## **Supplementary Online Content**

Novak I, Morgan C, Adde L, et al. Early, accurate diagnosis and early intervention in cerebral palsy: advances in diagnosis and treatment. *JAMA Pediatr*. Published online July 17, 2017. doi:10.1001/jamapediatrics.2017.1689

eTable. Major Diagnostic Advances in Cerebral Palsy Best-Available Evidence eFigure. PRISMA Diagram of Study Flow

This supplementary material has been provided by the authors to give readers additional information about their work.

eTable. Major Diagnostic Advances in Cerebral Palsy Best-Available Evidence

Tool	Citation	Type of Evidence	Numbe r Studie s	Numbe r Patient s	Populatio n	Age at Exam	Accuracy of Detecting Risk for Cerebral Palsy	QUADA S Quality
MOTOR DYS				1			-	T .
AIMS Norm- referenced gross motor assessment	Heinema n 2008	Systemati c review	2 AIMS	Unable to calculat e from review	Newborn high risk for CP & other diagnoses	3-18 month s	Cerebral Palsy Authors conclude moderate predictive validity but no values reported	14/14
	Spittle 2008	Systemati c review	2 AIMS	205	Preterm newborn high risk for CP	4, 6 & 8 month s	Cerebral Palsy No data available in review  Abnormal Motor Sensitivity = 77-86% Specificity = 81-93%	14/14
GMs Quality of movement assessment	Bosanque t 2013	Systemati c review with meta- analysis	6 GMs	1358	Newborn high risk for CP	3-5 month s CA	Cerebral Palsy Sensitivity = 98% Specificity = 91%  A trajectory of longitudinal assessments are more predictive  GMs with MRI provided more accurate prognostic information than the individual tools	14/14
	Burger	Systemati	17 <i>GMs</i>	1820	Newborn	3-5	Cerebral	14/14

	2009	c review			high risk for CP	month s CA	Palsy Sensitivity = 92% Specificity = 82%	
	Darsaklis 2011	Systemati c review	39 <i>GMs</i>	Unable to calculat e from review	Newborn high risk for CP & other diagnoses	3-5 month s CA	Cerebral Palsy Sensitivity = 100% Specificity = 100%	14/14
							Development al Delay Sensitivity = 38-63% Specificity = 54%	
Tool	Citation	Type of Evidence	Numbe r Studie s	Numbe r Patient s	Populatio n	Age at Exam	Accuracy of Detecting Risk for Cerebral Palsy	QUADA S Quality
GMs continued	Heinema n 2008	Systemati c review	7 GMs	Unable to calculat e from review	Newborn high risk for CP & other diagnoses	3-18 month s	Cerebral Palsy Authors conclude good predictive validity but no values reported	14/14
	Spittle 2008	Systemati c review	5 GMs	344	Preterm, term & IUGR newborn high risk for CP	3-5 month s CA	Cerebral Palsy Sensitivity = 83-100% Specificity = 57-96%	14/14
DAYC Parent- reported scored checklist of motor skills	Maitre 2013	Retro- spective chart audit	1 DAYC	606	Preterm, and HIE newborn high risk for CP	6, 12, 24 month s CA	Cerebral Palsy Predictive C- index = 89%  Falling DAYC scores 6-12 months Probability for CP = low for score decreases within 1 SD of	N/A

							the mean i.e. 15 points Probability for CP = high (35%), as the score decreases (>1SD) i.e. 20 points Probability for CP = very high (83%), as the score decreases (>2SD) i.e. 30 points	
MAI Standardize d motor assessment , also assessing tone and reflexes	Heinema n 2008	Systemati c review	7 MAI	Unable to calculat e from review	Newborn high risk for CP & other diagnoses	3-18 month s	Cerebral Palsy Authors conclude moderate predictive validity but no values reported	14/14
	Spittle 2008	Systemati c review	5 MAI	856	Preterm, term newborn high risk for CP & social risk	4 & 8 month s	Cerebral Palsy Sensitivity = 73-96% Specificity = 62-78% Abnormal Motor Sensitivity = 63-85% Specificity = 53-93%	14/14

Tool	Citation	Type of Evidence	Numbe r Studie s	Numbe r Patient s	Populatio n	Age at Exam	Accurac y of Detectin g Risk for Cerebral Palsy	QUADA S Quality
NSMDA Standardize d gross and fine motor assessment	Spittle 2008	Systematic review	1 NSMD A	148	Low birth weight infants	1, 4, 8, 12 month s	Cerebral Palsy No data available in review  Abnormal Motor at 1 month: Sensitivit y = 68% Specificit y = 72%  Abnormal Motor at 4 months: Sensitivit y = 80% Specificit y = 56%  Abnormal Motor at 8 months: Sensitivit y = 82% Specificit y = 82% Specificit y = 83%	14/14
PDMS-2 Norm- referenced gross and fine motor assessment	Heinema n 2008	Systemati c review	2 PDMS- 2	Unable to calculat e from review	Newborn high risk for CP & other diagnoses	3-18 month s	Cerebral Palsy Authors conclude moderate predictive validity but no values reported	14/14
TIMP Standardize d motor assessment	Heinema n 2008	Systemati c review	5 TIMP	Unable to calculat e from	Newborn high risk for CP & other	3-18 month s	Cerebral Palsy Authors conclude	14/14

			review	diagnoses		good predictive validity but no values reported	
Spittle 2008	Systemati c review	3 TIMP	192	Preterm newborn high risk for CP	32 weeks - 12 month s	Cerebral Palsy No data available in review  Abnormal Motor Sensitivit y = 33- 91% Specificit y = 68- 100% Note: Most sensitive at 4 months	14/14

Tool	Citation	Type of Evidence	Numbe r Studie s	Numbe r Patient s	Populatio n	Age at Exam	Accuracy of Detecting Risk for Cerebral Palsy	QUADA S Quality
NEUROLOG								
Amiel- Tison Neurologic al examinatio n	Heinema n 2008	Systemati c review	6 Amiel- Tison	Unable to calculat e from review	Newborn high risk for CP & other diagnoses	3-18 month s	Cerebral Palsy Authors conclude good predictive validity but no values reported Note: The test's purpose is discriminativ e not predictive	14/14
HINE Scored neurologic al examinatio n	Romeo 2015	Systemati c review	10 HINE	3452 n=2621 preterm n=831 termbor n	Preterm & HIE newborn high risk for CP	3-12 month s CA	Cerebral Palsy HINE ≤56 at 3 months: Sensitivity = 90% Specificity = 90% HINE ≤65 at 12 months: Sensitivity = 90% Specificity = 90% Scores <40 = severe CP only A trajectory of longitudinal assessment s are more predictive  HINE with MRI provided more	14/14

Hainama	Cyatamati	4 HIME	Linghia	Nowborn	3-18	accurate prognostic information than the individual tools. Normal neonatal MRI or moderate white matter lesions = HINE scores (>73) and normal motor outcome. Severe basal ganglia lesions were associated with HINE scores (<40) and CP	14/14
Heinema n 2008	Systemati c review	4 HINE	Unable to calculat e from review	Newborn high risk for CP & other diagnoses	month s	Cerebral Palsy Authors conclude good predictive validity but no values reported	14/14

Tool	Citation	Type of Evidenc e	Numb er Studie s	Numbe r Patient s	Populatio n	Age at Exam	Accuracy of Detecting Risk for Cerebral Palsy	QUADA S Quality
Infanib Neurologic al examinatio n	Heinema n 2008	Systemat ic review	3 Infanib	Unable to calculat e from review	Newborn high risk for CP & other diagnoses	3-18 months	Cerebral Palsy Authors conclude moderate predictive validity but no values reported	14/14
Muscle Power Neurologic al examinatio n	Heinema n 2008	Systemat ic review	3 Muscle Power	Unable to calculat e from review	Newborn high risk for CP & other diagnoses	3-18 months	Cerebral Palsy Authors conclude good predictive validity but no values reported Note: The test's purpose is discriminati ve not predictive	14/14
Touwen Neurologic al examinatio n	Bosanqu et 2013	Systemat ic review with meta- analysis	2 Touwe n	142	Preterm newborn high risk for CP	5-12 months CA	Cerebral Palsy Sensitivity = 88% Specificity = 87%	14/14
	Heinema n 2008	Systemat ic review	2 Touwe n	Unable to calculat e from review	Newborn high risk for CP & other diagnoses	3-18 months	Cerebral Palsy Authors conclude good predictive validity but no values reported	14/14
NEUROIMA			40	0044	Dunta ::: C	Dunting	Couck	4.4/4.4
CUS	Bosanqu et 2013	Systemat ic review with meta- analysis	10 CUS	2644	Preterm & HIE newborn high risk for CP	Preterm or term equivale nt age	Cerebral Palsy Sensitivity = 88% Specificity =	14/14

			87%	

Tool	Citatio n	Type of Evidenc e	Numbe r Studie s	Numbe r Patient s	Populatio n	Age at Exam	Accuracy of Detecting Risk for Cerebral Palsy	QUADA S Quality
CUS continue d	Ment 2002	Clinical Practice Guidelin e	7 CUS	2045	Preterm <30 wks as newborn high risk for CP & other outcomes	7-14 days of age	Cerebral Palsy Grade 3 & 4 IVH, cystic PVL, and/or moderate- severe ventriculomegal y injuries were predictive of CP CUS = established as predictive  Recommendatio n: Routine CUS screening should be performed on all infants of <30- weeks' GA once between 7-14 days of age. Plus repeated	14/14
СТ	Ashwal 2004	Clinical Practice Guidelin e	9 CT	782	СР	7month s-16yrs	between 36-40 weeks'  Cerebral Palsy Sensitivity = 77% Yield varied by CP type (hemiplegic > ataxic > mixed > diplegic > quadriplegic > hypotonic > dyskinetic)  Recommendatio n: Neuroimaging should be conducted using MRI, preferably to CT, because MRI is more accurate	14/14

Too I	Citation	Type of Evidence	Numbe r Studie s	Numbe r Patient s	Populatio n	Age at Exam	Accuracy of Detecting Risk for Cerebral Palsy	QUADA S Quality
MRI	Ashwal 2004	Clinical Practice Guideline	10 MRI	644	СР	3months- 41yrs	Cerebral Palsy Sensitivity = 89%  Recommendatio n: 1. Neuroimaging should be conducted using MRI, preferably to CT. Yield varied by CP type (mixed > quadriplegic > hemiplegic > hemiplegic > hemiplegic > hypotonic) and was different to yields from CT MRI = established as predictive 2. Screen for intellectual disability, ophthalmologic and hearing impairments, speech and language disorders, and oral-motor dysfunction as these are common co-occurring impairments	14/14
	Bosanqu et 2013	Systemati c review with meta- analysis	3 MRI	702	Newborn high risk for CP	Term equivale nt age	Cerebral Palsy Sensitivity = 86% Specificity = 89%	14/14

Tool	Citatio n	Type of Evidenc e	Numbe r Studie s	Numbe r Patient s	Populatio n	Age at Exa m	Accuracy of Detecting Risk for Cerebral Palsy	QUADA S Quality
MRI continue d	Ment 2002	Clinical Practice Guidelin e	13 <i>MRI</i>	410	Encephal- opathy newborn high risk for CP & other outcomes	2-8 days	Cerebral Palsy Basal ganglia injury was predictive of CP (50-94%) MRI = established as predictive DWI = probably predictive MRS = possibly predictive  Recommendatio n: MRI should be performed 2-8 days to assess the location and extent of injury DWI and MRS, when available, should also be performed to provide additional prognostic data	14/14

AIMS = Alberta Infant Motor Scale; Amiel-Tison = Amiel-Tison Neurological Examination; CA = Chronological Age; CP = Cerebral Palsy; CT = Computed Tomography; CUS = Cranial Ultrasound; DAYC = Developmental Assessment of Young Infants; DWI = Diffusion Weighted Imaging; GMs= Prechtl's General Movements Assessment; HIE = Hypoxic Ischaemic Encephalopathy; HINE = Hammersmith Infant Neurological Examination; Infanib = Infant Neurological International Battery; MAI = Movement Assessment of Infants; MRI = Magnetic Resonance Imaging; Muscle Power = Active and Passive Muscle Power; MRS = Magnetic Resonance Spectroscopy; NSMDA = Neuro Sensory Motor Development Assessment; PDMS-2 = Peabody Developmental Motor Scales; SD = standard deviation; TIMP = Test of Infant Motor Performance; Touwen = Touwen Infant Neurological Examination.

eFigure. PRISMA Diagram of Study Flow 21 additional 1341 records identified records identified through database through other sources searching 125 records after duplicates removed 125 records 103 records records excluded screened 14 full-text articles **22** full-text excluded: articles • n=12 due to assessed for design eligibility • n=2 due to participants 8 studies included in qualitative synthesis (n=6 systematic reviews; n=2 evidence-based guidelines)