal neurological outcomes, especially when combined with GMA. Prolonged hyperechogenicity and conditions like periventricular leukomalacia were significant predictors of CP. These findings underscore the importance of early multimodal diagnostics, particularly CUS and GMA, in predicting long-term neurological outcomes and guiding interventions.

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Conflicts of interest: None declared.

**270** Zlatanović D. et al.

#### **REFERENCES**

- Colver A, Fairhurst C, Pharoah P. Cerebral palsy. Lancet. 2014;383(9924):1240–9. [DOI: 10.1016/S0140-6736(13)61835-8] [PMID: 24268104]
- Fahey MC, MacLennan AH, Kretzschmar D, Gecz J, Kruer MC. The genetic basis of cerebral palsy. Dev Med Child Neurol. 2017;59(5):462–9. [DOI: 10.1111/dmcn.13363] [PMID: 28042670]
- Sato J, Vandewouw MM, Bando N, Branson HM, O'Connor DL, Unger SL, et al. White matter alterations and cognitive outcomes in children born very low birth weight. Neuroimage Clin. 2021;32:102843. [DOI: 10.1016/j.nicl.2021.102843] [PMID: 34601309]
- Kyono Y, Nishiyama M, Kawamura A, Oikawa S, Tokumoto S, Yamaguchi H, et al. Association of perinatal factors with neurodevelopmental referrals in a population-based cohort study in Japan. Sci Rep. 2024;14:3492. [DOI: 10.1038/s41598-024-54167-w] [PMID: 38347064]
- Yoshida T, Matsumura K, Hatakeyama T, Inadera H; Japan Environment, Children's Study Group. Association between Cesarean section and neurodevelopmental disorders in a Japanese birth cohort. BMC Pediatr. 2023;23:306. [DOI: 10.1186/s12887-023-04128-5] [PMID: 37331958]
- Haga M, Kanai M, Ishiguro A, Nishimura E, Minamitani Y, Iwatani A, et al. Changes in in-hospital survival and long-term neurodevelopmental outcomes of extremely preterm infants: a retrospective study of a Japanese tertiary center. J Pediatr. 2023;255:166–74.e4. [DOI: 10.1016/j.jpeds.2022.11.024] [PMID: 36462685]
- Boileau P, Letouzey M, Morgan AS, Lorthe E, Kaminski M, Coquelin A, et al. Circumstances, causes and timing of death in extremely preterm infants admitted to NICU: The EPIPAGE-2 study. Acta Paediatr. 2023;112(10):2066–74. [DOI: 10.1111/apa.16894] [PMID: 37402152]
- Horne RS, Harrewijn I, Hunt CE. Physiology during sleep in preterm infants: implications for increased risk for the sudden infant death syndrome. Sleep Med Rev. 2024;78:101990.
   [DOI: 10.1016/j.smrv.2024.101990] [PMID: 39116607]
- Bourneuf AL, Pabic EL, de Visme S, Levieux K, Pladys P; OMIN study group. Risk factors for unexpected infant death among very premature infants in France. Arch Pediatr. 2024;31(3):195–201. [DOI: 10.1016/j.arcped.2023.12.003] [PMID: 38538469]
- Richer EJ, Riedesel EL, Linam LE. Review of neonatal and infant cranial US. Radiographics. 2021;41(7):E206–7.
   [DOI: 10.1148/rg.2021210089] [PMID: 34723697]
- Gumus M, Oommen KC, Squires JH. Contrast-enhanced ultrasound of the neonatal brain. Pediatr Radiol. 2022;52(4):837– 46. [DOI: 10.1007/s00247-021-05157-x] [PMID: 34333692]
- Plut D, Prutki M, Slak P. The use of contrast-enhanced ultrasound in the evaluation of the neonatal brain. Children (Basel). 2023;10(8):1303. [DOI: 10.3390/children10081303] [PMID: 37628302]

- Kuban K, Adler I, Allred EN, Batton D, Bezinque S, Betz BW, et al. Observer variability assessing US scans of the preterm brain: the ELGAN study. Pediatr Radiol. 2007;37(12):1201–8.
   [DOI: 10.1007/s00247-007-0605-z] [PMID: 17901950]
- Prechtl HF.The neurological examination of the full-term newborn infant. Cambridge (UK): Cambridge University Press; 1991.
- Prechtl HF. Qualitative changes of spontaneous movements in fetus and preterm infant are a marker of neurological dysfunction. Early Hum Dev. 1990;23(3):151–8.
   [DOI: 10.1016/0378-3782(90)90011-7] [PMID: 2253578]
- Zlatanovic D, Čolović H, Živković V, Stanković A, Kostić M, Vučić J, et al. The importance of assessing general motor activity in premature infants for predicting neurological outcomes. Folia Neuropathol. 2022;60(4):427–35.
   [DOI: 10.5114/fn.2022.119593] [PMID: 36734385]
- Hadders-Algra M, Heineman KR, Bos AF, Middelburg KJ. The assessment of minor neurological dysfunction in infancy using the Touwen Infant Neurological Examination. Dev Med Child Neurol. 2010;52(1):87–92. [DOI: 10.1111/j.1469-8749.2009.03305.x] [PMID: 19549207]
- Straathof EJM, Hamer EG, Hensens KJ, La Bastide-van Gemert S, Heineman KR, et al. Development of muscle tone impairments in high-risk infants: associations with cerebral palsy and cystic periventricular leukomalacia. Eur J Paediatr Neurol. 2022;37:12–8. [DOI: 10.1016/j.ejpn.2021.12.015] [PMID: 35007848]
- Surveillance of Cerebral Palsy in Europe (SCPE). Surveillance of cerebral palsy in Europe: a collaboration of cerebral palsy surveys and registers. Dev Med Child Neurol. 2000;42(12):816–24.
   [DOI: 10.1017/S0012162200001511] [PMID: 11132255]
- Cai Y, Li X, Zhao X, Song Y, Zhou W. Risk assessment and early prediction of intraventricular hemorrhage in extremely preterm infants. Sci Rep. 2025;15:17346.
   [DOI: 10.1038/s41598-025-02061-4] [PMID: 40389542]
- Diwakar RK, Khurana O. Cranial sonography in preterm infants: short review of literature. J Pediatr Neurosci. 2018;13(2):141–9. [DOI: 10.4103/jpn.JPN\_60\_17] [PMID: 30090126]
- Ballardini E, Tarocco A, Baldan A, Antoniazzi E, Garani G, Borgna-Pignatti C. Universal cranial ultrasound screening in preterm infants with gestational age 33–36 weeks. Pediatr Neurol. 2014;51(6):790–4. [DOI: 10.1016/j.pediatrneurol.2014.08.012] [PMID: 25456299]
- Syböck K, Hartmann B, Kirchengast S. Maternal prepregnancy obesity affects foetal growth, birth outcome, mode of delivery, and miscarriage rate in Austrian women. Int J Environ Res Public Health. 2023;20(5):4139. [DOI: 10.3390/ijerph20054139] [PMID: 36901147]
- Skovgaard AL, Zachariassen G. Cranial ultrasound findings in preterm infants predict the development of cerebral palsy. Dan Med J. 2017;64(2):A5330. [PMID: 28157062]
- Ploegstra WM, Bos AF, de Vries NK. General movements in healthy full-term infants during the first week after birth. Early Hum Dev. 2014;90(1):55–60. [DOI: 10.1016/j.earlhumdev.2013.10.004] [PMID: 24231002]

## Кранијални ултразвук као комплементарна метода процени општих покрета код превремено рођене деце за предвиђање коначног неуролошког исхода – искуство једног центра

Драган Златановић<sup>1,2</sup>, Христина Чоловић<sup>1,2</sup>, Весна Живковић<sup>1,2</sup>, Анита Станковић<sup>1,2</sup>, Милица Костић<sup>2</sup>, Вук Пејчић<sup>2</sup>, Вукота Радовановић<sup>2</sup>

<sup>1</sup>Универзитет у Нишу, Медицински факултет, Катедра за физикалну медицину и рехабилитацију, Ниш, Србија; <sup>2</sup>Универзитетски клинички центар у Нишу, Клиника за физикалну медицину и рехабилитацију, Ниш, Србија

#### САЖЕТАК

Увод/Циљ Примена ендокранијалног ултразвука (ЕУЗ) у свакодневној клиничкој пракси представља велики напредак у дијагностици и лечењу новорођенчади. Превремени порођај сматра се фактором ризика за поремећај неуролошког развоја. Циљ студије био је да се процени значај ЕУЗ код превремено рођене деце као комплементарне методе процени општих покрета за предвиђање коначног неуролошког исхода. Студија је била фокусирана на кохорту новорођенчади без значајног неонаталног морбидитета.

**Методе** Студија је обухватила 160 превремено рођене деце и осмишљена је као проспективна клиничка студија. Статистичка анализа је обухватила налазе ЕУЗ и процену спонтане моторичке активности у првих пет дана након рођења (*writhing* период), перинаталне податке и податке о трудноћи.

**Резултати** Статистички значајна повезаност постојала је између групе деце код које је церебрална парализа забе-

лежена у коначном неуролошком исходу и абнормалних налаза EУЗ (p < 0,001). Статистички значајно чешћи патолошки EУЗ налаз пронађен је код превремено рођене деце пре 30. недеље гестације (p < 0,001), новорођенчади рођене царским резом (p < 0,001) и оних који су имали Апгар скор мањи од осам у првом и петом минуту након рођења (p < 0,001). Специфичност нормалног EУЗ је 86%, али се повећава на 100% када се комбинује са нормалним налазом опште моторичке активности код превремено рођене деце.

**Закључак** Ово истраживање потврђује да ЕУЗ може бити значајна метода за предвиђање неуролошких исхода, може пружити податке за критичку употребу различитих метода праћења превремено рођене деце и може рационализовати њихове прегледе.

**Кључне речи**: превремено рођена деца; ендокранијални ултразвук; општи покрети; Прехтлова метода; неуроразвојни исход