

LONG-TERM OUTCOMES OF NEONATAL ABSTINENCE SYNDROME: IMPLICATIONS FOR PROVIDERS AND CAREGIVERS

October 29, 2018 2:30 pm – 3:30 pm EST



Today's Speakers



Peggy Honein, PhD, MPH Director, Division of Congenital and Developmental Disorders



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Introduction and Welcome

Rebecca Russell, MSPH

SVP (Interim) Science and Strategy Senior Director, Applied Research and Evaluation, March of Dimes



National Center on Birth Defects and Developmental Disabilities



Maternal and Child Health Impact of the U.S. Opioid Epidemic

Margaret (Peggy) Honein, PhD, MPH

Director, Division of Congenital and Developmental Disorders National Center on Birth Defects and Developmental Disabilities Centers for Disease Control and Prevention

October 29, 2018

Overview of the Opioid Epidemic

- In 2016, about 11.8 million people in the U.S. misused opioids in the past year, including:
 - 11.5 million pain reliever misusers
 - 948,000 heroin users
- Increase in drug overdose deaths
- Vulnerable populations affected include pregnant women and infants



U.S. State Opioid Prescribing Rates, 2016



Opioid Use among Women

About **1** in **3** women of reproductive age filled an opioid prescription between 2008 – 2012.



Opioid use disorder rates at delivery increased by more than **4-fold**

during 1999 to 2014.



Ailes EC, Dawson AL, Lind JN, et al. MMWR. 2015 Jan 23;64(2):37-41.

Haight SC, Ko JY, Tong VT, et al. MMWR. 2018 Aug 10; 67(31):845-849.

Babies Born with Neonatal Abstinence Syndrome (NAS)

Every 15 minutes, a baby was born with NAS



Nearly **100** babies each day

Babies born with NAS experience serious medical problems



In 2014, for NAS total hospital costs in the US were over \$563 million



Protecting Our Infants Act, 2015

One Hundred Fourteenth Congress of the United States of America

AT THE FIRST SESSION

Begun and held at the City of Washington on Tuesday, the sixth day of January, two thousand and fifteen

An Act

To address problems related to prenatal opioid use.

Be it enacted by the Senate and House of Representatives of the United States of America in Congress assembled,

SECTION 1. SHORT TITLE.

This Act may be cited as the "Protecting Our Infants Act of 2015".

SEC. 2. ADDRESSING PROBLEMS RELATED TO PRENATAL OPIOID USE.

- Department of Health and Human Services:
 - Review and improve coordination
 - Develop a strategy to address gaps in research and federal programs
 - Study and develop recommendations for preventing and treating prenatal opioid use and neonatal abstinence syndrome
 - Improve data and public health response by supporting states and tribes

HHS: U.S. Department of Health and Human Services Public Law No: 114-91

Outcomes Associated With Prenatal Opioid Exposure



Current NCBDDD-Supported Efforts

- With March of Dimes on two NAS pilot projects
 - NAS surveillance based on birth defects surveillance
 - Grantees: Illinois, New Mexico, Vermont
 - Readmissions and adverse outcomes through one year of age
 - Inform NAS surveillance and prevention efforts in other states
 - Understanding the long-term outcomes of NAS: Tennessee Pilot
- With other groups at CDC and other partners
 - Assess various aspects about NAS across the U.S.
 - Broader impact of prenatal opioid exposure on the infant



FY19 Budget Initiatives

- \$10 million for surveillance of emerging threats to mothers and babies
 - Leverage Zika pregnancy and birth defects surveillance system
 - Capture real-time data that can rapidly be translated into clinical guidance
 - Understand long-term implications of known or emerging threats, including infectious agents, vaccines, or medications, such as opioids
- \$2 million for surveillance of neonatal abstinence syndrome <u>ht</u>

PROGRAM HIGHLIGHT

Emerging Threats to Mothers and Babies Initiative

Mothers and babies are often at higher risk during any kind of public health crisis. Supported through supplemental Zika response funding, the innovative Zika pregnancy and infant registry was established to ensure that mothers and babies are adequately monitored and quickly informed about the impact of an emerging threat. The continuation and expansion of this surveillance system is critical to better understanding the long-term implications of not only Zika, but also for other emerging infectious diseases, such as pandemic influenza, and other emerging threats, including prenatal exposure to opioids. The FY 2019 Budget includes an additional \$10 million to support an initiative that will enable CDC to continue to work with states to maintain ongoing registries and continue to monitor mothers and babies for the impact of Zika and serve as a tool for other emerging public health threats.

This additional surveillance will inform public health action for mothers and babies, including prevention strategies, clinical guidance, enhanced follow-up, targeted screening and evaluation, and identification of medical and early interventions to help children thrive. In addition, CDC will work collaboratively with state, local, and territorial health departments to extend the monitoring of babies born to mothers with evidence of Zika infection to better understand the full impact of Zika on child development, and work with healthcare providers and others to develop better assessment and communication tools.

https://www.hhs.gov/sites/default /files/fy-2019-budget-in-brief.pdf

Leverage Zika Infrastructure for Prenatal Opioid Exposure



State Spotlight: Pennsylvania

Background: On January 10, 2018, PA Governor added neonatal abstinence syndrome (NAS) as a reportable condition as part of a 90-day state of emergency for the opioid epidemic. Prior to the 2017 implementation of PA's Zika Birth Defects Surveillance (ZBDS), the state had never collected data on birth defects or NAS.

Methods	Results	
 Developed strategy for facility outreach based on live birth counts and reported neonatal intensive care units (NICUs) Created a brief one-page NAS case report Created electronic survey using REDCap Cloud 	 REDCap Cloud survey for NAS surveillance created in 2 days After 1 month: 342 cases of NAS reported from 57 (61% of) facilities 7 weeks after distribution: 520 cases of NAS reported 	 Rapid tracking of NAS data within the short 90- day timeframe of the opioid state of emergency Fast turn-around to inform targeted community outreach Blueprint for Pennsylvania's disaster preparedness for other emerging surveillance needs

Aligns with CDC's Mission

- Protect the health, safety, and security of the nation
- Put science into action

Bottom line:

- Pregnancy and birth defects surveillance are key components of CDC's preparedness work.
- Birth defects can be the first sign that an emerging infection causes serious harm.





Thank you

For more information, contact CDC 1-800-CDC-INFO (232-4636) TTY: 1-888-232-6348 www.cdc.gov

The findings and conclusions in this report are those of the authors and do not necessarily represent the official position of the Centers for Disease Control and Prevention.



Long-Term Outcomes of Neonatal Abstinence Syndrome: Implications for Providers and Caregivers

> Mary-Margaret A. Fill, MD Michael D. Warren, MD, MPH, FAAP Tennessee Department of Health







Objectives

 Review the clinical presentation and treatment options for infants with NAS

• Discuss possible long-term outcomes of NAS

 Outline opportunities for prevention or early intervention in children and families at risk for NAS



Neonatal Abstinence Syndrome (NAS)

NAS is a postnatal drug withdrawal syndrome that most commonly occurs after intrauterine opioid exposure.



















NAS Treatment

- Nonpharmacologic supportive care
 - Swaddling
 - Minimize environmental stimuli



- Pharmacologic therapy
 - Morphine
 - Buprenorphine
 - Methadone





A Problem of Pandemic Proportions



In the United States, every



a baby is born affected by opioid withdrawal

NAS: A Growing Problem in Tennessee



East Tennessee Disproportionately Impacted



Rate of NAS per 1,000 live births





East Tennessee Disproportionately Impacted



Rate of NAS per 1,000 live births





"The Call"

- Anecdotal reports from educators in east Tennessee
- Children with a history of NAS had learning challenges
- No studies examining educational outcomes in the United States







Examine associations between a history of NAS and educational outcomes.







Potential Educational Data

- Standardized reading / math test scores
 - TN Comprehensive Assessment Program: statewide (3rd grade)
 - Stanford Achievement Test: optional in some districts (K, 1st & 2nd)
- Absenteeism data
 - Excused / unexcused
- Disciplinary data
 - Suspension / expulsion
- Special education data
 - IEP
 - Accommodations
 - Therapies (PT/OT/ST)



Special Education Services in Tennessee





Qualifying Educational Disabilities in TN

Intellectually Gifted Autism **Multiple Disabilities** Deaf-Blindness **Orthopedic Impairment** Deafness **Other Health Impairment Developmental Delay Emotional Disturbance Specific Learning Disabilities** Speech or Language Impairment **Functional Delay Hearing Impairment** Traumatic Brain Disorder Intellectual Disability Visual Impairment














Process Flow





Process Flow



* Individualized Education Program



Process Flow

Referral
Evaluation
Eligibility Determination
Development of IEP
Implementation of Services



Methods: Creation of Dataset

Tennessee Birth Cohort 2008–2011



ICD-9 Diagnosis Code: 779.5 (Drug withdrawal syndrome in newborn)



1:3 matched pairs Birth certificate data Enrolled in TennCare



Methods: Creation of Dataset





















Data Analysis

- Pearson's Chi Square
 - Descriptive comparisons between groups

- Conditional multivariable logistic regression
 - Associations between a history of NAS and outcomes of interest

• SAS 9.4



Matched Demographic Characteristics

	NAS (+)	NAS (–)
Characteristic	N = 1815 n (%)	N = 5441 n (%)
Male	967 (53.3)	2898 (53.3)
White	1694 (93.4)	5080 (93.4)
DOB 8/2010-8/2011	631 (34.8)	1893 (34.8)
East TN residence	1405 (77.4)	4213 (77.4)
TennCare insurance	1815 (100.0)	5441 (100.0)



Delivery and Birth Characteristics

	NAS (+)	NAS (–)	
Characteristic	n (%)	n (%)	P Value
Birth weight <2500g	435 (24.0)	500 (9.2)	<0.0001
Gestational age <37 weeks	392 (21.6)	625 (11.5)	<0.0001
NICU admission	379 (20.9)	315 (5.8)	<0.0001
Maternal tobacco use in pregnancy	1196 (65.9)	1640 (30.1)	<0.0001







Outcome #1: Referral for Evaluation









Outcome #2: Eligibility Determination





Qualifying Educational Disabilities in TN

Intellectually Gifted Autism **Multiple Disabilities** Deaf-Blindness **Orthopedic Impairment** Deafness **Other Health Impairment Developmental Delay Emotional Disturbance Specific Learning Disabilities** Speech or Language Impairment **Functional Delay Hearing Impairment** Traumatic Brain Disorder Intellectual Disability Visual Impairment



Qualifying Educational Disabilities in TN

Autism

Deaf-Blindness

Deafness

Developmental Delay

Emotional Disturbance

Functional Delay

Hearing Impairment

Intellectual Disability

Intellectually Gifted

Multiple Disabilities

Orthopedic Impairment

Other Health Impairment

Specific Learning Disabilities

Speech or Language Impairment

Traumatic Brain Disorder

Visual Impairment



Special Education Exceptionalities

Outcome	NAS (+) n (%)	NAS (–) n (%)	P Value
Autism	6 (0.3)	22 (0.4)	0.8
Developmental Delay	96 (5.3)	19 <mark>3 (3.6)</mark>	0.001
Other Health Impairment	12 (0.7)	27 (0.5)	0.5
Specific Learning Disability	7 (0.4)	16 (0.3)	0.6
Speech / Language Impairment	187 (10.3)	451 (8.3)	0.009







Outcome #3: Implementation of Services





Types of Services Received

Service	NAS (+) n (%)	NAS (–) n (%)	P Value
Accommodations	98 (5.4)	225 (4.1)	0.02
Aide / Paraprofessional	3 (0.2)	12 (0.2)	0.2
Occupational Therapy	55 (3.0)	126 (2.3)	0.09
Physical Therapy	17 (0.9)	54 (1.0)	0.8
Speech Therapy	255 (14.0)	586 (10.8)	0.0002



Types of Services Received

Service	NAS (+) n (%)	NAS (–) n (%)	P Value
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Conditional Logistic Regression

Outcome	Adjusted Odds Ratio	95% Cl
Referred for evaluation	1.44	1.23–1.67
Eligible for services	1.36	1.15–1.60
Received therapies/services	1.37	1.16-1.61

* Controlled for matching factors, maternal education status, and maternal tobacco use during pregnancy.



Conditional Logistic Regression

Outcome	Adjusted Odds Ratio	95% Cl
Developmental Delay	1.34	1.03–1.76
Speech / Language Impairment	1.26	1.04–1.52

* Controlled for matching factors, maternal education status, and maternal tobacco use during pregnancy.

Conditional Logistic Regression

Outcome	Adjusted Odds Ratio	95% Cl
Accommodations	1.32	1.03–1.69
Speech Therapy	1.33	1.12–1.57

* Controlled for matching factors, maternal education status, and maternal tobacco use during pregnancy.

Additional Regression Models

Outcome	aOR	95% CI
Model: maternal education, maternal tobacco,	birthweigh	t, NICU
Referred for evaluation	1.32	1.13–1.55
Eligible for services	1.26	1.07–1.49
Received therapies/services	1.27	1.07–1.51
Model: maternal education, maternal tobacco,	gestationa	l age, NICU
Referred for evaluation	1.37	1.17–1.60
Eligible for services	1.30	1.10–1.54
Received therapies/services	1.31	1.10–1.55
Model: maternal education, maternal tobacco,	birthweigh	t, gest age
Referred for evaluation	1.34	1.14–1.58
Eligible for services	1.28	1.08–1.51
Received therapies/services	1.28	1.09–1.52

Growing Body of Evidence?

Neonatal Abstinence Syndrome and High School Performance

Ju Lee Oei, MD,^{a,b,c} Edward Melhuish, PhD,^{4,e,t} Hannah Uebel,^a Nadin Azzam,^a Courtney Breen, PhD,⁶ Lucinda Burns, PhD,⁶ Lisa Hilder, MBBS,^b Barbara Bajuk, MPH,ⁱ Mohamed E. Abdel-Latif, MD,^{i,k} Meredith Ward, FRACP,^{a,b} John M. Feller, FRACP,^{a,j} Janet Falconer, CNC,^m Sara Clews, CNC,^m John Eastwood, FRACP, PhD,^{a,e,n,a,p} Annie Li,^a Ian M. Wright, FRACP^{d,a,r}



FIGURE 1

Linkage rates between children with NAS, control, and rest of NSW population to NAPLAN results.



Composite Test Score Differences





Limitations

- 1. Unable to analyze all children born with NAS in Tennessee during 2008–2011
- 2. Could not validate that all children in our sample had in utero opioid exposure
- **3.** Matching to special education database may have failed to match some children who had indeed been referred
- 4. Unable to control for some factors which have been shown to increase the risk of NAS
- 5. Potential differential referral patterns among children with a history of NAS compared to those without
- 6. Unable to verify the diagnostic coding of NAS, or stratify results based on severity of NAS



Summary of Results

- Novel analysis linking health and education datasets
- Children with a history of neonatal abstinence syndrome were significantly more likely to
 - be referred for evaluation of an educational disability
 - meet criteria for a disability, specifically developmental delay, or speech or language impairment
 - receive therapies or services, specifically accommodations or speech therapy



Public Health Implications

- Ongoing primary prevention efforts are needed to reduce intrauterine opioid exposure and NAS.
- Identification of infants with a history of NAS, and prompt referral to early intervention services is important for the early diagnosis and treatment of possible developmental or learning disabilities.
- Additional resources may be needed for school systems in areas with high rates of NAS in order to provide students with needed services



Individuals with Disabilities Act (IDEA)

- Federal law
- Originally established 1975
 - Last reauthorized 12/2004



- Ensures that children (3–21 years of age) with disabilities have the opportunity to receive free, appropriate public education (Part B)
- Provides assessments and early intervention services to children with disabilities as early as birth through 2 years of age (Part C)

Benefits of Early Intervention ...

- Infants/toddlers participating in Part C demonstrate:
 - Increased motor, social, and cognitive functioning
 - Acquisition of age-appropriate skills
 - Reduced negative impacts of their disabilities
 - Greater than expected growth in social relationships, use of knowledge & skills, taking action to meet needs
















Acknowledgments

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Tennessee Department of Education Rachel Wilkinson Dave Williams

March of Dimes

TennCare

Mary Lou Mangan Wesley Thompson

Vanderbilt University Medical Center Dr. Stephen Patrick

Centers for Disease Control and Prevention Dr. Stacey Bosch Dr. Jennifer Lind Dr. Daisy Christensen Dr. Marshalyn Yeargin-Allsopp Dr. Elizabeth Ailes







MEDICAL CENTER



Thank You!



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THANK YOU

marchofdimes.org



Extra Slides

















Types of Prenatal Opioid Exposure in TN



Examples of Classroom & Assessment Accommodations

1. Presentation

- Repeat directions, read aloud, use of larger bubbles on answer sheet
- 2. Response
 - Use of computer, use reference aids, mark answers in book
- 3. Timing/Scheduling
 - Extended time, frequent breaks
- 4. Setting
 - Study carrel, special lighting, separate room



Other Demographic Characteristics

	NAS (+)	NAS (–)	
Characteristic	n (%)	n (%)	P Value
Household Income <\$35,000	1184 (95.6)	3440 (89.7)	<0.0001
Mother married	532 (29.3)	2182 (40.1)	<0.0001
Mother education <hs degree<="" td=""><td>611 (33.7)</td><td>1571 (28.9)</td><td><0.0001</td></hs>	611 (33.7)	1571 (28.9)	<0.0001
Enrolled in WIC	1281 (70.6)	4358 (80.1)	<0.0001



	NAS (+)	NAS (–)	
Characteristic	n (%)	n (%)	P Value
Prenatal care	1677 (92.7)	5351 (98.6)	<0.0001
Mean no. prenatal visits (range)	9.4 (9.1–9.6)	11.8 (11.6–11.9)	<0.0001



Sub-analysis of 'Referred'

	NAS (+)	NAS (–)	
Characteristic	n/N (%)	n/N (%)	P Value
Referred	351/1815 (19.3)	745/5351 (13.7)	<0.0001
Eligible for Services	284/351 (80.9)	634/745 (85.1)	0.08
Receipt of Services	278/284 (97.9)	620/634 (97.8)	0.93



Matching Factors

- **1.** Sex
- 2. Race/ethnicity
- 3. Kindergarten cohort (~ age)
- 4. Public health region of residence
- 5. TennCare enrollment status



RESOURCES

Educational Disabilities Among Children Born With Neonatal Abstinence Syndrome

Mary Margaret J, FU, MA^{1,1,4} Angela H, Miller, Po's M2P L, Fachal J, Shlowara, MFP¹ Maturali, Charran MJ, MPN¹ John F, June, 1712, 762^{1,1} Killion Schoffner, Mir Tmeetry F, Janes, Hi^{1,4}

Increment Neural addresses syndrome (XAS) are posterial drug withdowed syndrome dot can accurate instructuring opicie Assesses for a solutional outcomes have not been donessing y saminar. We analyze Emorse of air to understand the resel for special datactional services are grither than the release in the SAS.

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mentations Results of this nevel analysis linking health and education data revealed dust children with a bistory of NAS were significantly more likely to have a subsequent advantional deabling.

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MS PREVENTING NAS IN YOUR BABY

WHAT IS NAS?



Neonutal abstinence syndrome (also called NAS) is a group of conditions caused writen a baby withdraws from certain drugs he's exposed to its the womb before birth. NAS most often happens when a baby's mother takes opiolds during tregnancy.

Babies born with NAS can have health problems at birth, including body shakes and problems with brosthing, cating and alcoping. They also may have learning problems later in life.

Opioids are drugs that often are used to treat pain. Your health care provider may prescribe you an opioid. They're also sold illegally. Opioids can be dangerous and addictive. Any opioid can cause NAS.

IF YOU'RE PREGNANT AND USING OPIOIDS

- Tell your prenatal care provider right away. Ask about taking a safer medicine during pregnancy.
- Don't stop taking any opicid until you talk to your provider. Quitting suddenly can cause problems for you and your baby.
- Make sure any provider who prescribes you an opicid knows you're pregnant.
- falk with your provider to make sure the hospitpl where you plan to have your baby has providers who have experience taking care of babies with NAS.

 If you're addicted to opioids, talk to your provider about medication assisted treatment (MAT). NAS may be easier to treat for babies

whose moms get MAT during pregnancy.

MARCHOFDIMES.ORG/OPIOIDS

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CARING FOR A BABY WITH NAS

WHAT ARE THE SIGNS AND SYMPTOMS OF NAS?

> Babies born with neonstal abstinence syndrome (also called MAS) can have health problems at binth and may need treatment in a hospital. Babies

born with NAS may have low birthweight and jourdice (when the eyes and skin look yellow). They also may have signs and symptoms like:

- Problems with breathing, feeding and sleeping
 Being fussy—crying a lot and having a
- high-pitched cry - Body shakes and sexures
- Body snakes and setzun
 Feuer

As they grow older, children who had NAS may have problems with speech, lenguage and learning. They may need early intervention services to help them learn to wolk, talk and interact with others.

NAS is a group of conditions caused when a baby withdraws from certain drugs he's exposed to in the womb before birth. NAS most often happens when a baby's mother takes opioids during pregnancy.

CARING FOR A BABY WITH NAS

- After birth, your baby may need to stay in the hospital for treatment.
- Make sure your baby gots engoing care from a health care provider.
 Don't trybs tried NNS on your own. Tell your baby's provider if you're womted about your baby's development.

3. Try these things to help calm your baby:

- Keep your baby's room quiet and the lights dim.

Give your baby skin-to-skin care. Put your baby only in a disper and lay him on your bare chest.

If possible, breastfeed your baby.
 Read to your baby.

 Always put your baby on his back to slotp to reduce the risk of sudden infant death syndrome (also called SIDS).

 Ask your baby's health care provider about early intervention services. To find services, visit: cdc.opv/ncbddd/actearly

MARCHOFDIMES.ORG/NAS

See The set of the second seco

Fill M-MA, Miller AM, Wilkinson RH, et al. Educational Disabilities Among Children Born With Neonatal Abstinence Syndrome. Pediatrics. 2018;142(3):e20180562

NEW MOD INFOGRAPHICS COMING MONDAY, NOV 5™ Preventing NAS in your baby & Caring for a baby with NAS



RESOURCES

MARCHOFDIMES.ORG & NACERSANO.ORG

- Neonatal abstinence syndrome
- Prescription medicine during pregnancy; includes video: Prescription medicine before pregnancy
- Prescription opioids during pregnancy; includes link to the Health Action Sheet: Are you taking any of these prescription painkillers?

MARCHOFDIMES.ORG/NURSING

- Assessment of neonatal abstinence
- Impact of prenatal drug use: Managing the consequences of opioid and marijuana use
- Understanding addiction, drug use and abuse among women



