Neurodynamics derived from early clinical electroencephalography predict neurodevelopmental function in infants with cerebral palsy



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Background and Objectives

Infants with cerebral palsy (CP) have impairments in movement, balance, and posture associated with static brain injuries often sustained before or surrounding birth. CP is primarily a clinical diagnosis, but there has been great interest in developing objective biomarkers to predict prognosis and guide early interventions. Neonates with neurological concerns often receive clinical electroencephalography (EEG) as standard-of-care. We assessed if neonatal EEG corresponded with standardized evaluations, Hammersmith Infant Neurological Examination (HINE) and the Bayley Scale of Infant & Toddler Development, 3rd edition (BAYLEY), in the first three years of life in children with CP.

Materials and Methods

Study Design

Retrospective cohort study

Study Participants & Setting

24 children with CP and neonatal EEG at a tertiary care pediatric hospital

EEG recordings were preprocessed and analyzed for absolute band power, phase-lag index connectivity (PLI), and entropy-based dynamic features (spectral entropy, spectral flatness, fractal dimension, median frequency). Uncorrected **Pearson's correlation coefficients** were calculated from EEG features (yearly average and over-time slope) and clinical measure subscales.

Results - Pearson's Correlation Coefficient

Figure 1. Availability and sampling times of clinical EEGs and clinical evaluations by the Hammersmith Infant Neurological Examination (HINE) and the Bayley Scale of Infant & Toddler Development (BAYLEY) from 24

children with cerebral palsy.



Table 1.1. Correlation coefficients of prior to 1 year of age **EEG power spectral band averages** and Clinical outcome scores prior to 1 year of age (upper panel), at 1 year of age (lower panel).* denotes p-value-0.05. Multiple comparisons were not corrected. Some other correlations are worth reporting within the sample.

HINE scaled scores prior to 1 year of age (n=8)

Subscale Band	Asymmetry	Movements	Nerve	Posture	Reflexes	Tone	Tota
delta	.66	.15	.59	.57	.31	.47	.52
theta	.20	.25	.71*	.75*	.51	.55	.67
alpha1	.17	.34	.69	.76*	.61	.57	.70
alpha2	.10	.21	.66	.65	.53	.49	.61
beta	.14	.10	.57	.55	.37	.42	.50
gamma	11	42	.33	.02	29	02	01
Subso Band	ales HINE:	scaled scores at	1 year of age	e (n=14)		caled score of age (n=14	
	Nerve	Posture	Reflexes	Tone	(Cognitive	
delta	59*	.35	44	.08		.34	
theta	62*	.33	64*	03		.41	
alpha	149	.51	49	.21	.38		
alpha	227	.44	46	.22		.29	
					.25		
beta	01	.38	17	.47		.25	

Table 1.2. Correlation coefficients of prior to 1 year of age EEG band power over-time slope and clinical outcome scores prior to 1 year of age (upper panel), at 1 year of age (lower panel), * denotes p-value-0.05, ** denotes p-value-0.01. Multiple comparisons were not corrected. Some other correlations are worth reporting within the sample.

HINE scale scores prior to 1 year of age (n=5)

Subscales Band	Asymmetry	Movements	Nerve	Posture	Reflexes	Tone	Total
delta	.92*	17	.46	.27	.10	.12	.17
theta	.78	.21	.74	.33	.47	.41	.45
alpha1	.78	06	.47	16	.16	.11	.06
alpha2	.77	.34	.83	.30	.52	.54	.51
beta	.69	.55	.92*	.43	.57	.79	.67
gamma	.66	.57	.95*	.55	.66	.79	.74
HINE scaled scores at 1 year of age BAYLEY scaled scores at 1 year of age (n=9) (n=10)							
Subscales	Δsymme	try Con	nitive (n=10)	Fine (Gross Ian	anage	Total

Subscales	Asymmetry	Cognitive (n=10)	Fine Motor	Gross Motor	Language Expressive	Total
delta	17	12	.75*	10	.20	.11
theta	42	22	.89**	.14	.44	.36
alpha1	35	54	.73*	.04	.40	.23
alpha2	19	73*	.86**	.17	.24	.38
beta	16	69*	.31	.52	.03	.54
gamma	23	62	.04	.56	.02	.51

Results

		HINE scaled scores prior to 1 year of age (n=8)						
Subscales Band	Asymmetry	Movements	Nerve	Posture	Reflexes	Tone	Total	
delta	.03	56	95***	71*	50	85**	82*	
theta	06	57	95***	76*	50	89**	84**	
alpha1	.02	60	92**	64	43	81*	77*	
alpha2	17	65	86**	71*	45	86**	80*	
beta	41	42	49	68	26	68	59	
gamma	45	.07	.39	03	.22	.10	.17	

gamma	45		.39	03
Subscales	HINE scale scores 1 year of age (n=14)	BAYLEY scaled of age	year	
	Asymmetry	Cognitive	Fine Mo	tor
delta	.10	.66**	.49	
theta	47	.28	.29	
alpha1	33	.41	.27	
alpha2	50	.25	.26	
beta	07	54*	.02	
gamma	.12	38	.02	

Table 2.1. Correlation coefficients between prior to 1 year of age EEG connectivity band average and clinical outcome scores prior to 1 year of age (upper panel), at 1 year of age (uoyer panel), denotes p-value-0.05, "denotes p-value-0.01", "denotes p-value-0.01", "denotes p-value-0.01", which was the correlations are worth reporting within the sample

Table 2.2. Correlation coefficients between EEG Band Connectivity Slope and BAYLEY subscale scores at 1 year of age. * denotes p-value-0.05, ** denotes p-value-0.01. Multiple comparisons were not corrected. Some other correlations are worth reporting within the sample.

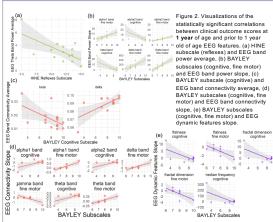
BAYLEY scaled scores 1 year of age (n=9)									
Subscales Band	Cognitive (n=10)	Fine Motor	Language Composite	Language Total	Language Expressive	Language Receptive			
delta	.54	.80*	21	20	16	23			
theta	.70*	.77*	.43	.44	.49	.36			
alpha1	.65*	.83**	.08	.09	.13	.04			
alpha2	.68*	.65	.33	.34	.37	.28			
beta	.28	65	.44	.43	.41	.42			
gamma	.18	75*	.34	.33	.30	.35			

Table 3.1. Correlation coefficients of prior to 1 year of age EEG dynamic feature average with HINE subscale scores prior to 1 year of age. * denotes p-value-0.05. ** denotes p-value-0.01 Multiple comparisons were not corrected. Some other correlations are worth reporting within the sample.

	HINE scaled scores prior to 1 year of age (n=8)									
Subscales Feature	Movements	Nerve	Posture	Reflexes	Tone	Total				
Entropy	61	80*	74*	44	89**	80*				
Flatness	56	87**	80*	48	86**	82*				
HFD	46	80*	77*	45	76*	75*				
MedFreq	60	84**	70	42	86**	78*				

Table 3.2. Correlation coefficients of prior to 1 year of age **EEG dynamic feature slope** with HINE and BAYLEY subscale scores at 1 year of age. * denotes p-value<0.05. Multiple comparisons were not corrected. Some other correlations are worth reporting within the sample.

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Subscales Feature	Asymmetry	Movement s	Postur e	Tone	Cognitiv e (n=10)	Fine Motor	Gross Motor	Language Composite	Language Total
Entropy	38	.54	34	.36	45	58	.34	.43	.42
Flatness	86	.47	.29	.24	65*	70*	.17	13	14
HFD	40	06	.51	08	67*	69*	.04	32	33
MedFreq	37	.60	27	.45	66*	47	.51	.20	.19



Conclusions and Significance

Early clinical EEGs demonstrate predictive value for neurological and developmental outcomes prior to 1 year and during 1 year of age in infants diagnosed with CP. Quantitative analysis of available clinical EEGs months prior to any structured clinical evaluations may inform prognosis and/or referral early intervention.

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