Early Identification and Screening of Neonates At-Risk for Neonatal Encephalopathy

A Statewide Guideline



DKLAHOMA PERINATAL QUALIT IMPROVEMENT COLLABORATIVE

Early Identification and Screening of Neonates At-Risk for Neonatal Encephalopathy

- Aim of the task force was to create a state-wide guideline:
 - Decrease "missed" infants and improve newborn outcomes across the state
 - Common guidelines for NICUs providing therapeutic hypothermia
 - Provide guidance to birthing hospitals and NICUS not providing therapeutic hypothermia on admission criteria



Hypoxic Ischemic Encephalopathy (HIE)

Ulana Pogribna, MD/MPH November 2023

OBJECTIVES

> Define HIE

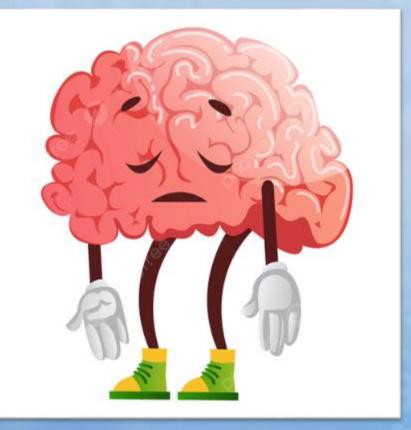
> Incidence of HIE

- Review underlying pathophysiology of HIE
- > Review outcomes for infants with HIE

 Outline diagnostic considerations, including MRI and EEG

Definitions

- > Neonatal Encephalopathy
 - Clinically defined syndrome
 - Neurologic dysfunction in infants in early days of life at or beyond 35 weeks gestation
 - Spectrum of neurologic dysfunction
- Hypoxic ischemic encephalopathy (HIE)
 - "Birth asphyxia" an outdated term
 - Most common type of neonatal encephalopathy
 - Accounts for 15-35% of encephalopathy in late preterm and term infants
 - Represents global hypoxic insult to the brain



HIE Incidence

> 1 to 8 per 1000 live births in developed countries

> As high as 26 per 1000 live births in underdeveloped countries

Variation occurs based on variability of resources

Pathophysiology – Key Points

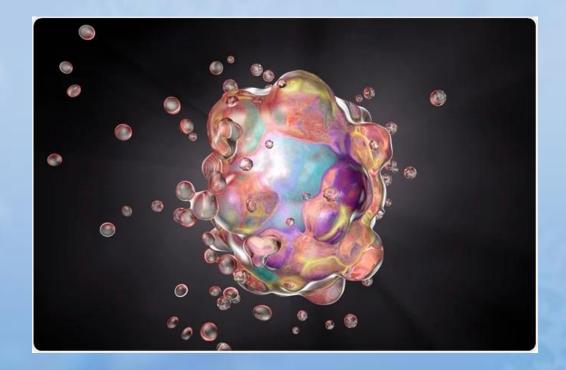
> Brain is dependent on appropriate delivery of oxygen and glucose to function

- > Adequate cerebral blood flow is essential for oxygen and glucose delivery
 - Maintenance of homeostasis
 - Meeting of cellular energy demands

> Variety of conditions compromise appropriate placental perfusion and disrupt oxygen and glucose delivery to the umbilical cord

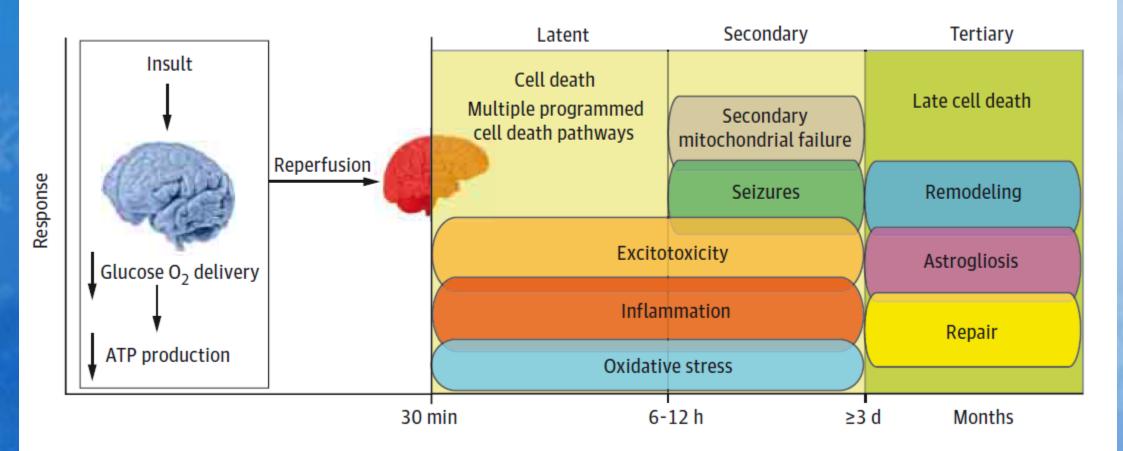
HIE – 3 Main Phases of Injury

- > Initial hypoxic-ischemic insult
- Secondary effects of oxidative stress, mitochondrial deficiency, excitotoxicity, inflammation, and early stages of neuronal necrosis and apoptosis
- Long-term cell death, inflammation, cell turnover/repair, and gliosis



HIE – Timeline of Injury

Figure 1. Schematic Overview of the Pathophysiological Features of Hypoxic-Ischemic Encephalopathy



HIE – TIMELINE OF INJURY

> Primary energy failure

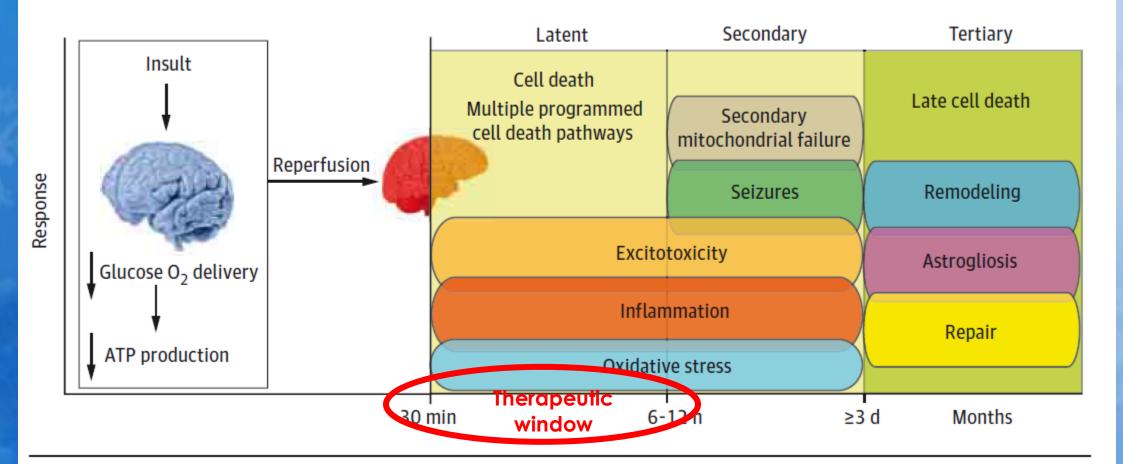
- Occurs early, around hypoxic event
- May resolve based on timing and effectiveness of resuscitative efforts
- Neuronal injury necrotic

> Secondary energy failure

- Occurs 6-12 hours later, may be progressive
- Leads to tissue damage brain and other organs
- Neuronal injury apoptotic

HIE – Timeline of Injury

Figure 1. Schematic Overview of the Pathophysiological Features of Hypoxic-Ischemic Encephalopathy



Douglas-Escobar M. JAMA Pediatrics. 2015.

Therapeutic Hypothermia

> The only approved treatment for HIE

> Works within the "therapeutic window" to decrease brain injury during the phase of secondary energy failure

> Two types:

- Selective head cooling
- Whole body hypothermia
 - > Easier to use and less expensive

Therapeutic Hypothermia

SELECTIVE HEAD COOLING

WHOLE BODY





So, Why Cool?

> Multiple trials to date showing benefits of cooling on outcomes for babies with moderate to severe HIE

> 2 largest multi-center trials showed:

- 18-22 months of age:
 - > Lower mortality and moderate/severe disability
 - Decreased rate of cerebral palsy
 - > Decrease in hearing and visual impairments

– At 6-7 years of age:

- > Benefits seen at 18-22 months persisted at school age
- > Lower rate of mortality and moderate/severe disability
- > Lower rate of cerebral palsy

Reduction in death DID NOT result in increase in survivors with disability

Imaging - MRI

> Preferred modality

Need conventional sequences (T1 and T2) and diffusion weighted imaging (DWI)

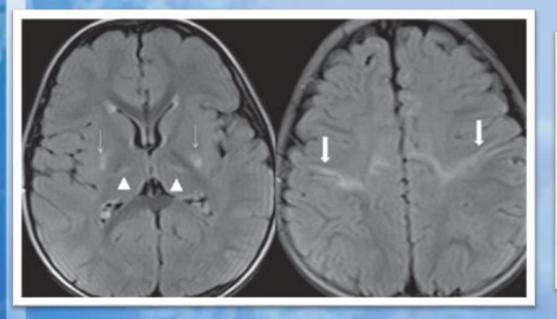
> When to get it

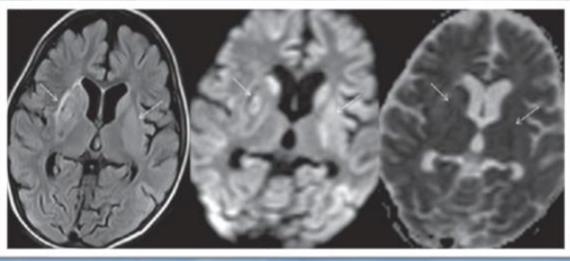
- 3-5 days ideally
- Prior to 2 days too early, may only show subtle changes
- After 7 days can have some "pseudo-normalization" and can miss injury

> Patterns of injury

- Watershed infarcts involving cortical gray matter and subcortical white matter
- Involvement of basal ganglia, thalami, brainstem, hippocampus more reflective of global hypoxic injury

MRI Findings





1 year-old with severe HIE: injury in thalami (arrow head), posterior putamen (thin arrow), and cortex (thick arrow) 2.5 yo male with severe HIE: basal ganglia infarcts

Bano S, et al. J Pediatr Neurosci. 2017

Seizures

Common sign of CNS injury, thus common in neonatal encephalopathy

In neonatal period, HIE is the most common cause

> EEG

- Identifies and characterizes seizures
- 2011 guidelines by the American Clinical Neurophysiology Society recommend
 24 hour continuous EEG monitoring great idea but not always feasible

> aEEG

- Amplitude-integrated EEG
- Continuously monitors cerebral activity and helpful in detecting seizures
- More readily available than continuous EEG

References

- Kurinczuk JJ, White-Koning M, Badawi N. Epidemiology of neonatal encephalopathy and hypoxic-ischaemic incephalopathy. Early Hum Dev. 2010; 86(6): 329-338.
- > Buss JB, Simmons MD, Glass HC. Neonatal encephalopathy: beyond hypoxic-ischemic encephalopathy. Neoreviews. 2021 Mar 22(3): e148-e162.
- > Douglas-Escobar M, Weiss MD. Hypoxic-ischemic encephalopathy: a review for the clinician. JAMA Pediatrics. 2015 169(4): 397-403.
- Bano S, Chaudhary V, Garga UC. Neonatal Hypoxic-ischemic Encephalopathy: A Radiological Review. J Pediatr Neurosci. 2017 Jan-Mar;12(1):1-6. doi: 10.4103/1817-1745.205646. PMID: 28553370; PMCID: PMC5437770.
- > Allen KA. Moderate hypothermia: is selective head cooling or whole body cooling better? Adv Neonatal Care. 2014 Apr;14(2):113-8.
- > Laptook AR, McGowan EC. Outcomes in the era of therapeutic hypothermia. Neoreviews. 2015. Sept 15(9): e386-e395.
- Natarajan G, Pappas A, Shankaran S. Outcomes in childhood following therapeutic hypothermia for neonatal hypoxic-ischemic encephalopathy (HIE). Semin Perinatol. 2016 Dec;40(8):549-555.
- > Bonifacio SL, Hutson S. The Term Newborn: Evaluation for Hypoxic-Ischemic Encephalopathy. Clin Perinatol. 2021 Aug;48(3):681-695.
- > Shah NA, Wusthoff CJ. How to use: amplitude-integrated EEG (aEEG). Arch Dis Child Educ Pract Ed. 2015 Apr;100(2):75-81. doi: 10.1136/archdischild-2013-305676. Epub 2014 Jul 17.

Early Identification and Screening of Neonates At-Risk for Neonatal Encephalopathy Guidelines

Dr. Arlen Foulks, Dr. Susan Bedwell





Oklahoma HIE Task Force

The Oklahoma HIE (Hypoxic Ischemic Encephalopathy) Task Force was assembled to develop state-wide guidelines for the identification and screening of infants at-risk for neonatal encephalopathy. Using the most recent evidencebased information, the task force worked to create a guiding document to ensure all newborns across Oklahoma have the opportunity to receive life-saving treatment at a skilled Oklahoma Neonatal Intensive Care Unit.

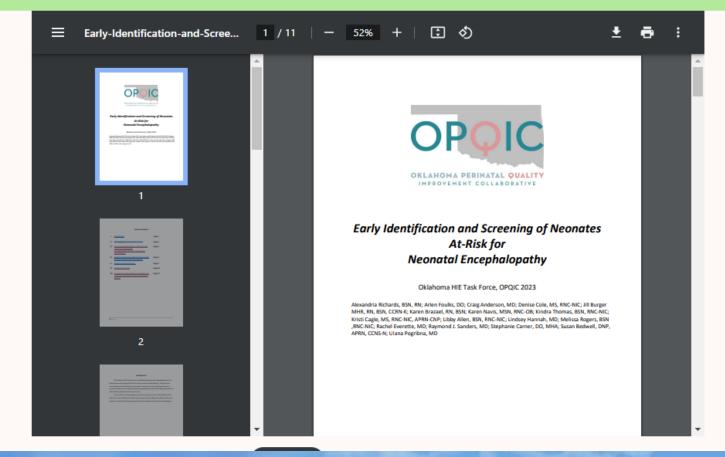


Table of Contents

Early Identification and Screening of Neonates At-Risk for Neonatal Encephalopathy

I.	Introduction	Page 3
II.	HIE Highlights for Medical Providers	Page 4
III.	<u>Recommended Guidelines: Screening and</u> <u>Treatment of Neonatal</u> <u>Encephalopathy Requiring Therapeutic</u> <u>Hypothermia</u>	Page 6
IV.	<u>Decision Tree for Neonatal Encephalopathy</u> <u>Requiring Therapeutic Hypothermia</u>	Page 8
v.	Passive Cooling Directions	Page 9
VI.	Sarnat Scoring Tool	Page 11
VII.	<u>Neonatal Encephalopathy and Hypothermic</u> <u>Treatment Questions Frequently Asked by</u> <u>Parents</u>	Page 12

Birthing Hospital Screening Criteria

Page 6

> Because the window for treatment for hypoxic ischemic
 encephalopathy is short (6 hours), any infant equal to or greater
 than 35 weeks gestation with a history of an acute perinatal event
 with the potential for hypoxia accompanied by any one of the
 following should be immediately referred to a neonatal intensive
 care unit that provides therapeutic hypothermia:

- . APGAR less than or equal to 5 at 10 minutes of life
- Assisted ventilation required at 10 minutes of life

Birthing Hospital Screening Criteria

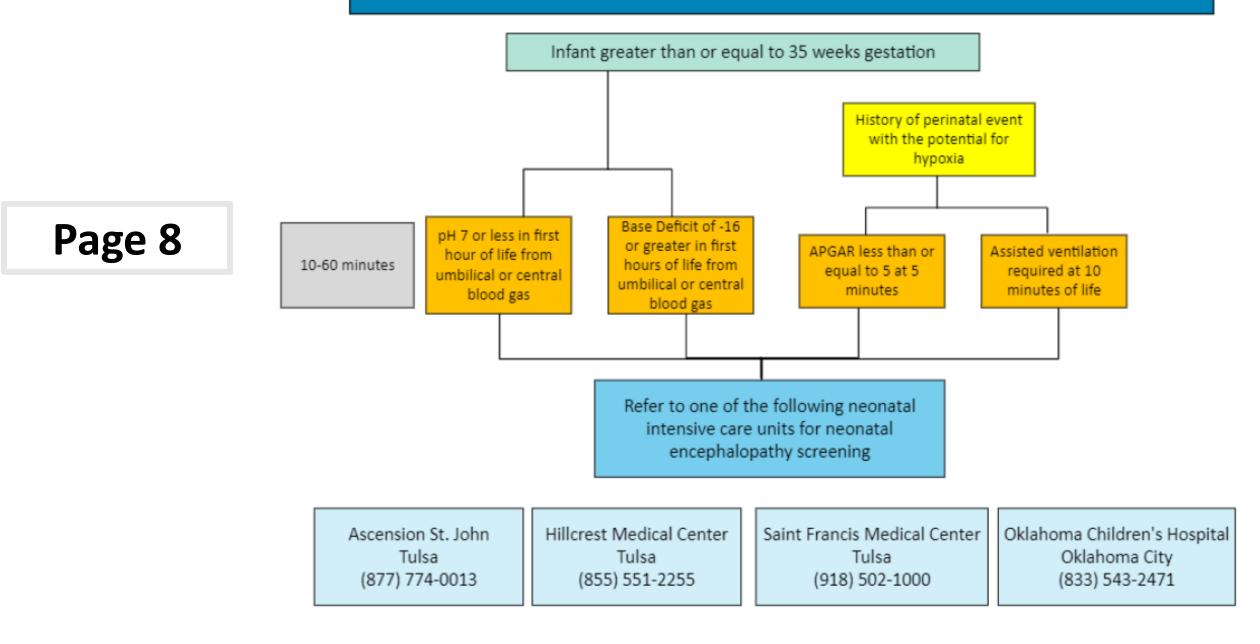
 Infants regardless of perinatal history with an umbilical cord or central blood gas obtained within 1 hour of life with one of the following should be immediately referred to a neonatal intensive care unit that provides therapeutic hypothermia:

. pH of 7 or less

Page 6

Base Deficit of negative 16 mEq or greater

 Infants with severe IUGR, severe chromosomal or congenital anomalies or who are unlikely to benefit from or respond to aggressive life support are not eligible for therapeutic hypothermia.



Neonatal Intensive Care Confirming Criteria

Page 6

The presence of seizure activity or presence of moderate to severe neonatal encephalopathy qualifies the infant for treatment with therapeutic hypothermia.

Neonatal Intensive Care Confirming Criteria

A Neonatologist, or a clinician should examine the infant for signs of encephalopathy using the <u>Modified</u> <u>Sarnat Scoring Tool</u>.

Page 6

The presence of abnormal neurologic signs affecting 3 of the 6 categories indicates moderate to severe encephalopathy and the infant should receive treatment with therapeutic hypothermia.

Neonatal Intensive Care Confirming Criteria

Page

	Sarnat Level			
	Category	Moderate	Severe	
	Level of	Lethargic	Stupor/coma	
	consciousness			
11	Spontaneous activity	Decreased activity	No activity	
	Posture	Distal flexion, full extension	Decerebrate	
-	Tone	Hypotonia (focal, general)	Flaccid	
	Primitive reflexes:			
	Suck	Weak	Absent	
	Moro	Incomplete	Absent	
	Autonomic system:			
	Pupils	Constricted	Skewed	
	Heart rate	Bradycardia	deviation/dilated/nonreactive	
	Respirations	Periodic breathing	Variable heart rate Apnea	

Passive Cooling

Modest hypothermia (core temperature of 33 to 35 degrees C [91.4 to 95 degrees F]) may be indicated for some infants with moderate to severe hypoxic ischemic encephalopathy.

Page 9

- Passive cooling may be advantageous for some infants with signs of perinatal depression prior to and during transport when eligibility for hypothermia therapy is being considered.
- > During passive cooling, no external sources of cooling are applied. Care must be taken to avoid over-cooling because passive cooling is unregulated and, depending on patientspecific and environmental circumstances, the infant's temperature may fall too low.
- CAUTION: Passive cooling should ONLY be initiated under the direction of a neonatologist/clinician from a NICU providing therapeutic hypothermia.

Confirming Criteria Not Met

If the infant is not having seizure activity or the neurological exam using the Modified Sarnat Scoring Tool does not indicate moderate to severe encephalopathy:

Page 7

> Keep the infant normothermic

 Transfer to a neonatal intensive care unit that provides therapeutic hypothermia for continued observation.

The Modified Sarnat Scoring Tool is repeated hourly until the scoring tool indicates moderate to severe encephalopathy or the infant reaches 6-hours post birth

Page 12

Therapeutic Hypothermia (Cooling)

What is cooling?

 Cooling is a treatment for suspected hypoxic ischemic encephalopathy (HIE). Within 6 hours of birth, your baby's temperature is cooled below normal body temperature and is kept in this low range of 33°C to 34°C (91.4°F to 93.2°F) for 72 hours. Cooling may stop further injury to your baby's brain and may reduce or prevent physical or mental conditions that limit movement, senses and activities.

What to expect/side effects

- Sleepy/sedated
- · Pale, patchy, slightly blue skin coloring
- Possible shivering
- Slowing of the heart rate (will return to normal rate after cooling)
- Monitors to track heart rate, oxygen levels and breathing, lines into the umbilical cord, tubes in the mouth
- Medication administration While the cooling process has not been shown to cause pain, measures to comfort your baby, including pain medication, may be provided

What can I do?

- · Allow your baby to rest
- · Gently whisper to your baby
- Provide a calm and quiet environment
- Ask the health care team when hands-on time will occur
- Pump and store your breast milk
- Ask questions when you have them

Feel free to reach out to your baby's health care team For more information about HIE, visit the www.hopeforhie.org



THANK YOU! Oklahoma HIE Task Force, OPQIC 2023

- Alexandria Richards, BSN, RN
- Arlen Foulks, DO
- Craig Anderson, MD
- Denise Cole, MS, RNC-NIC
- Jill Burger MHR, RN, BSN, CCRN-K
- Karen Brazael, RN, BSN
- Karen Navis MSN, RNC-OB
- Kindra Thomas, BSN, RNC-NIC
- Barbara O'Brien MS, RN

- Libby Allen, BSN, RNC-NIC
- Lindsey Hannah, MD
- Melissa Rogers, BSN, RNC-NIC
- Rachel Everette, MD
- Raymond J. Sanders, MD
- Stephanie Carner, DO, MHA
- Susan Bedwell, DNP, APRN, CCNS-N
- Ulana Pogribna, MD/MPH FAAP